

**UNITED STATES
SECURITIES AND EXCHANGE COMMISSION
Washington, D.C. 20549**

FORM S-1
REGISTRATION STATEMENT
UNDER
THE SECURITIES ACT OF 1933

Clearside Biomedical, Inc.

(Exact name of registrant as specified in its charter)

Delaware
(State or other jurisdiction of
incorporation or organization)

2834
(Primary Standard Industrial
Classification Code Number)

45-2437375
(I.R.S. Employer
Identification Number)

1220 Old Alpharetta Road, Suite 300
Alpharetta, GA 30005
(678) 270-3631

(Address, including zip code, and telephone number, including area code, of registrant's principal executive offices)

Daniel H. White
Chief Executive Officer
Clearside Biomedical, Inc.
1220 Old Alpharetta Road, Suite 300
Alpharetta, GA 30005
(678) 270-3631

(Name, address, including zip code, and telephone number, including area code, of agent for service)

Copies to:

Brent B. Siler
Darren K. DeStefano
Brian F. Leaf
Cooley LLP
11951 Freedom Drive
Reston, VA 20190-5656
(703) 456-8000

Peter N. Handrinos
Latham & Watkins LLP
John Hancock Tower
200 Clarendon Street
Boston, MA 02116
(617) 948-6000

Approximate date of commencement of proposed sale to the public: As soon as practicable after the effective date of this registration statement.

If any of the securities being registered on this Form are to be offered on a delayed or continuous basis pursuant to Rule 415 under the Securities Act of 1933, check the following box.

If this Form is filed to register additional securities for an offering pursuant to Rule 462(b) under the Securities Act, check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering.

If this Form is a post-effective amendment filed pursuant to Rule 462(c) under the Securities Act, check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering.

If this Form is a post-effective amendment filed pursuant to Rule 462(d) under the Securities Act, check the following box and list the Securities Act registration number of the earlier effective registration statement for the same offering.

CALCULATION OF REGISTRATION FEE

Title of Securities being Registered	Proposed Maximum Aggregate Offering Price(1)(2)	Amount of Registration Fee
Common Stock, \$0.001 par value per share	\$	\$

- (1) In accordance with Rule 457(o) under the Securities Act of 1933, as amended, the number of shares being registered and the proposed maximum offering price per share are not included in this table.
 (2) Estimated solely for purposes of computing the amount of the registration fee pursuant to Rule 457(o) under the Securities Act. Includes offering price of additional shares that underwriters have the option to purchase.

Indicate by check mark whether the registrant is a large accelerated filer, an accelerated filer, a non-accelerated filer, or a smaller reporting company. See the definitions of "large accelerated filer," "accelerated filer" and "smaller reporting company" in Rule 12b-2 under the Securities Exchange Act of 1934. (Check one):

Large Accelerated Filer Accelerated Filer Non-accelerated Filer Smaller Reporting Company

The registrant hereby amends this Registration Statement on such date or dates as may be necessary to delay its effective date until the registrant shall file a further amendment that specifically states that this Registration Statement shall thereafter become effective in accordance with Section 8(a) of the Securities Act of 1933, as amended, or until the registration statement shall become effective on such date as the Commission, acting pursuant to said Section 8(a), may determine.

[Table of Contents](#)

The information in this preliminary prospectus is not complete and may be changed. We may not sell these securities until the registration statement filed with the Securities and Exchange Commission is declared effective. This preliminary prospectus is not an offer to sell these securities and we are not soliciting offers to buy these securities in any jurisdiction where the offer or sale is not permitted.

Subject to Completion, dated September 9, 2014

Shares



COMMON STOCK

We are offering _____ shares of our common stock. This is our initial public offering and no public market currently exists for our common stock. We expect the initial public offering price to be between \$ _____ and \$ _____ per share.

We have applied to list our common stock on The NASDAQ Global Market under the symbol "CLSD."

We are an emerging growth company, as defined in the Jumpstart Our Business Startups Act of 2012, and will be subject to reduced public company reporting requirements.

Investing in our common stock involves a high degree of risk. See "[Risk Factors](#)" beginning on page 15.

	<u>PER SHARE</u>	<u>TOTAL</u>
Initial public offering price	\$ _____	\$ _____
Underwriting discount and commissions(1)	\$ _____	\$ _____
Proceeds, before expenses, to us	\$ _____	\$ _____

(1) See "Underwriting" in this prospectus for a description of compensation payable to the underwriters.

We have granted the underwriters an option to purchase up to _____ additional shares of common stock, exercisable at any time until 30 days after the date of this prospectus. If the underwriters exercise the option in full, the total underwriting discounts and commissions payable by us will be \$ _____ and the total proceeds to us, before expenses, will be \$ _____.

Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities or passed upon the adequacy or accuracy of this prospectus. Any representation to the contrary is a criminal offense.

The underwriters expect to deliver the shares on or about _____, 2014.

RBC CAPITAL MARKETS

WELLS FARGO SECURITIES

NEEDHAM & COMPANY

OPPENHEIMER & CO.

Prospectus dated _____, 2014

TABLE OF CONTENTS

	<u>Page</u>		<u>Page</u>
Prospectus Summary	1	Executive Compensation	139
Risk Factors	15	Certain Relationships and Related Party Transactions	151
Special Note Regarding Forward-Looking Statements and Industry Data	56	Principal Stockholders	157
Use of Proceeds	58	Description of Capital Stock	161
Dividend Policy	59	Shares Eligible for Future Sale	167
Capitalization	60	Material U.S. Federal Income Tax Consequences to Non-U.S. Holders	170
Dilution	63	Underwriting	174
Selected Financial Data	65	Legal Matters	182
Management's Discussion and Analysis of Financial Condition and Results of Operations	67	Experts	182
Business	86	Where You Can Find Additional Information	182
Management	131	Index to Financial Statements	F-1

You should rely only on the information contained in this prospectus and any free writing prospectus prepared by or on behalf of us or to which we have referred you. We have not authorized anyone to provide you with information that is different from that contained in such prospectuses. We are offering to sell shares of our common stock, and seeking offers to buy shares of our common stock, only in jurisdictions where such offers and sales are permitted. The information in this prospectus is accurate only as of the date of this prospectus, regardless of the time of delivery of this prospectus or any sale of our common stock.

Until and including _____, 2014, 25 days after the date of this prospectus, all dealers that buy, sell or trade our common stock, whether or not participating in this offering, may be required to deliver a prospectus. This is in addition to the dealer's obligation to deliver a prospectus when acting as an underwriter and with respect to unsold allotments or subscriptions.

For investors outside of the United States: neither we nor any of the underwriters have done anything that would permit this offering or possession or distribution of this prospectus in any jurisdiction where action for that purpose is required, other than in the United States. You are required to inform yourselves about and to observe any restrictions relating to this offering and the distribution of this prospectus.

PROSPECTUS SUMMARY

This summary highlights information contained elsewhere in this prospectus and does not contain all of the information that you should consider in making your investment decision. Before investing in our common stock, you should carefully read this entire prospectus, including our financial statements and the related notes thereto and the information set forth under the sections “Risk Factors” and “Management’s Discussion and Analysis of Financial Condition and Results of Operations,” in each case included in this prospectus. Unless the context otherwise requires, we use the terms “Clearside,” “company,” “we,” “us” and “our” in this prospectus to refer to Clearside Biomedical, Inc.

Overview of Clearside Biomedical

We are a clinical-stage biopharmaceutical company developing first-in-class drug therapies to treat chronic, blinding diseases of the eye. Our current product candidates focus on diseases affecting the retina and choroid, especially diseases associated with macular edema, and are injected non-surgically into the suprachoroidal space, or SCS, adjacent to the choroid, using our proprietary microinjector. With the SCS injection procedure, our product candidates are more directly administered to the retina and choroid as compared to currently used intravitreal injections, which we believe may provide a number of benefits, including lower frequency of necessary administration and faster onset of therapeutic effect. We hold the exclusive rights to develop and commercialize drugs for non-surgical administration into the SCS. Our most advanced product candidates, CLS-1001 and CLS-1003, are based on commonly used ophthalmic therapeutic agents, which we believe will allow us to more efficiently and predictably pursue the regulatory approval of these product candidates under Section 505(b)(2) of the Federal Food, Drug, and Cosmetic Act. Under Section 505(b)(2), if we can establish that reliance on the FDA’s previous findings of safety and effectiveness for an approved product is scientifically appropriate, or there is scientific literature to support the product’s safety or effectiveness, it may eliminate the need for us to conduct some of the preclinical studies or clinical trials for the new product candidate that might otherwise have been required. CLS-1001, for macular edema associated with non-infectious uveitis, is in an ongoing Phase 1/2 clinical trial, from which we expect final results in the first half of 2015. We expect to be able to conduct a single pivotal Phase 3 clinical trial in approximately 150 patients beginning in the first half of 2015, the results of which we believe, based on a meeting with the FDA and subsequent written correspondence, will be sufficient to support the filing of a New Drug Application, or NDA, to the U.S. Food and Drug Administration, or FDA, in the first half of 2017. For CLS-1003, for macular edema associated with retinal vein occlusion, or RVO, we plan to initiate a Phase 2 clinical trial before the end of 2014, with data expected in the second half of 2015. In our CLS-1002 program, for wet age-related macular degeneration, or wet AMD, we are evaluating drug compounds with anti-VEGF activity or dual anti-VEGF and anti-platelet-derived growth factor, or anti-PDGF, activity for SCS injection, and plan to file an investigational new drug application, or IND, by the end of 2015. We are also considering a development program for drug compounds to treat diabetic macular edema, or DME. We estimate that there are nearly five million people in the United States diagnosed with our target indications and that worldwide annual sales of drugs to treat these indications were approximately \$6 billion in 2013.

Our current and planned development programs aim to restore or improve visual function primarily by reducing the macular edema associated with a number of diseases affecting the retina, the tissue that lines the inside of the eye and is the part of the eye primarily responsible for vision, and the choroid, the layer adjacent to the retina that supplies the retina with blood, oxygen and nourishment. These diseases include

uveitis, RVO, wet AMD and DME. Macular edema is the build-up of fluid that can cause abnormal swelling of the macula, the portion of the retina responsible for central vision and color perception. This swelling can rapidly result in deterioration of vision and can eventually lead to blindness.

The most common treatments for diseases of the retina and choroid are corticosteroids and drugs that inhibit vascular endothelial growth factor, or anti-VEGF drugs. These corticosteroids and anti-VEGF drugs are injected into the vitreous, a jelly-like substance that occupies the central portion of the eye. Intravitreal injections rely on drug to diffuse outward from the vitreous to the retina and choroid, which diffusion can result in lower concentrations of drug in these targeted areas. At the same time, this diffusion can result in drug spreading to unintended parts of the eye, where, for some drugs, it can cause significant side effects. By contrast, with our approach, drug is injected into and spreads within the SCS, which is the space located between the choroid and the outer protective layer of the eye known as the sclera, to reach the back of the eye. We believe SCS administration may provide improved bioavailability to the targeted retina and choroid, with drug remaining more localized and away from other parts of the eye, where the drug could cause side effects. Corticosteroids are effective both at treating the inflammation aspect of ocular disease and at reducing macular edema, but when delivered intravitreally, they have been associated with significant side effects, such as cataracts and elevated intraocular pressure, or IOP, which can lead to glaucoma. Anti-VEGF drugs, while highly effective in reducing macular edema, require frequent administration, in some cases as often as monthly. We believe that our differentiated approach of injecting ocular drugs into the SCS may result in fewer side effects and improved bioavailability of drug at the diseased retina and choroid, as compared to intravitreal administration, a faster onset of therapeutic effect, a similar efficacy profile with lower drug amounts required, and longer duration of therapeutic effect with less frequent required injections. The procedure for SCS injection with our microinjector is conducted in an in-office setting and is similar to the procedure for intravitreal injection, and we therefore expect our products, if they achieve regulatory approval, to be easily incorporated into retinal specialists' standard medical practice.

We are developing CLS-1001 for the treatment of macular edema associated with non-infectious uveitis, an indication for which we are seeking orphan drug designation and exclusivity. CLS-1001 consists of an SCS injection of CLS-TA, our proprietary, preservative-free formulation of the corticosteroid triamcinolone acetonide, or TA, specifically designed to be administered through our SCS microinjector. We expect final results from our ongoing Phase 1/2 clinical trial with Triesence, a commonly used TA, in the first half of 2015, and we plan to initiate a Phase 2 clinical trial with CLS-TA in approximately 30 patients in the second half of 2014, with data expected in the first half of 2015. We also plan to initiate a single pivotal Phase 3 clinical trial in approximately 150 patients in the first half of 2015, with the goal of filing a Section 505(b) (2) NDA in the first half of 2017. Based on our discussions with the FDA, which included both a pre-IND meeting in 2012 and subsequent written correspondence from the FDA, we believe we will be able to initiate our pivotal Phase 3 clinical trial of CLS-1001 before we have the final results from our planned Phase 2 clinical trial. We are developing CLS-1003 for the treatment of macular edema associated with RVO. CLS-1003 consists of an SCS injection of CLS-TA with our microinjector, for administration together with an intravitreal injection of an anti-VEGF drug. We plan to initiate a Phase 2 clinical trial in approximately 40 patients before the end of 2014, with data expected in the second half of 2015. Under our CLS-1002 program, we are evaluating a number of compounds to develop as an SCS-injected therapy for the treatment of wet AMD, including compounds with anti-VEGF activity and compounds with dual anti-VEGF and anti-PDGF activity. We expect to select a lead drug candidate under this program for IND submission in 2015.

Our drug candidates, microinjector and method of non-surgical drug delivery into the SCS are protected by four issued U.S. patents and over 20 patent applications filed in the United States and major international markets, including the European Union, Canada, India, Japan, China and Australia. Our issued patents are broadly directed to methods of administering drugs of any type into the SCS by injection and are not scheduled to expire until between 2027 and 2029. Our patent applications relate to SCS delivery technology and methods, our current therapeutic drug candidates and formulations, various therapeutic uses, including treatment of specific indications such as macular edema associated with non-infectious uveitis and RVO, as well as wet AMD and DME, and administration of specific classes of drugs to the SCS, including anti-inflammatory drugs and anti-VEGF drugs. Our pending patent applications are not projected to expire until between 2027 and 2034.

If our product candidates are approved, we plan to commercialize them with a specialty sales force of 30 to 40 representatives to target retinal specialists in the United States, and we may also pursue collaborations with third parties to commercialize any of our drugs approved for marketing outside the United States.

The members of our management team have held senior leadership positions at a number of pharmaceutical and biotechnology companies with approved ophthalmic drugs, including Alcon, Alimera Sciences, CIBA Vision and ISTA Pharmaceuticals. Members of our team have contributed to the development and commercialization of several retinal products, such as Triesence, Iluvien, Nevanac, Visudyne and Xibrom.

Anticipated Benefits from the Clearside Approach

We are developing drug candidates for back of the eye diseases to be administered to the retina and choroid through the SCS using our proprietary microinjector. We believe that our novel, patented approach of SCS drug administration may provide a number of potential benefits, including:

- **Improved bioavailability at the site of disease and faster onset of therapeutic effect.** In preclinical studies, we observed higher amounts of drug present in the retina and choroid following SCS injection, as compared to the distribution of the same drug following intravitreal injection. We believe this suggests that SCS injection may have similar or better efficacy with a faster onset of therapeutic effect than intravitreal injection in diseases of the retina and choroid like uveitis, RVO, wet AMD and DME.
- **Similar efficacy profile with lower drug amounts required.** In a preclinical study in a model of uveitis, SCS injection of TA achieved similar efficacy results with only 10% of the drug amount commonly administered through intravitreal injection. We believe this suggests our approach may allow us to develop ocular therapies with similar efficacy, but with lower amounts of active drug, compared to current therapies.
- **Less frequent injections.** Current treatments for RVO require monthly intravitreal injections of anti-VEGF drugs, while current treatments for wet AMD require such injections as often as seven times per year. For RVO patients, we believe that a combination of a standard intravitreal injection of an anti-VEGF drug that addresses the vascular aspect of the disease, together with an SCS injection of CLS-TA, which addresses the inflammatory aspect of RVO, may have similar or better efficacy with a reduction in the frequency of required anti-VEGF treatments from once every 30 days to once every 90 days. In wet AMD, we believe that more direct application of drugs with anti-VEGF or dual anti-VEGF and anti-PDGF activity to the neovascularization in the choroid through SCS injection may block the process of additional new vascular growth within the choroid before the

vessels break into the retina and create further damage through leakage. We believe that such SCS injection of treatment therefore has the potential to delay vision loss for a longer duration of time than the current standard intravitreal administration of anti-VEGF drugs and, accordingly, could require less frequent administration.

- **Enhanced safety profile.** Intravitreal injections result in drugs diffusing throughout the eye, including into the lens, iris and ciliary body at the front of the eye, which for some drugs, has been associated with safety issues, such as cataracts and elevated IOP levels. Intravitreal administration of TA has been associated with cataracts and increases in IOP levels in 20% to 60% of patients. Because SCS injection of drugs appears to result in drug remaining localized in the retina and choroid without substantial diffusion to the vitreous or the front portion of the eye, we believe SCS injection has the potential to reduce the incidence of these side effects. Based on interim results from our ongoing Phase 1/2 clinical trial, none of the eight patients for whom data is available have experienced cataracts, abnormally high levels of IOP or other treatment-related serious adverse events over the course of 26 weeks following a single SCS injection of TA.
- **Incorporated into standard medical practice.** If approved for marketing, our drugs will be packaged together with our SCS microinjector for one-time use by retinal specialists in their offices. Our product candidates are designed to be administered using a procedure similar to that used for intravitreal injections, without the need for any capital equipment to be installed in the office. Accordingly, we expect our products, if approved, will be easily incorporated into retinal specialists' standard medical practice.

Strategy

Our goal is to be a leading biopharmaceutical company focused on the development and commercialization of therapeutics to treat chronic, blinding diseases of the eye, with a particular emphasis on diseases affecting the choroid and the retina. The key elements of our strategy are:

- **Advancing CLS-1001 and CLS-1003 to FDA approval under the Section 505(b)(2) NDA regulatory pathway.** Our most advanced product candidates, CLS-1001 and CLS-1003, utilize CLS-TA, our proprietary formulation of TA, a corticosteroid that has been used for decades to treat ocular inflammatory conditions. We believe this approach will allow us to more efficiently and predictably pursue the regulatory approval of these product candidates under Section 505(b)(2) of the Federal Food, Drug and Cosmetic Act. We plan to advance CLS-1001 into a single pivotal Phase 3 clinical trial for macular edema associated with non-infectious uveitis and, based on our pre-IND meeting in 2012 and subsequent written correspondence from the FDA, we expect this single trial to be sufficient to support a 505(b)(2) NDA filing. Our goal is to submit the 505(b)(2) NDA for CLS-1001 in the first half of 2017. Separately, we expect to initiate a Phase 2 clinical trial for CLS-1003 for macular edema associated with RVO before the end of 2014, with data expected in the second half of 2015.
- **Maximizing the commercial potential of our product candidates.** If either CLS-1001 or CLS-1003 is approved, we plan to build a specialized sales force of approximately 30 to 40 representatives to target the approximately 1,700 retinal specialists in the United States. We expect that this sales force would be able to effectively sell more than one of our products, in the event that multiple product candidates are approved for marketing. We have retained the worldwide commercialization rights to

all of our product candidates, except we have granted a third party the exclusive right to commercialize our product candidates in Russia and specified adjacent territories pursuant to a license agreement, as described later in this prospectus. We may pursue additional collaborations with third parties to commercialize any of our drugs approved for marketing outside the United States, either through territorial licenses or distributor relationships.

- **Developing therapies for other back of the eye diseases that can be treated more effectively with SCS injection.** We believe that SCS injection of ocular therapies could have benefits in a variety of other back of the eye indications. Our initial areas of focus for development include:
 - *Advancing our CLS-1002 wet AMD development program.* We are investigating opportunities to improve the treatment of wet AMD through SCS injection of compounds that are commonly used for this disease, such as anti-VEGF drugs, or have shown clinical promise, such as anti-VEGF drugs in combination with anti-PDGF drugs. These therapies are commonly administered by intravitreal injection. We have completed an exploratory Phase 1 clinical trial using an anti-VEGF drug, Avastin, administered into the SCS, rather than intravitreally, in four patients with wet AMD, from which we observed encouraging safety and efficacy results. We are conducting preclinical studies evaluating other compounds with dual anti-VEGF and anti-PDGF activity, with the goal of selecting a lead drug candidate under this program for IND submission in 2015.
 - *Developing a product candidate to treat DME.* DME, like uveitis and RVO, is characterized by an inflammatory aspect. Once the data from the CLS-1003 Phase 2 clinical trial have been analyzed, we intend to initiate clinical trials to develop a product candidate to treat DME modeled after our approach for addressing macular edema associated with RVO.
 - *Developing additional therapies through collaborations with third parties.* We plan to explore collaborations with third parties to develop SCS-administered treatments for back of the eye diseases. These collaborations may take the form of in-licenses of the third parties' drugs or drug candidates for our own development for SCS injection, or out-licenses for third parties to use our intellectual property covering SCS injection as part of the development of their own drugs. We are currently studying several compounds under a research collaboration with Santen Pharmaceutical Co., Ltd.
- **Maintaining and strengthening our intellectual property portfolio.** Our intellectual property strategy aims to control the commercialization of SCS access, with patents and patent applications covering non-surgical SCS delivery methods, novel formulations of drugs, and microinjectors used to access the SCS. Our current patents and patent applications provide us with exclusivity in the United States and major international markets, and are not projected to expire until between 2027 and 2034. We will seek to broaden the scope of and increase the geographic reach of our patent protection throughout the world.

Overview of our Product Candidates

The current development status of our product candidates is summarized in the chart below:

Development Program	Indication	Route of Administration and Active Ingredient	Status and Upcoming Milestones
CLS-1001	Macular edema associated with non-infectious uveitis	SCS injection of CLS-TA	<ul style="list-style-type: none"> • Final results from ongoing Phase 1/2 clinical trial with Triesence expected 1H 2015 • Initiate ~30-patient Phase 2 clinical trial in 2H 2014 with CLS-TA, with data expected 1H 2015 • Initiate ~150-patient single pivotal Phase 3 clinical trial in 1H 2015 with CLS-TA, with 505(b)(2) NDA filing expected 1H 2017
CLS-1003	Macular edema associated with RVO	SCS injection of CLS-TA together with intravitreal injection of anti-VEGF compound	<ul style="list-style-type: none"> • Initiate ~40-patient Phase 2 clinical trial in 2H 2014, with data expected 2H 2015
CLS-1002	Wet AMD	SCS injection of anti-VEGF compound or compound with dual anti-VEGF and anti-PDGF activity	<ul style="list-style-type: none"> • Exploratory Phase 1 clinical trial with Avastin completed outside the U.S. • Selection of lead drug candidate for IND submission expected in 2015

CLS-1001 Program Targeting Macular Edema Associated with Non-infectious Uveitis

The most common treatment for non-infectious uveitis, a condition characterized by inflammation of the choroid and surrounding tissues, involves the use of corticosteroids, such as TA. CLS-1001 consists of an SCS injection of CLS-TA with our microinjector. Under our CLS-1001 program, we are conducting a Phase 1/2 clinical trial in which we are evaluating the safety of SCS injection of Triesence, a TA formulation that is similar to CLS-TA and that has been approved by the FDA to treat non-infectious uveitis. While the trial is primarily a safety and tolerability study, we are also assessing efficacy measures. We intended to enroll approximately 10 patients in this trial. We completed enrollment after dosing the eighth patient. Interim data from the eight patients for whom data is available indicate that SCS injection of the drug has generally been well tolerated, with none of the eight patients having developed cataracts or experienced elevated IOP. At eight weeks after a single SCS injection, the visual acuity of all seven patients who currently have reached week 8 had improved by five or more letters on a standard eye chart over baseline, a clinically meaningful improvement, and the visual acuity of five of these seven patients improved by at least 15 letters. Because CLS-1001 is based on our formulation of TA, and TA is known to be effective in treating uveitis, we believe that our product candidate will be at least as effective as commonly used formulations of TA in treating all aspects of the disease, including the associated macular edema. Our preclinical studies suggest that SCS injection of TA may provide improved bioavailability of drug at the diseased retina and choroid, which may result in faster onset of therapeutic effect and a similar efficacy profile with lower drug amounts required. Our early clinical trial results suggest the potential for CLS-1001 to effectively treat uveitis for at least 90 days following a single SCS injection. Currently used

ocular injections of TA are typically effective for approximately 90 days. In the second half of 2014, we plan to initiate a Phase 2 clinical trial in approximately 30 patients with macular edema associated with non-infectious uveitis to evaluate the safety and efficacy of our CLS-TA administered into the SCS. In the first half of 2015, we also expect to initiate a single pivotal Phase 3 clinical trial with our CLS-TA administered into the SCS in approximately 150 patients with macular edema associated with non-infectious uveitis, with the goal of being able to submit an NDA to the FDA in the first half of 2017. Based on our pre-IND meeting in 2012 and subsequent written correspondence from the FDA, we believe this single pivotal clinical trial will be sufficient to support a 505(b)(2) NDA filing for CLS-1001 for macular edema associated with non-infectious uveitis. If approved, CLS-1001 would be the first drug specifically indicated for macular edema associated with non-infectious uveitis.

CLS-1003 Program Targeting Macular Edema Associated with Retinal Vein Occlusion

Under our CLS-1003 program, we are planning to initiate a Phase 2 clinical trial before the end of 2014 in approximately 40 patients with macular edema associated with RVO, a sight-threatening disorder resulting from the blockage of a retinal vein. The current standard of care for RVO involves monthly intravitreal injections of an anti-VEGF drug, such as Lucentis (ranibizumab), Eylea (aflibercept) or Avastin (bevacizumab). Although studies have shown that corticosteroids effectively address the inflammatory aspect of RVO, they are not used as first-line therapy because they are generally not considered to be as effective as anti-VEGF drugs and also have an unattractive side effect profile when injected intravitreally. In our Phase 2 clinical trial, we will evaluate the safety and efficacy of the injection of our CLS-TA into the SCS together with Eylea injected into the vitreous, as compared to an intravitreal injection of Eylea alone. We believe that SCS administration of our CLS-TA, by addressing the inflammatory aspect of RVO, in combination with intravitreal anti-VEGF treatment, addressing the vascular aspect of RVO, may result in faster onset of therapeutic effect and may be able to improve visual acuity, compared to currently used intravitreal anti-VEGF treatment alone, while also potentially reducing the frequency of necessary anti-VEGF treatments from once every 30 days to once every 90 days. In addition, we believe that the side effect profile of CLS-1003 will be similar to that of current intravitreal anti-VEGF treatments alone, as SCS injections of CLS-TA should allow the corticosteroid to remain substantially localized in the retina and choroid without introducing the side effects that are commonly observed with other routes of administration of corticosteroids.

CLS-1002 Program Targeting Wet Age-Related Macular Degeneration

Under our CLS-1002 program, we are evaluating a number of compounds as potential therapies for the treatment of wet AMD by SCS injection, including compounds with anti-VEGF activity and compounds with dual anti-VEGF and anti-PDGF activity. Wet AMD is a condition involving the abnormal formation and growth of new blood vessels in the choroid, their intrusion into the outer layers of the retina and the leakage of blood and fluid into the retina. The current standard of care for the treatment of wet AMD is intravitreal injection of anti-VEGF drugs as a standalone therapy, as often as seven times per year. We have completed a Phase 1 clinical trial evaluating the safety and efficacy of SCS injection of Avastin, an anti-VEGF drug, in four patients with wet AMD. In this trial, SCS injection of Avastin was observed to be well tolerated, with no treatment-related serious adverse events. In addition, the four patients showed an average improvement in visual acuity of nine letters on a standard eye chart at the end of the eight-week post-treatment observation period. Two of the four patients showed an improvement in visual acuity of at least 13 letters. Because wet AMD manifests in the retina and choroid, we believe that SCS injection of drug

more directly to the retina and choroid may result in faster onset of therapeutic effect as compared to intravitreal injection, and could reduce the necessary frequency of treatment to once every 90 days.

Future Potential Product Candidates

We believe that our SCS-focused approach has the potential for application in treating other back of the eye diseases, and we intend to develop additional product candidates for SCS injection based on the results of our current and planned clinical trials. In addition to uveitis, RVO and wet AMD, there are a number of ophthalmic conditions affecting the retina and choroid with unmet medical need for which SCS injection of therapy may be beneficial. These indications include DME, a common ocular complication of diabetes that results in swelling in the macula and is the primary cause of vision loss associated with diabetes, polypoidal choroidal vasculopathy, or PCV, a disorder resulting from the abnormal growth of new blood vessels in the choroid adjacent to the macula, and geographic atrophy, an advanced form of AMD.

Risks Associated with Our Business

Our business is subject to a number of risks of which you should be aware before making a decision to invest in our common stock. These risks are discussed more fully in the “Risk Factors” section of this prospectus. These risks include the following:

- We have incurred significant losses since our inception, expect to incur losses over the next several years and may never achieve or maintain profitability.
- We are very early in our development efforts and none of our product candidates have completed a clinical trial.
- We have not yet dosed any patients using CLS-TA, our proprietary drug formulation of TA, for our two most advanced clinical programs.
- We have recently revised the design of our microinjector that we intend to utilize with any product candidate for which we ultimately seek marketing approval, but we do not yet have significant experience with this microinjector in humans.
- Clinical drug development involves a lengthy and expensive process, with an uncertain outcome, and we may incur additional costs or experience delays in completing, or ultimately be unable to complete, the development and commercialization of our product candidates.
- We rely, and expect to continue to rely, on third parties to conduct our clinical trials for our product candidates, and those third parties may not perform satisfactorily, including failing to meet deadlines for the completion of such trials.
- If the FDA does not conclude that our product candidates satisfy the requirements for the Section 505(b)(2) NDA regulatory approval pathway, or if the requirements under Section 505(b)(2) are not as we expect, the approval pathway for our product candidates will likely take significantly longer, cost significantly more and entail significantly greater complications and risks than anticipated, and in either case may not be successful.
- Our research and development efforts are focused on the development of product candidates for SCS administration, which is a novel approach and may fail to achieve and sustain market acceptance.
- If we are unable to establish sales, marketing and distribution capabilities for our product candidates, we may not be successful in commercializing those product candidates, if and when they are approved.

- If we are unable to obtain and maintain patent protection for our technology and product candidates, or if our licensors are unable to obtain and maintain patent protection for the technology or product candidates that we license from them, or if the scope of the patent protection obtained is not sufficiently broad, our competitors could develop and commercialize technology and drugs similar or identical to ours, and our ability to successfully commercialize our technology and product candidates may be impaired.
- An active trading market for our common stock may not develop and you may not be able to resell your shares of our common stock at or above the initial offering price, if at all.
- We are an “emerging growth company” and as a result of the reduced disclosure and governance requirements applicable to emerging growth companies, our common stock may be less attractive to investors.

Corporate Information

We were incorporated under the laws of the State of Delaware in May 2011. Our principal executive offices are located at 1220 Old Alpharetta Road, Suite 300, Alpharetta, Georgia 30005 and our telephone number is (678) 270-3631. Our website address is www.clearsidebio.com. The information contained on our website is not incorporated by reference into this prospectus, and you should not consider any information contained on, or that can be accessed through, our website as part of this prospectus or in deciding whether to purchase our common stock.

We have proprietary rights to a number of trademarks used in this prospectus which are important to our business, including Clearside® and the Clearside logo. Solely for convenience, the trademarks and trade names in this prospectus are referred to without the ® and TM symbols, but such references should not be construed as any indicator that their respective owners will not assert, to the fullest extent under applicable law, their rights thereto. All other trademarks, trade names and service marks appearing in this prospectus are the property of their respective owners.

Implications of Being an Emerging Growth Company

As a company with less than \$1 billion in revenue during our last fiscal year, we qualify as an “emerging growth company” as defined in the Jumpstart Our Business Startups Act of 2012, or the JOBS Act. For so long as we remain an emerging growth company, we are permitted and intend to rely on exemptions from some of the reporting requirements that are applicable to other public companies that are not emerging growth companies. These exemptions include:

- being permitted to provide only two years of audited financial statements, in addition to any required unaudited interim financial statements, with correspondingly reduced “Management’s Discussion and Analysis of Financial Condition and Results of Operations” disclosure;
- not being required to comply with the auditor attestation requirements in the assessment of our internal control over financial reporting;
- not being required to comply with any requirement that may be adopted by the Public Company Accounting Oversight Board regarding mandatory audit firm rotation or a supplement to the auditor’s report providing additional information about the audit and the financial statements;
- reduced disclosure obligations regarding executive compensation; and

- not being required to hold a non-binding advisory vote on executive compensation or obtain stockholder approval of any golden parachute payments not previously approved.

We may take advantage of these reporting exemptions until we are no longer an emerging growth company. We will remain an emerging growth company until the earlier of (1) the last day of the fiscal year (a) following the fifth anniversary of the completion of this offering, (b) in which we have total annual gross revenue of at least \$1 billion or (c) in which we are deemed to be a large accelerated filer, which means the market value of our common stock that is held by non-affiliates exceeds \$700 million as of the prior June 30th and (2) the date on which we have issued more than \$1 billion in non-convertible debt during the prior three-year period. We may choose to take advantage of some, but not all, of the available exemptions. We have taken advantage of some reduced reporting burdens in this prospectus. Accordingly, the information contained herein may be different than the information you receive from other public companies in which you hold stock.

In addition, under the JOBS Act, emerging growth companies can delay adopting new or revised accounting standards until such time as those standards apply to private companies. We have irrevocably elected not to avail ourselves of this exemption from new or revised accounting standards and, therefore, we will be subject to the same new or revised accounting standards as other public companies that are not emerging growth companies.

THE OFFERING

Common stock offered by us	shares
Common stock to be outstanding immediately after this offering	shares
Option to purchase additional shares	We have granted the underwriters an option for a period of 30 days from the date of this prospectus to purchase up to additional shares of our common stock.
Use of proceeds	<p>We estimate that the net proceeds to us from this offering, after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us, will be approximately \$ million, assuming the shares are offered at \$ per share, which is the midpoint of the estimated offering price range set forth on the cover page of this prospectus.</p> <p>We anticipate that the majority of the net proceeds from this offering will be used to fund Phase 2 and Phase 3 clinical trials for CLS-1001, as well as a Phase 2 clinical trial for CLS-1003. The remaining proceeds will be used for further development of CLS-1003 and continued research and development of our earlier-stage programs, and for working capital and general corporate purposes. See “Use of Proceeds” for additional information.</p>
Risk factors	You should read the “Risk Factors” section of this prospectus for a discussion of factors to consider carefully before deciding to invest in shares of our common stock.
Proposed NASDAQ Global Market symbol	CLSD

The number of shares of our common stock that will be outstanding after this offering is based on 19,828,234 shares of common stock outstanding as of August 31, 2014, after giving effect to the automatic conversion of 15,564,959 shares of our convertible preferred stock outstanding as of August 31, 2014 and the exercise of warrants to purchase an aggregate of 248,175 shares of common stock that would otherwise expire upon the closing of this offering, and excludes:

- 1,685,902 shares of our common stock issuable upon the exercise of stock options outstanding under our 2011 stock incentive plan as of August 31, 2014, at a weighted average exercise price of \$0.21 per share;
- 16,550 shares of our common stock issuable upon exercise of a warrant outstanding as of August 31, 2014, at an exercise price of \$1.81 per share; and
- shares of our common stock reserved for future issuance under our equity incentive plans following this offering.

Except as otherwise indicated herein, all information in this prospectus, including the number of shares that will be outstanding after this offering, assumes or gives effect to:

- -for- reverse stock split of our common stock expected to be completed prior to the completion of this offering;
- the conversion of all outstanding shares of our convertible preferred stock into an aggregate of 15,564,959 shares of our common stock, which will occur automatically upon the closing of this offering;
- the exercise of all outstanding warrants to purchase shares of common stock that will otherwise expire upon the closing of this offering for an aggregate of 248,175 shares of common stock;
- the conversion of an outstanding warrant to purchase shares of Series A-1 convertible preferred stock into a warrant to purchase 16,550 shares of common stock upon the closing of this offering; and
- no exercise of the underwriters' option to purchase additional shares of common stock.

Summary Financial Data

In the tables below, we provide you with our summary financial data for the periods indicated. We have derived the following summary of our statement of operations data for the years ended December 31, 2012 and 2013 from our audited financial statements appearing elsewhere in this prospectus. We have derived the following summary of our statement of operations data for the six months ended June 30, 2013 and 2014 and for the period from May 26, 2011 (date of inception) through June 30, 2014 and our balance sheet data as of June 30, 2014 from our unaudited interim financial statements appearing elsewhere in this prospectus.

The financial data for the six months ended June 30, 2013 and 2014, for the period from May 26, 2011 (date of inception) through June 30, 2014 and as of June 30, 2014 includes, in the opinion of our management, all adjustments, consisting only of normal recurring adjustments that are necessary for a fair presentation of our financial position and results of operations for these periods. Our historical results are not necessarily indicative of the results to be expected in the future, and our operating results for the six months ended June 30, 2014 are not necessarily indicative of the results that may be expected for the entire year ending December 31, 2014 or any other future period.

You should read this summary financial data together with the historical financial statements and related notes to those statements, as well as “Management’s Discussion and Analysis of Financial Condition and Results of Operations,” which are included elsewhere in this prospectus.

	Year Ended December 31,		Six Months Ended June 30,		Period From May 26, 2011 (Date of Inception) to June 30, 2014
	2012	2013	2013	2014	
(in thousands, except share and per share data)					
Statement of Operations Data:					
Costs and expenses:					
Research and development	\$ 2,354	\$ 5,045	\$ 2,204	\$ 2,562	\$ 10,168
General and administrative	1,575	2,193	1,101	1,731	5,640
Total costs and expenses	<u>3,929</u>	<u>7,238</u>	<u>3,305</u>	<u>4,293</u>	<u>15,808</u>
Loss from operations	(3,929)	(7,238)	(3,305)	(4,293)	(15,808)
Other income (expense):					
Interest expense	(3)	(23)	(6)	(99)	(127)
Interest income	1	7	5	1	9
Total other expense	<u>(2)</u>	<u>(16)</u>	<u>(1)</u>	<u>(98)</u>	<u>(118)</u>
Net loss	<u>\$ (3,931)</u>	<u>\$ (7,254)</u>	<u>\$ (3,306)</u>	<u>\$ (4,391)</u>	<u>\$ (15,926)</u>
Net loss per share of common stock — basic and diluted	<u>\$ (2.12)</u>	<u>\$ (2.45)</u>	<u>\$ (1.21)</u>	<u>\$ (1.20)</u>	<u>\$ (6.78)</u>
Weighted average shares outstanding, basic and diluted	1,853,423	2,956,285	2,728,321	3,673,629	2,349,838
Pro forma net loss per share — basic and diluted		<u>\$ (0.58)</u>		<u>\$ (0.33)</u>	
Pro forma weighted average shares outstanding — basic and diluted		12,512,042		13,229,386	

The following table presents our summary balance sheet data as of June 30, 2014:

- on an actual basis;
- on a pro forma basis to give effect to:
 - our issuance of an aggregate of 6,009,202 shares of Series B convertible preferred stock in August 2014 at a purchase price of \$2.70 per share, which included the conversion of principal and interest under outstanding convertible promissory notes with a principal balance of \$3.1 million into shares of Series B convertible preferred stock, and our receipt of \$13.0 million in net cash proceeds therefrom;
 - the conversion of all outstanding shares of our convertible preferred stock, including the 6,009,202 shares of Series B convertible preferred stock issued in August 2014, into an aggregate of 15,564,959 shares of our common stock, which will occur automatically upon the closing of this offering; and
 - the exercise of all outstanding warrants to purchase shares of common stock that will otherwise expire upon the closing of this offering for an aggregate of 248,175 shares of common stock; and
- on a pro forma as adjusted basis to give further effect to our sale of _____ shares of common stock in this offering at an assumed initial public offering price of \$ _____ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us.

	As of June 30, 2014		
	Actual	Pro forma	Pro forma as adjusted
	(in thousands)		
Balance Sheet Data:			
Cash and cash equivalents	\$ 1,453	\$	\$
Total assets	2,862		
Total liabilities	5,622		
Total convertible preferred stock	11,883		
Total stockholders' equity (deficit)	(14,643)		

The pro forma as adjusted information presented in the summary balance sheet data is illustrative only and will change based on the actual initial public offering price and other terms of this offering determined at pricing. Each \$1.00 increase or decrease in the assumed initial public offering price of \$ _____ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, would increase or decrease each of cash and cash equivalents, total assets and total stockholders' equity on a pro forma as adjusted basis by \$ _____ million, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same. Similarly, each increase or decrease of 1.0 million shares offered by us at the assumed initial public offering price would increase or decrease each of cash and cash equivalents, total assets and total stockholders' equity on a pro forma as adjusted basis by \$ _____ million.

RISK FACTORS

Investing in our common stock involves a high degree of risk. Before you invest in our common stock, you should carefully consider the following risks, as well as general economic and business risks, and all of the other information contained in this prospectus. Any of the following risks could have a material adverse effect on our business, operating results and financial condition and cause the trading price of our common stock to decline, which would cause you to lose all or part of your investment. When determining whether to invest, you should also refer to the other information contained in this prospectus, including our financial statements and the related notes thereto.

Risks Related to Our Financial Position and Capital Needs

We have incurred significant losses since our inception. We expect to incur losses over the next several years and may never achieve or maintain profitability.

Since our inception through June 30, 2014, we have incurred net losses of \$15.9 million. We incurred net losses of \$3.9 million, \$7.3 million and \$4.4 million for the years ended December 31, 2012 and 2013 and the six months ended June 30, 2014, respectively. As of June 30, 2014, we had a deficit accumulated during the development stage of \$15.9 million. We financed our operations through June 30, 2014 with approximately \$14.7 million of net cash proceeds raised in private placements of convertible preferred stock and convertible promissory notes, and in August 2014 we raised an additional \$13.0 million of net cash proceeds through a private placement of convertible preferred stock.

We have devoted substantially all of our financial resources and efforts to research and development, including preclinical studies and clinical trials. We are still in the early stages of development of our product candidates, and we have not completed development of any drugs. We expect to continue to incur significant expenses and operating losses over the next several years. Our net losses may fluctuate significantly from quarter to quarter and year to year. We anticipate that our expenses will increase substantially as we:

- continue our ongoing Phase 1/2 clinical trial evaluating the safety of SCS administration of a commercially available formulation of TA with our microinjector, and initiate clinical trials using CLS-TA in our CLS-1001 and CLS-1003 drug programs;
- continue the research and development of our other product candidates;
- seek to discover and develop additional product candidates;
- seek regulatory approvals for any product candidates that successfully complete clinical trials;
- ultimately establish a sales, marketing and distribution infrastructure and scale up external manufacturing capabilities to commercialize any drugs for which we may obtain regulatory approval;
- maintain, expand and protect our intellectual property portfolio;
- hire additional clinical, manufacturing and scientific personnel;
- add operational, financial and management information systems and personnel, including personnel to support our drug development and planned future commercialization efforts; and
- incur additional legal, accounting and other expenses in operating as a public company.

[Table of Contents](#)

To become and remain profitable, we must succeed in developing and eventually commercializing drugs that generate significant revenue. This will require us to be successful in a range of challenging activities, including completing preclinical testing and clinical trials of our product candidates, obtaining regulatory approval and manufacturing, marketing and selling any products for which we may obtain regulatory approval, as well as discovering and developing additional product candidates. We are only in the preliminary stages of most of these activities. We may never succeed in these activities and, even if we do, may never generate revenues that are significant enough to achieve profitability.

Because of the numerous risks and uncertainties associated with drug development, we are unable to accurately predict the timing or amount of expenses or when, or if, we will be able to achieve profitability. If we are required by regulatory authorities to perform studies in addition to those currently expected, or if there are any delays in the initiation and completion of our clinical trials or the development of any of our product candidates, our expenses could increase.

Even if we achieve profitability, we may not be able to sustain or increase profitability on a quarterly or annual basis. Our failure to become and remain profitable would depress our value and could impair our ability to raise capital, expand our business, maintain our research and development efforts, diversify our product offerings or continue our operations. A decline in our value could also cause you to lose all or part of your investment.

We will need substantial additional funding to meet our financial obligations and to pursue our business objectives. If we are unable to raise capital when needed, we could be forced to delay, reduce or altogether cease our drug development programs or commercialization efforts.

We believe that the net proceeds from this offering, together with our existing cash and cash equivalents, will enable us to fund our operating expenses and capital expenditure requirements for at least the next 12 months. However, we will need to obtain substantial additional funding in connection with our continuing operations. Our future capital requirements will depend on many factors, including:

- the progress and results of our planned Phase 2 and Phase 3 clinical trials, both for CLS-1001 and for CLS-1003;
- the scope, progress, results and costs of preclinical development, laboratory testing and clinical trials for our other product candidates;
- the number and development requirements of other product candidates that we may pursue;
- the costs, timing and outcome of regulatory review of our product candidates;
- the costs and timing of future commercialization activities, including product manufacturing, marketing, sales and distribution, for any of our product candidates for which we receive marketing approval;
- the revenue, if any, received from commercial sales of our product candidates for which we receive marketing approval;
- the costs and timing of preparing, filing and prosecuting patent applications, maintaining and enforcing our intellectual property rights and defending any intellectual property-related claims; and
- the extent to which we acquire or in-license other product candidates and technologies.

[Table of Contents](#)

Identifying potential product candidates and conducting preclinical testing and clinical trials is a time-consuming, expensive and uncertain process that takes years to complete, and we may never generate the necessary data or results required to obtain regulatory approval and achieve product sales. In addition, our product candidates, if approved, may not achieve commercial success. Our commercial revenues, if any, will be derived from sales of drugs that we do not expect to be commercially available for several years, if at all. Accordingly, we will need to continue to rely on additional financing to achieve our business objectives. Adequate additional financing may not be available to us on acceptable terms, or at all. In addition, we may seek additional capital due to favorable market conditions or strategic considerations even if we believe we have sufficient funds for our current or future operating plans. If we are unable to raise capital when needed or on attractive terms, we could be forced to delay, reduce or altogether cease our research and development programs or future commercialization efforts.

Raising additional capital may cause dilution to our stockholders, including purchasers of common stock in this offering, restrict our operations or require us to relinquish rights to our technologies or product candidates.

Until such time, if ever, as we can generate substantial product revenue, we expect to finance our cash needs through a combination of equity offerings, debt financings and potential collaboration, license and development agreements. We do not currently have any committed external source of funds. To the extent that we raise additional capital through the sale of equity or convertible debt securities, your ownership interest will be diluted, and the terms of these securities may include liquidation or other preferences that adversely affect your rights as a common stockholder. Debt financing and preferred equity financing, if available, may involve agreements that include covenants limiting or restricting our ability to take specific actions, such as incurring additional debt, making capital expenditures or declaring dividends.

If we raise additional funds through collaborations, strategic alliances or marketing, distribution or licensing arrangements with third parties, we may be required to relinquish valuable rights to our technologies, future revenue streams, research programs or product candidates or to grant licenses on terms that may not be favorable to us. If we are unable to raise additional funds through equity or debt financings when needed, we may be required to delay, limit, reduce or terminate our drug development or future commercialization efforts or grant rights to develop and market product candidates that we would otherwise prefer to develop and market ourselves.

Our operating history may make it difficult for you to evaluate the success of our business to date and to assess our future viability.

We commenced active operations in 2012, and our operations to date have been largely focused on raising capital, identifying potential product candidates, broadening our expertise in the development of our product candidates and our microinjector, undertaking preclinical studies and conducting early clinical trials. We have not yet demonstrated an ability to successfully complete later-stage clinical trials, obtain regulatory approvals, manufacture a commercial scale product or arrange for a third party to do so on our behalf, or conduct sales and marketing activities necessary for successful commercialization. Consequently, any predictions you make about our future success or viability may not be as accurate as they could be if we had a longer operating history.

We may encounter unforeseen expenses, difficulties, complications, delays and other known or unknown factors in achieving our business objectives. We will need to transition at some point from a

company with a research and development focus to a company capable of supporting commercial activities. We may not be successful in such a transition.

We expect our financial condition and operating results to continue to fluctuate significantly from quarter to quarter and year to year due to a variety of factors, many of which are beyond our control. Accordingly, you should not rely upon the results of any quarterly or annual periods as indications of future operating performance.

Risks Related to the Development of Our Product Candidates

Our research and development efforts are focused on the development of product candidates for SCS administration, which is a novel approach and may fail to achieve and sustain market acceptance.

Administering drugs into the SCS is a novel approach for ophthalmic therapies, and there is no guarantee this approach will be accepted by physicians, patients or third-party payors. We believe we are the first and only company developing drugs specifically for non-surgical SCS administration. The scientific evidence to support the feasibility of developing drugs based on this approach is both preliminary and limited. Although preclinical studies suggest that SCS administration of drugs, such as CLS-1001, may be effective at treating back of the eye diseases, to date no company has developed a drug for administration through the SCS that has received marketing approval.

Additionally, we have very limited clinical experience in SCS drug administration. Our most advanced clinical development program, CLS-1001 for the treatment of macular edema associated with non-infectious uveitis, is currently in a Phase 1/2 clinical trial in which we have dosed eight patients to date with a commercially available formulation of the active ingredient in CLS-TA, only four of which have completed the 26 week post-treatment observation period. In addition to the preliminary data from the eight patients for whom data is available in this trial, our only other human data involving SCS drug administration is from an exploratory Phase 1 clinical trial in four patients with wet AMD, which was conducted in Mexico without an IND from the FDA. To date, between these two trials, we have only dosed a total of twelve humans using our proprietary SCS injection method. Therefore, we cannot guarantee that SCS administration of drugs will prove in clinical trials to be a safe or effective approach for treating back of the eye diseases in humans.

Even if we are able to achieve marketing approval for one of our product candidates, the novelty of SCS injection may make it difficult to demonstrate to physicians and third-party payors that SCS administration of our drugs is the appropriate approach for treating diseases such as non-infectious uveitis, RVO and wet AMD and is superior to the current standard of care. Further, if we are not successful in conveying to physicians, patients and third-party payors that the administration of our drug candidates with our proprietary microinjector improves patient outcomes, we may experience reluctance, or refusal, on the part of physicians to order and use, and third-party payors to cover and provide adequate payment for, our product candidates. Additionally, in some cases, our product candidates will complement the current standard of care, rather than serve as a replacement for the current standard of care. Therefore, we may encounter significant difficulty in gaining broad market acceptance by physicians, third-party payors and potential patients using CLS-1001, CLS-1003 and our other product candidates.

We are very early in our development efforts and none of our product candidates have completed a clinical trial. Our most advanced program, CLS-1001, is currently in a Phase 1/2 clinical trial. For our second program, CLS-1003, we plan to initiate our first clinical trial, a Phase 2 trial, before the end of

2014. All of our other product candidates are still in preclinical development. If we are unable to commercialize our product candidates or if we experience significant delays in doing so, our business may be harmed.

We are very early in our development efforts. For our most advanced clinical development program, CLS-1001 for the treatment of macular edema associated with non-infectious uveitis, we are currently conducting a Phase 1/2 clinical trial in which we have dosed eight patients to date with a commercially available formulation of the active ingredient in CLS-TA, only four of whom have completed the 26 week post-treatment observation period, and we plan to initiate a Phase 2 trial administering CLS-TA with our microinjector in the second half of 2014. For our second clinical development program, CLS-1003 for the treatment of macular edema associated with RVO, we also plan to initiate our first clinical trial, a Phase 2 trial, administering CLS-TA to patients before the end of 2014. In addition to the preliminary data from the eight patients for whom data is available in our ongoing Phase 1/2 clinical trial, our only other human data is from an exploratory Phase 1 clinical trial conducted in Mexico without an IND from the FDA for our CLS-1002 program. Given our limited human experience with our clinical programs, the successful development of any of our product candidates is extremely uncertain, and we cannot guarantee that we will be successful in developing any of our product candidates.

Further, the FDA may conclude that our clinical trials are not sufficient to support approval of our product candidates. For example, we are planning to initiate our clinical development program for CLS-1003 with a Phase 2 clinical trial without having conducted a separate Phase 1 clinical trial for this program. We believe that we will be able to rely on our ongoing toxicology study of CLS-TA in rabbits, the preclinical studies and safety data generated by our ongoing Phase 1/2 clinical trial for CLS-1001 and other supportive literature to be able proceed directly into the Phase 2 clinical trial for CLS-1003, which has the same active pharmaceutical ingredient, CLS-TA, as CLS-1001. However, we have not yet confirmed this approach with the FDA, and the FDA may require that we conduct additional safety studies or trials before we are able to commence clinical trials under an IND for the CLS-1003 program. For our CLS-1002 program, the only clinical trial we have conducted to date is the Phase 1 exploratory trial in Mexico. Although the exploratory trial was conducted in accordance with good clinical practices and had approval and oversight of institutional review boards and institutional ethics committees, the FDA could conclude that we may not rely on the results of the trial conducted in Mexico as part of our regulatory application seeking marketing approval for CLS-1002.

We have not completed the development of any product candidates, we currently generate no revenues from sales of any products and we may never be able to develop a marketable product. We have invested substantially all of our efforts and financial resources in the development of our proprietary microinjector for SCS administration of drugs and the identification of potential drug candidates using that technology. Our ability to generate revenue from our product candidates, which we do not expect will occur for several years, if ever, will depend heavily on their successful development and eventual commercialization. The success of those product candidates will depend on several factors, including the following:

- successful completion of preclinical studies and requisite clinical trials;
- receipt of marketing approvals from applicable regulatory authorities;
- obtaining marketing approvals with labeling for sufficiently broad patient populations and indications, without unduly restrictive distribution limitations or safety warnings;
- obtaining and maintaining patent, trademark and trade secret protection and regulatory exclusivity for our product candidates;

[Table of Contents](#)

- making arrangements with third-party manufacturers for, or establishing, commercial manufacturing capabilities;
- launching commercial sales of products, if and when approved, whether alone or in collaboration with others;
- successful training of physicians in the proper use of our microinjector to administer drugs into the SCS;
- acceptance of the therapies and of the concept of SCS administration of drugs, if and when approved, by physicians, patients and third-party payors;
- competing effectively with other therapies;
- obtaining and maintaining healthcare coverage and adequate reimbursement;
- protecting our rights in our intellectual property portfolio; and
- maintaining a continued acceptable safety profile of the drugs and microinjector following approval.

If we do not achieve one or more of these factors in a timely manner or at all, we could experience significant delays or an inability to successfully commercialize our product candidates, which would materially harm our business.

We have not yet dosed any patients using CLS-TA, our proprietary drug formulation of TA, for our two most advanced clinical programs.

To date, our clinical trial experience has involved the SCS injection of commercially available therapeutic agents. In our ongoing Phase 1/2 trial in our uveitis program, we are administering Triesence, a commercially available TA formulation, via SCS injection. However, in future clinical trials of our most advanced product candidates, including our planned Phase 2 clinical trials of CLS-1001 and CLS-1003, we intend to treat patients with CLS-TA, our proprietary drug formulation of TA. Although we have completed preclinical studies comparing CLS-TA to Triesence, we have never dosed this formulation in a human patient. Therefore, in addition to the risks associated with advancing our uveitis program from smaller scale to larger scale clinical trials and initiating our clinical trial program in RVO, there is also a risk that CLS-TA may have different pharmacokinetic or safety profiles in human subjects than the TA formulations that we have administered to date in animals. If CLS-TA does not exhibit similar safety and efficacy profiles to that of TA in humans, our ability to develop CLS-1001 and CLS-1003 may be harmed. Additionally, if the safety and efficacy profile of CLS-TA is not sufficiently comparable to that of previously commercialized TA formulations, we may be unable to rely on the Section 505(b)(2) regulatory approval pathway for CLS-1001 and CLS-1003, which would significantly lengthen our development process and the cost of developing and commercializing our product candidates.

The administration of CLS-1003 as a therapy together with anti-VEGF drugs for the treatment of macular edema associated with RVO is a novel approach and may fail to be successful.

We are developing our second drug program, CLS-1003, as a therapy to complement the current standard of care in the treatment of patients experiencing macular edema associated with RVO, with the goal of reducing current required monthly anti-vascular endothelial growth factor, or anti-VEGF, injections to quarterly injections. The scientific evidence to support the potential efficacy of this treatment approach is limited and based on third party clinical trials studying intravitreal injections of steroids in patients with

[Table of Contents](#)

RVO, which, although effective in reducing edema, has been associated with side effects. While our preliminary clinical trial experience involving the SCS injection of steroids suggest that these adverse side effects may be avoided using SCS injection, to date no company has explored this concomitant treatment approach in clinical trials or preclinical studies.

Even if we are able to successfully develop, and achieve marketing approval of, CLS-1003, it may be difficult to demonstrate to physicians and third-party payors that the administration of CLS-1003 concomitantly with anti-VEGF drugs, and the reduction in frequency of anti-VEGF treatments, is the appropriate approach for treating RVO and is superior to the current standard of care. Further, if we are not successful in conveying to physicians, patients and third-party payors that the administration of CLS-1003 concomitantly with anti-VEGF drugs improves patient outcomes, we may experience reluctance, or refusal, on the part of physicians to order and use, and third-party payors to pay for, CLS-1003. Therefore, we may encounter significant difficulty in gaining broad market acceptance by physicians, third-party payors and potential patients using CLS-1003.

We have recently revised the design of our microinjector that we intend to utilize with any product candidate for which we ultimately seek marketing approval, but we do not yet have significant experience with this microinjector in humans.

We used an earlier design of our microinjector in our preclinical studies and clinical trials, including the first four patients dosed in our ongoing Phase 1/2 clinical trial. However, we have recently completed the design of a modified microinjector that we used in the most recent four patients in the Phase 1/2 trial, and we intend to use this design in our planned pivotal Phase 3 clinical trial of CLS-1001 and, if successful, ultimately include in the NDA of any drug for which we seek marketing approval. Accordingly, in addition to the risks associated with drug development, we are also subject to the risks associated with developing the microinjector. For example, in our ongoing Phase 1/2 clinical trial, the needle of our earlier microinjector was not long enough to penetrate the scleras of two patients screened for the trial. If we encounter similar limitations with this design, or if it does not function properly in any way, we could be required to expend significant additional time and resources to redesign our microinjector, which would delay or compromise our drug development efforts. Additionally, our ability to successfully commercialize our product candidates will depend on retinal specialists being comfortable with the design and functionality of our microinjector. If, for any reason, retinal specialists were unsatisfied with the form or function of our microinjector, it would harm the market acceptance and potential commercial success of our product candidates, if any, that receive marketing approval.

We may not be successful in our efforts to build a pipeline of product candidates.

A key element of our strategy is to build a pipeline of product candidates for SCS administration and to progress these product candidates through clinical development for the treatment of a variety of diseases of the back of the eye. We may not be able to develop product candidates that are safe and effective. Even if we are successful in continuing to build our pipeline, the potential product candidates that we identify may not be suitable for clinical development, including as a result of being shown to have significant side effects or other characteristics that indicate that they are unlikely to be drugs that will receive marketing approval or achieve market acceptance. If we do not successfully develop and commercialize product candidates based upon our approach, we will not be able to obtain product revenue in future periods, which could significantly harm our financial position and adversely affect our stock price.

Clinical drug development involves a lengthy and expensive process, with an uncertain outcome. We may incur additional costs or experience delays in completing, or ultimately be unable to complete, the development and commercialization of our product candidates.

The risk of failure for our product candidates is high. It is impossible to predict when or if any of our product candidates will prove effective or safe in humans and will receive regulatory approval. Before obtaining marketing approval from regulatory authorities for the sale of any product candidate, we must complete preclinical development and then conduct extensive clinical trials to demonstrate the safety and efficacy of our product candidates in humans. Clinical testing is expensive, difficult to design and implement, can take many years to complete and is uncertain as to outcome. A failure of one or more clinical trials can occur at any stage of testing. The outcome of preclinical testing and early clinical trials may not be predictive of the success of later clinical trials, and interim results of a clinical trial do not necessarily predict final results. For example, the preliminary data obtained from our Phase 1/2 clinical trial for our CLS-1001 program is based on only eight patients and might not be achieved in any other patients treated with CLS-1001. Moreover, preclinical and clinical data are often susceptible to varying interpretations and analyses, and many companies that have believed their product candidates performed satisfactorily in preclinical studies and clinical trials have nonetheless failed to obtain marketing approval of their products.

We may experience numerous unforeseen events during, or as a result of, clinical trials that could delay or prevent our ability to receive marketing approval or commercialize our product candidates, including:

- regulators or institutional review boards may not authorize us or our investigators to commence a clinical trial or conduct a clinical trial at a prospective trial site;
- we may experience delays in reaching, or fail to reach, agreement on acceptable clinical trial contracts or clinical trial protocols with prospective trial sites;
- clinical trials of our product candidates may produce negative or inconclusive results, including failure to demonstrate statistical significance, and we may decide, or regulators may require us, to conduct additional clinical trials or abandon product development programs;
- the number of patients required for clinical trials of our product candidates may be larger than we anticipate, enrollment in these clinical trials may be slower than we anticipate or participants may drop out of these clinical trials at a higher rate than we anticipate;
- our third-party contractors may fail to comply with regulatory requirements or meet their contractual obligations to us in a timely manner, or at all;
- regulators or institutional review boards may require that we or our investigators suspend or terminate clinical research for various reasons, including noncompliance with regulatory requirements or a finding that the participants are being exposed to unacceptable health risks;
- the cost of clinical trials of our product candidates may be greater than we anticipate;
- the supply or quality of our product candidates or other materials necessary to conduct clinical trials of our product candidates may be insufficient or inadequate; and
- our product candidates may have undesirable side effects or other unexpected characteristics, causing us or our investigators, regulators or institutional review boards to suspend or terminate the trials.

[Table of Contents](#)

If we are required to conduct additional clinical trials or other testing of our product candidates beyond those that we currently contemplate, if we are unable to successfully complete clinical trials of our product candidates or other testing, if the results of these trials or tests are not positive or are only modestly positive or if there are safety concerns, we may:

- be delayed in obtaining marketing approval for our product candidates;
- not obtain marketing approval at all;
- obtain approval for indications or patient populations that are not as broad as intended or desired;
- obtain approval with labeling that includes significant use or distribution restrictions or safety warnings;
- be subject to additional post-marketing testing requirements; or
- have the product removed from the market after obtaining marketing approval.

Our drug development costs may also increase if we experience delays in testing or marketing approvals. We do not know whether any of our preclinical studies or clinical trials will begin as planned, will need to be restructured or will be completed on schedule, or at all. Significant preclinical study or clinical trial delays also could shorten any periods during which we may have the exclusive right to commercialize our product candidates or allow our competitors to bring products to market before we do and impair our ability to successfully commercialize our product candidates.

If we experience delays or difficulties in the enrollment of patients in clinical trials, our receipt of necessary regulatory approvals could be delayed or prevented.

We may not be able to initiate or continue clinical trials for our product candidates if we are unable to locate and enroll a sufficient number of eligible patients to participate in these trials as required by the FDA or similar regulatory authorities outside the United States. For example, we began enrolling patients in our Phase 1/2 uveitis trial in July 2013 and intended to enroll approximately 10 patients in the trial, but we completed enrollment after dosing the eighth patient in July 2014. We have very little experience enrolling patients in our clinical trials, and cannot predict how successful we will be at enrolling patients in future clinical trials. If we are not successful at enrolling patients in one clinical trial, it may affect when we are able to initiate our next clinical trial, which could result in significant delays in our efforts to pursue regulatory approval of and commercialize our product candidates. In addition, some of our competitors have ongoing clinical trials to treat the same indications as our product candidates, and patients who would otherwise be eligible for our clinical trials may instead enroll in clinical trials of our competitors. Patient enrollment is affected by other factors including:

- the severity of the disease under investigation;
- the eligibility criteria for the study in question;
- the perceived risks and benefits of the product candidate under study;
- the availability of drugs approved to treat the diseases under study;
- the efforts to facilitate timely enrollment in clinical trials;
- the patient referral practices of physicians;
- the ability to monitor patients adequately during and after treatment; and
- the proximity and availability of clinical trial sites for prospective patients.

[Table of Contents](#)

Our inability to enroll a sufficient number of patients for clinical trials would result in significant delays and could require us to abandon one or more clinical trials altogether. Enrollment delays in these clinical trials may result in increased development costs for our product candidates, which could cause our value to decline and limit our ability to obtain additional financing.

If serious adverse or unacceptable side effects are identified during the development of our product candidates, we may need to abandon or limit our development of some of our product candidates.

If our product candidates are associated with side effects in clinical trials or have characteristics that are unexpected, we may need to abandon their development or limit development to more narrow uses or subpopulations in which the side effects or other characteristics are less prevalent, less severe or more acceptable from a risk-benefit perspective. Many product candidates that initially showed promise in early stage testing have later been found to cause side effects that prevented further development of the product candidate.

We may expend our limited resources to pursue a particular product candidate or indication and fail to capitalize on product candidates or indications that may be more profitable or for which there is a greater likelihood of success.

Because we have limited financial and management resources, we focus on research programs and product candidates that we identify for specific indications. As a result, we may forego or delay pursuit of opportunities with other product candidates or for other indications that later prove to have greater commercial potential. Our resource allocation decisions may cause us to fail to capitalize on viable commercial drugs or profitable market opportunities. Our spending on current and future research and development programs and product candidates for specific indications may not yield any commercially viable products. If we do not accurately evaluate the commercial potential or target market for a particular product candidate, we may relinquish valuable rights to that product candidate through collaboration, licensing or other royalty arrangements in cases in which it would have been more advantageous for us to retain sole development and commercialization rights to such product candidate.

Risks Related to Our Dependence on Third Parties

We expect to rely on third parties to conduct our clinical trials for our product candidates, and those third parties may not perform satisfactorily, including failing to meet deadlines for the completion of such trials.

We expect to engage clinical research organizations, or CROs, for our planned Phase 2 and Phase 3 clinical trials for CLS-1001 and CLS-1003. We also expect to engage CROs for any of our other product candidates that may progress to clinical development. We expect to rely on CROs, as well as other third parties, such as clinical data management organizations, medical institutions and clinical investigators, to conduct those clinical trials. Agreements with such third parties might terminate for a variety of reasons, including a failure to perform by the third parties. If we need to enter into alternative arrangements, that would delay our drug development activities.

Our reliance on these third parties for research and development activities will reduce our control over these activities but will not relieve us of our responsibilities. For example, we will remain responsible for ensuring that each of our clinical trials is conducted in accordance with the general investigational plan and protocols for the trial. Moreover, the FDA requires us to comply with regulatory standards, commonly

[Table of Contents](#)

referred to as good clinical practices, or GCPs, for conducting, recording and reporting the results of clinical trials to assure that data and reported results are credible and accurate and that the rights, integrity and confidentiality of trial participants are protected. We also are required to register ongoing clinical trials and post the results of completed clinical trials on a government-sponsored database, ClinicalTrials.gov, within specified timeframes. Failure to do so can result in FDA refusal to approve applications based on the clinical data, fines, adverse publicity and civil and criminal sanctions.

Furthermore, these third parties may also have relationships with other entities, some of which may be our competitors. If these third parties do not successfully carry out their contractual duties, meet expected deadlines or conduct our clinical trials in accordance with regulatory requirements or our stated protocols, we will not be able to obtain, or may be delayed in obtaining, marketing approvals for our product candidates and will not be able to, or may be delayed in our efforts to, successfully commercialize our product candidates.

In addition, principal investigators for our clinical trials may serve as scientific advisors or consultants to us from time to time and may receive cash or equity compensation in connection with such services. If these relationships and any related compensation result in perceived or actual conflicts of interest, or the FDA concludes that the financial relationship may have affected the interpretation of the trial, the integrity of the data generated at the applicable clinical trial site may be questioned and the utility of the clinical trial itself may be jeopardized, which could result in the delay or rejection of any new drug application, or NDA, we submit by the FDA. Any such delay or rejection could prevent us from commercializing CLS-1001, CLS-1003 or our other future product candidates.

We also expect to rely on other third parties to store and distribute product supplies for our clinical trials. Any performance failure on the part of our distributors could delay clinical development or marketing approval of our product candidates or commercialization of our products, producing additional losses and depriving us of potential product revenue.

We contract with third parties for the manufacture of CLS-TA and our microinjector for preclinical and clinical testing and expect to continue to do so for commercialization. This reliance on third parties increases the risk that we will not have sufficient quantities of CLS-TA and our microinjector or such quantities at an acceptable cost, which could delay, prevent or impair our development or commercialization efforts.

We do not have any manufacturing facilities or personnel. We currently procure the active pharmaceutical ingredient in CLS-TA on a purchase order basis from a third-party manufacturer, and we anticipate entering into commercial supply agreements with this or another manufacturer at a later date. In addition, we obtain each of the components of our SCS microinjector on a purchase order basis from third-party suppliers. Some of our current suppliers are based outside of the United States. We expect to continue to rely on third parties as we proceed with preclinical and clinical testing using CLS-TA with our microinjector, as well as for commercial manufacture, if any of our product candidates receives marketing approval. This reliance on third parties increases the risk that we will not have sufficient quantities of CLS-TA and our microinjector or such quantities at an acceptable cost or quality, which could delay, prevent or impair our ability to timely conduct our clinical trials or our other development or commercialization efforts.

We also expect to rely on third-party manufacturers or collaborators for the manufacture of commercial supply of any other product candidates for which we obtain marketing approval. We may be unable to establish any agreements with third-party manufacturers or collaborators or to do so on acceptable terms.

[Table of Contents](#)

Even if we are able to establish agreements with third-party manufacturers or collaborators, reliance on third-party manufacturers or collaborators entails additional risks, including:

- reliance on the third party for regulatory compliance and quality assurance;
- the possible breach of the manufacturing agreement by the third party;
- the possible misappropriation of our proprietary information, including our trade secrets and know-how; and
- the possible termination or nonrenewal of the agreement by the third party at a time that is costly or inconvenient for us.

Our product candidates, including our proprietary drug formulations packaged together with our microinjector, are drug/device combination products that will be regulated under the drug regulations of the Federal Food, Drug, and Cosmetic Act, or FDCA, based on their primary mode of action as drugs. Third-party manufacturers may not be able to comply with current good manufacturing practices, or cGMP, regulations, regulations applicable to drug/device combination products, including applicable provisions of the FDA's drug cGMP regulations, device cGMP requirements embodied in the Quality System Regulation, or QSR, or similar regulatory requirements outside the United States. Our failure, or the failure of our third-party manufacturers, to comply with applicable regulations could result in sanctions being imposed on us, including clinical holds, fines, injunctions, civil penalties, delays, suspension or withdrawal of approvals, license revocation, seizures or recalls of product candidates, operating restrictions and criminal prosecutions, any of which could significantly affect supplies of our product candidates.

Our product candidates that we may develop may compete with other drugs and devices for access to manufacturing facilities. There are a limited number of manufacturers that operate under the drug and device cGMP regulations applicable to our product candidates and that might be capable of manufacturing for us. Any performance failure on the part of our existing or future manufacturers could delay clinical development or marketing approval. We do not currently have arrangements in place for redundant supply or a second source for bulk drug substance or our microinjector. If our current contract manufacturers cannot perform as agreed, we may be required to replace such manufacturers. Although we believe that there are several potential alternative manufacturers who could manufacture our drugs and the components of our microinjector, we may incur added costs and delays in identifying and qualifying any such replacement. For example, the FDA could require supplemental data if a new supplier is relied upon for the active pharmaceutical ingredient used in our products. Any interruption or delay in the supply of components and materials, or our inability to obtain components or materials from alternate sources at acceptable prices in a timely manner, could impair our ability to meet the demand of our customers and cause them to cancel orders.

Our current and anticipated future dependence upon others for the manufacture of our product candidates may compromise our future profit margins and our ability to commercialize any product candidates that receive marketing approval on a timely and competitive basis.

We may seek collaborations with third parties for the development and commercialization of our product candidates. If those collaborations are not successful, we may not be able to capitalize on the market potential of these product candidates.

We may seek third-party collaborators for the development and commercialization of our product candidates, particularly for the commercialization of any of our product candidates that are approved for

[Table of Contents](#)

marketing outside the United States or for product candidates targeted at larger indications in the United States such as wet AMD and DME. Our likely collaborators for any collaboration arrangements include large and mid-size pharmaceutical companies, regional and national pharmaceutical companies and biotechnology companies. If we do enter into any such arrangements with any third parties, we will likely have limited control over the amount and timing of resources that our collaborators dedicate to the development or commercialization of our product candidates. Our ability to generate revenues from these arrangements will depend on our collaborators' abilities to successfully perform the functions assigned to them in these arrangements.

Collaborations involving our product candidates would pose the following risks to us:

- collaborators have significant discretion in determining the efforts and resources that they will apply to these collaborations;
- collaborators may not perform their obligations as expected;
- collaborators may not pursue development and commercialization of any product candidates that achieve regulatory approval or may elect not to continue or renew development or commercialization programs based on clinical trial results, changes in the collaborators' strategic focus or available funding, or external factors, such as an acquisition, that divert resources or create competing priorities;
- collaborators may delay clinical trials, provide insufficient funding for a clinical trial program, stop a clinical trial or abandon a product candidate, repeat or conduct new clinical trials or require a new formulation of a product candidate for clinical testing;
- collaborators could independently develop, or develop with third parties, products that compete directly or indirectly with our product candidates if the collaborators believe that competitive products are more likely to be successfully developed or can be commercialized on terms that are more economically attractive than ours;
- product candidates discovered in collaboration with us may be viewed by our collaborators as competitive with their own product candidates or products, which may cause collaborators to cease to devote resources to the commercialization of our product candidates;
- a collaborator with marketing and distribution rights to one or more of our product candidates that achieve regulatory approval may not commit sufficient resources to the marketing and distribution of such products;
- disagreements with collaborators, including disagreements over proprietary rights, contract interpretation or the preferred course of development, might cause delays or termination of the research, development or commercialization of product candidates, might lead to additional responsibilities for us with respect to product candidates, or might result in litigation or arbitration, any of which would be time-consuming and expensive;
- collaborators may not properly maintain or defend our or their intellectual property rights or may use our or their proprietary information in such a way as to invite litigation that could jeopardize or invalidate such intellectual property or proprietary information or expose us to potential litigation;
- collaborators may infringe the intellectual property rights of third parties, which may expose us to litigation and potential liability; and

[Table of Contents](#)

- collaborations may be terminated for the convenience of the collaborator and, if terminated, we could be required to raise additional capital to pursue further development or commercialization of the applicable product candidates.

Collaboration agreements may not lead to development or commercialization of product candidates in the most efficient manner or at all. If a present or future collaborator of ours were to be involved in a business combination, the continued pursuit and emphasis on our drug development or commercialization program could be delayed, diminished or terminated.

If we are not able to establish collaborations, we may have to alter some of our future development and commercialization plans.

Our drug development programs and the potential commercialization of our product candidates will require substantial additional cash to fund expenses. For some of our product candidates, we may decide to collaborate with pharmaceutical and biotechnology companies for the future development and potential commercialization of those product candidates.

We face significant competition in seeking appropriate collaborators. Whether we reach a definitive agreement for a collaboration will depend, among other things, upon our assessment of the collaborator's resources and expertise, the terms and conditions of the proposed collaboration and the proposed collaborator's evaluation of a number of factors. Those factors may include the design or results of clinical trials, the likelihood of approval by the FDA or similar regulatory authorities outside the United States, the potential market for the subject product candidate, the costs and complexities of manufacturing and delivering such product candidate to patients, the potential of competing products, the existence of uncertainty with respect to our ownership of technology, which can exist if there is a challenge to such ownership without regard to the merits of the challenge, and industry and market conditions generally. The collaborator may also consider alternative product candidates or technologies for similar indications that may be available to collaborate on and whether such a collaboration could be more attractive than the one with us for our product candidate. We may also be restricted under existing license agreements from entering into agreements on certain terms with potential collaborators. Collaborations are complex and time-consuming to negotiate and document. In addition, there have been a significant number of recent business combinations among large pharmaceutical companies that have resulted in a reduced number of potential future collaborators.

We may not be able to negotiate collaborations on a timely basis, on acceptable terms, or at all. If we are unable to do so, we may have to curtail the development of such product candidate, reduce or delay its development program or one or more of our other development programs, delay its potential commercialization or reduce the scope of any sales or marketing activities, or increase our expenditures and undertake development or commercialization activities at our own expense. If we elect to increase our expenditures to fund development or commercialization activities on our own, we may need to obtain additional capital, which may not be available to us on acceptable terms or at all. If we do not have sufficient funds, we may not be able to further develop our product candidates or bring them to market and generate product revenue.

Risks Related to the Commercialization of Our Product Candidates

If we are unable to establish sales, marketing and distribution capabilities for our product candidates, we may not be successful in commercializing those product candidates, if and when they are approved.

We do not have a sales or marketing infrastructure and have no experience in the sale, marketing or distribution of pharmaceutical drugs or medical devices. To achieve commercial success for any product

candidate for which we may obtain marketing approval in the United States, we will need to establish a sales and marketing organization. In the future, we expect to build a focused sales and marketing infrastructure to target the approximately 1,700 retinal specialists in the United States for any of our product candidates that receive marketing approval. There are risks involved with establishing our own sales, marketing and distribution capabilities. For example, recruiting and training a sales force is expensive and time consuming and could delay any product launch. If the commercial launch of a product candidate for which we recruit a sales force and establish marketing capabilities is delayed or does not occur for any reason, we would have prematurely or unnecessarily incurred these commercialization expenses. This may be costly, and our investment would be lost if we cannot retain or reposition our sales and marketing personnel.

Factors that may inhibit our efforts to commercialize our drugs on our own include:

- our inability to recruit, train and retain adequate numbers of effective sales and marketing personnel;
- the inability of sales personnel to obtain access to physicians or persuade adequate numbers of physicians to prescribe any future drugs;
- the lack of complementary drugs to be offered by sales personnel, which may put us at a competitive disadvantage compared to companies with more extensive product lines; and
- unforeseen costs and expenses associated with creating an independent sales and marketing organization.

If we are unable to establish our own sales, marketing and distribution capabilities and, instead, enter into arrangements with third parties to perform these services, our product revenues and our profitability, if any, are likely to be lower than if we were to sell, market and distribute any product candidates that we develop ourselves. In addition, we may not be successful in entering into arrangements with third parties to sell, market and distribute our product candidates or may be unable to do so on terms that are favorable to us. We likely will have little control over such third parties, and any of them may fail to devote the necessary resources and attention to sell and market our product candidates effectively. If we do not establish sales, marketing and distribution capabilities successfully, either on our own or in collaboration with third parties, we will not be successful in commercializing our product candidates.

Even if any of our product candidates receives marketing approval, it may fail to achieve the degree of market acceptance by physicians, patients, third-party payors and others in the medical community necessary for commercial success.

If any of our product candidates receives marketing approval, it may nonetheless fail to gain sufficient market acceptance by physicians, patients, third-party payors and others in the medical community. SCS administration of drugs is a novel approach and physicians, patients or third-party payors may be hesitant to deviate from or change the current standard of care. If our product candidates do not achieve an adequate level of market acceptance, we may not generate significant product revenue and we may not become profitable. The degree of market acceptance of our product candidates, if approved for commercial sale, will depend on a number of factors, including:

- the efficacy and potential advantages compared to alternative treatments;
- our ability to offer our drugs for sale at competitive prices;
- the convenience and ease of administration compared to alternative treatments;

[Table of Contents](#)

- the willingness of the target patient population to try new therapies and of physicians to prescribe these therapies;
- the willingness of the healthcare community and patients to adopt new technologies and our novel approach of SCS administration of drugs;
- the willingness of retinal specialists to expend the time necessary to receive proper training on administering drugs into the SCS using our microinjector;
- the ability to manufacture our product in sufficient quantities and yields;
- the strength of marketing and distribution support;
- the availability of third-party coverage and adequate reimbursement;
- the prevalence and severity of any side effects; and
- any restrictions on the use of our drugs together with other medications.

We face substantial competition, which may result in others discovering, developing or commercializing products before or more successfully than we do.

The development and commercialization of new products is highly competitive. We face competition with respect to our current product candidates, and will face competition with respect to any product candidates that we may seek to develop or commercialize in the future, from major pharmaceutical companies, specialty pharmaceutical companies and biotechnology companies worldwide.

With respect to CLS-TA, we face competition from other commercially available forms of TA and other injectable and implantable corticosteroids. Bristol-Myers Squibb markets TA under the brand name Kenalog. Kenalog is indicated only for intramuscular or intraarticular injection, although it is commonly used off-label for intraocular inflammation. There are also a number of generic equivalents of Kenalog. In addition, Alcon's injectable TA, Triesence, is approved in the United States for the treatment of uveitis and other inflammatory conditions unresponsive to topical corticosteroids, although it is not indicated for the treatment of macular edema associated with uveitis or RVO or for the treatment of DME. Ozurdex, marketed by Allergan, is a bioerodable extended release implant that delivers the corticosteroid dexamethasone and is approved for the treatment of macular edema following non-infectious uveitis and RVO in both the United States and in the European Union, and recently received approval from the FDA to treat DME in adult patients who have an artificial lens implant or who are scheduled for cataract surgery. Bausch + Lomb markets Retisert, an intravitreal implant of the corticosteroid fluocinolone acetonide, for the treatment of non-infectious uveitis. Alimera Sciences is developing Iluvien, an injectable form of fluocinolone acetonide, as a therapy for DME in the United States. Iluvien has been approved in the European Union for the treatment of DME.

Our product candidates will also compete with anti-VEGF drugs, which are the current standard of care for RVO and wet AMD and have also been approved for DME. Genentech's products Lucentis and Avastin are both anti-VEGF antibodies. Lucentis is currently approved in the United States and European Union for the treatment of wet AMD and the treatment of macular edema following RVO. Lucentis is the only ophthalmic drug therapy approved to treat DME. Avastin is an anti-VEGF drug used by retinal specialists off-label in both the United States and in certain countries of the European Union in the treatment of numerous retinal diseases, but it is not indicated for ophthalmic use. Regeneron's anti-VEGF product, Eylea, is approved for the treatment of wet AMD, macular edema following central retinal vein

occlusion and DME in the United States. In the European Union, Eylea is approved for the treatment of wet AMD, DME and macular edema secondary to central retinal vein occlusion, and is being reviewed for the treatment of macular edema following branch retinal vein occlusion.

Ophthotech's drug candidate, Fovista, an anti-PDGF agent, is in Phase 3 clinical trials as a potential treatment for wet AMD to be administered together with an anti-VEGF drug. We are also aware of several other drug candidates in earlier stages of development as potential treatments for the diseases that we intend to target.

Our commercial opportunity could be reduced or eliminated if our competitors develop and commercialize products that are safer, more effective, have fewer or less severe side effects, are more convenient or are less expensive than any product candidates that we may develop. Our competitors also may obtain FDA or other regulatory approval for their product candidates more rapidly than we may obtain approval for ours, which could result in our competitors establishing a strong market position before we are able to enter the market. In addition, our ability to compete may be affected in many cases by insurers or other third-party payors seeking to encourage the use of generic drugs. For some of the indications that we are pursuing, drugs used off-label, such as Kenalog and Avastin, serve as a cheaper alternatives to our product candidates. Their lower prices could result in significant pricing pressure, even if our product candidates are otherwise viewed as a preferable therapy. Additional drugs may become available on a generic basis over the coming years. If our product candidates achieve marketing approval, we expect that they will be priced at a significant premium over competitive generic drugs.

Many of the companies against which we are competing or against which we may compete in the future have significantly greater financial resources and expertise in research and development, manufacturing, preclinical testing, conducting clinical trials, obtaining regulatory approvals and marketing approved drugs than we do. Mergers and acquisitions in the pharmaceutical and biotechnology industries may result in even more resources being concentrated among a smaller number of our competitors. Smaller and other early stage companies may also prove to be significant competitors, particularly through collaborative arrangements with large and established companies. These third parties compete with us in recruiting and retaining qualified scientific and management personnel, establishing clinical trial sites and patient registration for clinical trials, as well as in acquiring technologies complementary to, or necessary for, our programs.

Even if we are able to commercialize any product candidates, our product candidates may be subject to unfavorable pricing regulations, third-party coverage and reimbursement policies or healthcare reform initiatives.

Our ability to commercialize any product candidates successfully will depend, in part, on the extent to which coverage and adequate reimbursement for our product candidates will be available from government payor programs at the federal and state levels, including Medicare and Medicaid, private health insurers, managed care plans and other third-party payors. Government authorities and other third-party payors, such as private health insurers and health maintenance organizations, decide which medical products they will pay for and establish reimbursement levels. Increasingly, third-party payors are requiring that drug companies provide them with predetermined discounts from list prices and are challenging the prices charged for drugs and products. Coverage and reimbursement may not be available for any product that we commercialize and, even if these are available, the level of reimbursement may not be satisfactory. Inadequate reimbursement levels may adversely affect the demand for, or the price of, any product

candidate for which we obtain marketing approval. Obtaining and maintaining adequate reimbursement for our drugs may be difficult. We may be required to conduct expensive pharmacoeconomic studies to justify coverage and reimbursement or the level of reimbursement compared to other therapies. If coverage and adequate reimbursement are not available or reimbursement is available only to limited levels, we may not be able to successfully commercialize any product candidates for which marketing approval is obtained.

A primary trend in the U.S. healthcare industry and elsewhere is cost containment. Government authorities and other third-party payors have attempted to control costs by limiting coverage and the amount of reimbursement for particular medical products. By way of example, the Medicare Prescription Drug Improvement and Modernization Act of 2003, or MMA, changed the way Medicare covers and pays for pharmaceutical products. The legislation expanded Medicare coverage for outpatient drug purchases by those covered by Medicare under a new Part D and introduced a new reimbursement methodology based on average sales prices for Medicare Part B physician-administered drugs, including drugs currently on the market used by physicians to treat the clinical indications for which we are currently seeking FDA approval and likely our product candidates, if approved. As a result of this legislation and the expansion of federal coverage of drug products, there is additional pressure to contain and reduce costs. While the MMA applies only to drug benefits for Medicare beneficiaries, private payors often follow Medicare coverage policy and payment limitations in setting their own reimbursement rates, and any reduction in reimbursement that results from the MMA may result in a similar reduction in payments from private payors. These cost reduction initiatives and other provisions of the MMA could decrease the coverage and reimbursement that we receive for any approved products, and could seriously harm our business.

Further, from time to time, typically on an annual basis, payment amounts are updated and revised by third-party payors. Because we expect that customers who use our product candidates, if approved, will be separately reimbursed for the procedure administering our products, these updates could directly impact the demand for our products. An example of payment updates is the Medicare program updates to physician payments, which is done on an annual basis using a prescribed statutory formula. In the past, when the application of the formula resulted in lower payment, Congress has passed interim legislation to prevent the reductions. Most recently, the Protecting Access to Medicare Act of 2014, signed into law in April 2014, provided for a 0.5% increase from 2013 payment rates under the Medicare Physician Fee Schedule through 2014 and a 0% increase from January 1 until April 1, 2015. If Congress fails to intervene to prevent the negative update factor in future years, the resulting decrease in payment may adversely affect our revenue and results of operations. In addition, the Medicare physician fee schedule has been adapted by some private payors into their plan-specific physician payment schedule. We cannot predict how pending and future healthcare legislation will impact our business, and any changes in coverage and reimbursement that further restricts coverage of our products or lowers reimbursement for procedures using our products could materially affect our business.

There may be significant delays in obtaining coverage and reimbursement for newly approved drugs, and coverage may be more limited than the indications for which the product is approved by the FDA or similar regulatory authorities outside the United States. Moreover, eligibility for coverage and reimbursement does not imply that a product will be paid for in all cases or at a rate that covers our costs, including research, development, manufacture, sale and distribution expenses. Interim reimbursement levels for new drugs, if applicable, may also not be sufficient to cover our costs and may not be made permanent. Reimbursement rates may vary according to the use of the product and the clinical setting in which it is used, may be based on reimbursement levels already set for lower cost drugs and may be incorporated into existing payments for other services. Net prices for drugs may be reduced by mandatory discounts or

rebates required by government healthcare programs or private payors and by any future relaxation of laws that presently restrict imports of drugs from countries where they may be sold at lower prices than in the United States. Third-party payors often rely upon Medicare coverage policy and payment limitations in setting their own reimbursement policies. However, no uniform policy requirement for coverage and reimbursement for drug products exists among third-party payors in the United States. Therefore, coverage and reimbursement can differ significantly from payor to payor. As a result, the coverage determination process is often a time-consuming and costly process that will require us to provide scientific and clinical support for the use of our products to each payor separately, with no assurance that coverage and adequate reimbursement will be applied consistently or obtained at all. Our inability to promptly obtain coverage and adequate reimbursement rates from both government-funded and private payors for any approved drugs that we develop could significantly harm our operating results, our ability to raise capital needed to commercialize drugs and our overall financial condition.

The regulations that govern marketing approvals, pricing, coverage and reimbursement for new drugs vary widely from country to country. Current and future legislation may significantly change the approval requirements in ways that could involve additional costs and cause delays in obtaining approvals. Some countries require approval of the sale price of a product before it can be marketed. In many countries, the pricing review period begins after marketing or product licensing approval is granted. In some foreign markets, prescription pharmaceutical pricing remains subject to continuing governmental control even after initial approval is granted. As a result, we might obtain marketing approval for a product in a particular country, but then be subject to price regulations that delay commercial launch of the product, possibly for lengthy time periods, and negatively impact the revenues able to be generated from the sale of the product in that country. Adverse pricing limitations may hinder our ability to recoup our investment in one or more product candidates, even if our product candidates obtain marketing approval.

There can be no assurance that our product candidates, if they are approved for sale in the United States or in other countries, will be considered medically reasonable and necessary for a specific indication, that they will be considered cost-effective by third-party payors, that coverage or an adequate level of reimbursement will be available, or that third-party payors' reimbursement policies will not adversely affect our ability to sell our product candidates profitably if they are approved for sale.

Product liability lawsuits against us could cause us to incur substantial liabilities and to limit commercialization of any products that we may develop.

We face an inherent risk of product liability exposure related to the testing of our product candidates in human clinical trials and will face an even greater risk if we commercially sell any drugs that we may develop. If we cannot successfully defend ourselves against claims that our product candidates or products caused injuries, we will incur substantial liabilities. Regardless of merit or eventual outcome, liability claims may result in:

- decreased demand for any product candidates or products that we may develop;
- injury to our reputation and significant negative media attention;
- withdrawal of clinical trial participants;
- significant costs to defend the related litigation;
- substantial monetary awards paid to trial participants or patients;

[Table of Contents](#)

- loss of revenue;
- reduced resources of our management to pursue our business strategy; and
- the inability to commercialize any drugs that we may develop.

We currently hold \$5.0 million in product liability insurance coverage in the aggregate, with a per incident limit of \$5.0 million, which may not be adequate to cover all liabilities that we may incur. We may need to increase our insurance coverage as we expand our clinical trials or if we commence commercialization of our product candidates. Insurance coverage is increasingly expensive. We may not be able to maintain insurance coverage at a reasonable cost or in an amount adequate to satisfy any liability that may arise.

Risks Related to Employee Matters and Managing Our Growth

Our future success depends on our ability to retain key executives and to attract, retain and motivate qualified personnel.

We are highly dependent on the management, research and development, clinical, financial and business development expertise of Daniel White, our Chief Executive Officer, Charlie Deignan, our Chief Financial Officer, and Glenn Noronha, our Executive Vice President of Research and Development, as well as the other members of our scientific and clinical teams. Although we intend to enter into new employment agreements with our executive officers that will be effective upon the closing of this offering, each of them may currently terminate their employment with us at any time and will continue to be able to do so after the closing of this offering. We do not maintain “key person” insurance for any of our executives or employees.

Recruiting and retaining qualified scientific and clinical personnel and, if we progress the development of our product pipeline toward scaling up for commercialization, manufacturing and sales and marketing personnel, will also be critical to our success. The loss of the services of our executive officers or other key employees could impede the achievement of our research, development and commercialization objectives and seriously harm our ability to successfully implement our business strategy. Furthermore, replacing executive officers and key employees may be difficult and may take an extended period of time because of the limited number of individuals in our industry with the breadth of skills and experience required to successfully develop, gain regulatory approval of and commercialize our product candidates. Competition to hire from this limited pool is intense, and we may be unable to hire, train, retain or motivate these key personnel on acceptable terms given the competition among numerous pharmaceutical and biotechnology companies for similar personnel. We also experience competition for the hiring of scientific and clinical personnel from universities and research institutions. In addition, we rely on consultants and advisors, including scientific and clinical advisors, to assist us in formulating our research and development and commercialization strategy. Our consultants and advisors may have commitments under consulting or advisory contracts with other entities that may limit their availability to us. If we are unable to continue to attract and retain high quality personnel, our ability to pursue our growth strategy will be limited.

We expect to expand our development and regulatory capabilities and potentially implement sales, marketing and distribution capabilities, and as a result, we may encounter difficulties in managing our growth, which could disrupt our operations.

As of June 30, 2014, we had 18 employees. As our development progresses, we expect to experience significant growth in the number of our employees and the scope of our operations, particularly in the areas

of research, drug development, regulatory affairs and, if any of our product candidates receives marketing approval, sales, marketing and distribution. To manage our anticipated future growth, we must continue to implement and improve our managerial, operational and financial systems, expand our facilities and continue to recruit and train additional qualified personnel. Due to our limited financial resources and the limited experience of our management team in managing a company with such anticipated growth, we may not be able to effectively manage the expansion of our operations or recruit and train additional qualified personnel. The expansion of our operations may lead to significant costs and may divert our management and business development resources. Any inability to manage growth could delay the execution of our business plans or disrupt our operations.

Risks Related to Our Intellectual Property

If we are unable to obtain and maintain patent protection for our technology and product candidates, or if our licensors are unable to obtain and maintain patent protection for the technology or product candidates that we license from them, or if the scope of the patent protection obtained is not sufficiently broad, our competitors could develop and commercialize technology and drugs similar or identical to ours, and our ability to successfully commercialize our technology and product candidates may be impaired.

Our success depends in large part on our and our licensors' ability to obtain and maintain patent protection in the United States and other countries with respect to our product candidates. We and our licensors seek to protect our proprietary position by filing patent applications in the United States and abroad related to our technology and product candidates.

The patent prosecution process is expensive and time-consuming, and we may not be able to file and prosecute all necessary or desirable patent applications at a reasonable cost or in a timely manner. It is also possible that we will fail to identify patentable aspects of our research and development output before it is too late to obtain patent protection. We may not have the right to control the preparation, filing and prosecution of patent applications, or to maintain the rights to patents licensed to third parties. For example, we do not control the prosecution of the patent applications licensed to us under the Emory/GT License described below. Therefore, these patents and applications may not be prosecuted and enforced in a manner consistent with the best interests of our business. If such licensors fail to maintain such patents, or lose rights to those patents, the rights we have licensed may be reduced or eliminated.

The patent position of biotechnology and pharmaceutical companies generally is highly uncertain, involves complex legal and factual questions and has in recent years been the subject of much litigation. In addition, the laws of foreign countries may not protect our rights to the same extent as the laws of the United States or visa-versa. For example, European patent law restricts the patentability of methods of treatment of the human body more than United States law does. Publications of discoveries in the scientific literature often lag behind the actual discoveries, and patent applications in the United States and other jurisdictions are typically not published until 18 months after filing, or in some cases not at all. Therefore, we cannot know with certainty whether we were the first to make the inventions claimed in our owned or licensed patents or pending patent applications, or that we were the first to file for patent protection of such inventions. As a result, the issuance, scope, validity, enforceability and commercial value of our patent rights are highly uncertain. Our pending and future patent applications may not result in patents being issued that protect our technology or product candidates, in whole or in part, or which effectively prevent

others from commercializing competitive technologies and drugs. Changes in either the patent laws or interpretation of the patent laws in the United States and other countries may diminish the value of our patents or narrow the scope of our patent protection.

Recent patent reform legislation could increase the uncertainties and costs surrounding the prosecution of our patent applications and the enforcement or defense of our issued patents. On September 16, 2011, the Leahy-Smith America Invents Act, or the Leahy-Smith Act, was signed into law. The Leahy-Smith Act includes a number of significant changes to United States patent law. These include provisions that affect the way patent applications are prosecuted, redefine prior art and may also affect patent litigation. The United States Patent and Trademark Office recently developed new regulations and procedures to govern administration of the Leahy-Smith Act, and many of the substantive changes to patent law associated with the Leahy-Smith Act, and in particular, the first to file provisions, only became effective on March 16, 2013. Accordingly, it is not clear what, if any, impact the Leahy-Smith Act will have on the operation of our business. However, the Leahy-Smith Act and its implementation could increase the uncertainties and costs surrounding the prosecution of our patent applications and the enforcement or defense of our issued patents, all of which could significantly harm our business and financial condition. In addition, patent reform legislation may pass in the future that could lead to additional uncertainties and increased costs surrounding the prosecution, enforcement and defense of our owned and licensed patents and patent applications.

Moreover, we may be subject to a third-party preissuance submission of prior art to the U.S. Patent and Trademark Office, or become involved in opposition, derivation, reexamination, *inter partes* review, post-grant review or interference proceedings challenging our patent rights or the patent rights of others. An adverse determination in any such submission, proceeding or litigation could reduce the scope of, or invalidate, our patent rights, allow third parties to commercialize our technology or drugs and compete directly with us, without payment to us, or result in our inability to manufacture or commercialize drugs without infringing third-party patent rights. In addition, if the breadth or strength of protection provided by our patents and patent applications is threatened, it could dissuade companies from collaborating with us to license, develop or commercialize current or future product candidates.

Even if our owned and licensed patent applications issue as patents, they may not issue in a form that will provide us with any meaningful protection, and may not be of sufficient scope or strength to provide us with any commercial advantage. Our competitors may be able to design around our owned or licensed patents by developing similar or alternative technologies or drugs without infringing on our intellectual property rights.

In addition, the issuance of a patent is not conclusive as to its inventorship, scope, validity or enforceability, and our owned and licensed patents may be challenged in the courts or patent offices in the United States and abroad. Such challenges may result in loss of exclusivity or freedom to operate or in patent claims being narrowed, invalidated or held unenforceable, in whole or in part, which could limit our ability to stop others from using or commercializing similar or identical technology and drugs, or limit the duration of the patent protection of our technology and drugs. Given the amount of time required for the development, testing and regulatory review of new product candidates, patents protecting such candidates might expire before or shortly after such candidates are commercialized. As a result, our owned and licensed patent portfolio may not provide us with sufficient rights to exclude others from commercializing drugs similar or identical to ours, and our business will suffer.

We may become involved in lawsuits to protect or enforce our patents or other intellectual property, which could be expensive, time consuming and unsuccessful.

Competitors may infringe our issued patents or other intellectual property. To counter infringement or unauthorized use, we may be required to file infringement claims, which can be expensive and time consuming. Any claims we assert against perceived infringers could provoke these parties to assert counterclaims against us alleging that we infringe their patents. In addition, in a patent infringement proceeding, a court may decide that a patent of ours is invalid or unenforceable, in whole or in part, construe the patent's claims narrowly or refuse to stop the other party from using the technology at issue on the grounds that our patents do not cover the technology. An adverse result in any litigation proceeding could put one or more of our patents at risk of being invalidated or interpreted narrowly, which would undermine our competitive position.

Third parties may initiate legal proceedings alleging that we are infringing their intellectual property rights, the outcome of which would be uncertain and could significantly harm business.

Our commercial success depends upon our ability, and the ability of any collaborators, to develop, manufacture, market and sell our product candidates and use our proprietary technologies without infringing the proprietary rights of third parties. There is considerable intellectual property litigation in the biotechnology and pharmaceutical industries. In particular, we are focused on developing product candidates based on widely used therapeutic agents, many of which are protected by proprietary rights of third parties, and we are developing proprietary formulations of these therapeutic agents specifically for SCS administration using our proprietary microinjector. Although we seek to develop our proprietary drug formulations that don't infringe the intellectual property rights of others, we may become party to, or threatened with, future adversarial proceedings or litigation regarding intellectual property rights with respect to our drugs or other aspects of our technology, including interference or derivation proceedings before the U.S. Patent and Trademark Office. Third parties may assert infringement claims against us based on existing patents or patents that may be granted in the future.

If we are found to infringe a third party's intellectual property rights, we could be required to obtain a license from such third party to continue developing and marketing our technology and drugs. However, we may not be able to obtain any required license on commercially reasonable terms or at all. Even if we were able to obtain a license, it could be non-exclusive, thereby giving our competitors access to the same technologies licensed to us. We could be forced, including by court order, to cease commercializing the infringing technology or product. In addition, we could be found liable for monetary damages, including treble damages and attorneys' fees if we are found to have willfully infringed a patent. A finding of infringement could prevent us from commercializing our product candidates or force us to cease some of our business operations.

Competing products may also be sold in other countries in which our patent coverage might not exist or be as strong. If we lose a foreign patent lawsuit alleging our infringement of a competitor's patent, we could be prevented from marketing our products in one or more foreign countries. As a result, our ability to grow our business and compete in the market may be harmed.

Intellectual property litigation could cause us to spend substantial resources and distract our personnel from their normal responsibilities.

Even if resolved in our favor, litigation or other legal proceedings relating to intellectual property claims may cause us to incur significant expenses, and could distract our technical and management

[Table of Contents](#)

personnel from their normal responsibilities. In addition, there could be public announcements of the results of hearings, motions or other interim proceedings or developments. If securities analysts or investors perceive these results to be negative, it could hurt the price of our common stock. Such litigation or proceedings could substantially increase our operating losses and reduce the resources available for development activities or any future sales, marketing or distribution activities. We may not have sufficient financial or other resources to conduct such litigation or proceedings adequately. Some of our competitors may be able to sustain the costs of such litigation or proceedings more effectively than we can because of their greater financial resources. Uncertainties resulting from the initiation and continuation of patent litigation or other proceedings could compromise our ability to compete in the marketplace.

If we fail to comply with our obligations under our existing intellectual property licenses with third parties, we could lose license rights that are important to our business.

We are a party to a license agreement with Emory University and Georgia Tech Research Corporation, or the Emory/GT License, and may enter into additional license agreements in the future. Our existing license agreement imposes, and we expect that future license agreements would impose, various diligence, milestone payment, royalty, insurance and other obligations on us. For example, under the Emory/GT License, we are required to use commercially reasonable efforts to develop and commercialize licensed products under the agreement and to satisfy other specified obligations, including the payment of license fees and minimum royalty payments. If we fail to comply with our obligations under our license agreements, our licensors may have the right to terminate these license agreements, in which event we might not be able to market any product that is covered by these agreements, or to convert the license to a non-exclusive license, which could impair the value of the product candidate being developed under the license agreement. Termination of these license agreements or reduction or elimination of our licensed rights may result in our having to negotiate new or reinstated licenses with less favorable terms. If our licensors under the Emory/GT License were to terminate their license agreement with us for any reason, we would lose access to critical technology related to our microinjector.

We may need to license additional intellectual property from third parties, and such licenses may not be available or may not be available on commercially reasonable terms.

A third party may hold intellectual property rights, including patent rights, that are important or necessary to the development of our product candidates. It may be necessary for us to use the patented or proprietary technology of third parties to commercialize our product candidates, in which case we would be required to obtain a license from these third parties. Such a license may not be available on commercially reasonable terms, or at all, and we could be forced to accept unfavorable contractual terms. If we are unable to obtain such licenses on commercially reasonable terms, our business could be harmed.

We may be subject to claims by third parties asserting that we or our employees have misappropriated their intellectual property, or claiming ownership of what we regard as our own intellectual property.

Many of our employees were previously employed at universities or other biotechnology or pharmaceutical companies. Although we try to ensure that our employees do not use the proprietary information or know-how of others in their work for us, we may be subject to claims that these employees or we have inadvertently or otherwise used or disclosed intellectual property, including trade secrets or other proprietary information, of any such employee's former employer. We may also in the future be subject to claims that we have caused an employee to breach the terms of his or her non-competition or non-solicitation agreement. Litigation may be necessary to defend against these potential claims.

In addition, while it is our policy to require our employees and contractors who may be involved in the development of intellectual property to execute agreements assigning such intellectual property to us, we may be unsuccessful in executing such an agreement with each party who in fact develops intellectual property that we regard as our own. Our and their assignment agreements may not be self-executing or may be breached, and we may be forced to bring claims against third parties, or defend claims they may bring against us, to determine the ownership of what we regard as our intellectual property.

If we fail in prosecuting or defending any such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights or personnel. A court could prohibit us from using technologies or features that are essential to our products, if such technologies or features are found to incorporate or be derived from the trade secrets or other proprietary information of the former employers. Even if we are successful in prosecuting or defending against such claims, litigation could result in substantial costs and could be a distraction to management. In addition, any litigation or threat thereof may adversely affect our ability to hire employees or contract with independent service providers. Moreover, a loss of key personnel or their work product could hamper or prevent our ability to commercialize our products.

Any trademarks we may obtain may be infringed or successfully challenged, resulting in harm to our business.

We expect to rely on trademarks as one means to distinguish any of our product candidates that are approved for marketing from the products of our competitors. We have not yet selected trademarks for our product candidates and have not yet begun the process of applying to register trademarks for our product candidates. Once we select trademarks and apply to register them, our trademark applications may not be approved. Third parties may oppose our trademark applications, or otherwise challenge our use of the trademarks. In the event that our trademarks are successfully challenged, we could be forced to rebrand our products, which could result in loss of brand recognition and could require us to devote resources to advertising and marketing new brands. Our competitors may infringe our trademarks and we may not have adequate resources to enforce our trademarks.

If we are unable to protect the confidentiality of our trade secrets, our business and competitive position would be harmed.

In addition to seeking patent and trademark protection for our product candidates, we also rely on trade secrets, including unpatented know-how, technology and other proprietary information, to maintain our competitive position. We seek to protect our trade secrets, in part, by entering into non-disclosure and confidentiality agreements with parties who have access to them, such as our employees, corporate collaborators, outside scientific collaborators, contract manufacturers, consultants, advisors and other third parties. We also enter into confidentiality and invention or patent assignment agreements with our employees and consultants. Despite these efforts, any of these parties may breach the agreements and disclose our proprietary information, including our trade secrets. Monitoring unauthorized uses and disclosures of our intellectual property is difficult, and we do not know whether the steps we have taken to protect our intellectual property will be effective. In addition, we may not be able to obtain adequate remedies for any such breaches. Enforcing a claim that a party illegally disclosed or misappropriated a trade secret is difficult, expensive and time-consuming, and the outcome is unpredictable. In addition, some courts inside and outside the United States are less willing or unwilling to protect trade secrets.

Moreover, our competitors may independently develop knowledge, methods and know-how equivalent to our trade secrets. Competitors could purchase our products and replicate some or all of the competitive

advantages we derive from our development efforts for technologies on which we do not have patent protection. If any of our trade secrets were to be lawfully obtained or independently developed by a competitor, we would have no right to prevent them, or those to whom they communicate it, from using that technology or information to compete with us. If any of our trade secrets were to be disclosed to or independently developed by a competitor, our competitive position would be harmed.

Risks Related to Regulatory Approval of Our Product Candidates and Other Legal Compliance Matters

If we are not able to obtain, or if there are delays in obtaining, required regulatory approvals, we will not be able to commercialize our product candidates, and our ability to generate revenue will be materially impaired.

Our product candidates and the activities associated with their development and commercialization, including their design, research, testing, manufacture, safety, efficacy, quality control, recordkeeping, labeling, packaging, storage, approval, advertising, promotion, sale, distribution, import, export, and reporting of safety and other post-market information, are subject to comprehensive regulation by the FDA and other regulatory agencies in the United States and by the EMA and similar regulatory authorities outside the United States. Failure to obtain marketing approval for a product candidate will prevent us from commercializing the product candidate. We have not received approval to market any of our product candidates from regulatory authorities in any jurisdiction. We have only limited experience in filing and supporting the applications necessary to gain marketing approvals and expect to rely on third-party CROs to assist us in this process. Securing marketing approval requires the submission of extensive preclinical and clinical data and supporting information to regulatory authorities for each therapeutic indication to establish the product candidate's safety and efficacy. Securing marketing approval also requires the submission of information about the product manufacturing process to, and inspection of manufacturing facilities by, the regulatory authorities. Our product candidates may not be effective, may be only moderately effective or may prove to have undesirable or unintended side effects, toxicities or other characteristics that may preclude our obtaining marketing approval or prevent or limit commercial use. If any of our product candidates receives marketing approval, the accompanying label may limit its approved use, which could limit sales of the product.

The process of obtaining marketing approvals, both in the United States and abroad, is expensive and may take many years, if approval is obtained at all, and can vary substantially based upon a variety of factors, including the type, complexity and novelty of the product candidates involved. Changes in marketing approval policies during the development period, changes in or the enactment of additional statutes or regulations, or changes in regulatory review for each submitted product application, may cause delays in the approval or rejection of an application. Regulatory authorities have substantial discretion in the approval process and may refuse to accept any application or may decide that our data is insufficient for approval and require additional preclinical, clinical or other studies. In addition, varying interpretations of the data obtained from preclinical and clinical testing could delay, limit or prevent marketing approval of a product candidate. Any marketing approval we ultimately obtain may be limited or subject to restrictions or post-approval commitments that render the approved product not commercially viable.

If we experience delays in obtaining approval or if we fail to obtain approval of our product candidates, the commercial prospects for our product candidates may be harmed and our ability to generate revenues will be materially impaired.

If the FDA does not conclude that our product candidates satisfy the requirements for the Section 505(b)(2) regulatory approval pathway, or if the requirements under Section 505(b)(2) are not as we expect, the approval pathway for our product candidates will likely take significantly longer, cost significantly more and entail significantly greater complications and risks than anticipated, and in either case may not be successful.

We intend to seek FDA approval through the Section 505(b)(2) regulatory pathway for our product candidates, such as CLS-1001. Our product candidates, including our proprietary drug formulations packaged together with our microinjector, are drug/device combination products that will be regulated under the drug provisions of the FDCA, enabling us to submit NDAs for approval of our product candidates. The Drug Price Competition and Patent Term Restoration Act of 1984, also known as the Hatch-Waxman Amendments, added Section 505(b)(2) to the FDCA. Section 505(b)(2) permits the filing of an NDA where at least some of the information required for approval comes from studies not conducted by or for the applicant and for which the applicant has not obtained a right of reference.

If the FDA does not allow us to pursue the Section 505(b)(2) regulatory pathway as anticipated, we may need to conduct additional clinical trials, provide additional data and information and meet additional standards for regulatory approval. If this were to occur, the time and financial resources required to obtain FDA approval for our product candidates, and complications and risks associated with our product candidates, would likely substantially increase. We may need to obtain additional funding, which could result in significant dilution to the ownership interests of our then existing stockholders to the extent we issue equity securities or convertible debt. We cannot assure you that we would be able to obtain such additional financing on terms acceptable to us, if at all. Moreover, inability to pursue the Section 505(b)(2) regulatory pathway could result in competitive products reaching the market before our product candidates, which could impact our competitive position and prospects. Even if we are allowed to pursue the Section 505(b)(2) regulatory pathway, we cannot assure you that our product candidates will receive the requisite approvals for commercialization.

In addition, notwithstanding the approval of a number of products by the FDA under Section 505(b)(2) over the last few years, some pharmaceutical companies and others have objected to the FDA's interpretation of Section 505(b)(2). If the FDA's interpretation of Section 505(b)(2) is successfully challenged, the FDA may be required to change its Section 505(b)(2) policies and practices, which could delay or even prevent the FDA from approving any NDA that we submit under Section 505(b)(2).

Additional time may be required to obtain regulatory approval for our product candidates because they are combination products.

Our product candidates are drug/device combination products that require coordination within the FDA and similar foreign regulatory agencies for review of their device and drug components. Although the FDA and similar foreign regulatory agencies have systems in place for the review and approval of combination products such as ours, we may experience delays in the development and commercialization of our product candidates due to regulatory timing constraints and uncertainties in the product development and approval process.

Failure to obtain marketing approval in international jurisdictions would prevent our product candidates from being marketed abroad.

In order to market and sell our products in the European Union and any other jurisdictions, we must obtain separate marketing approvals and comply with numerous and varying regulatory requirements. The

[Table of Contents](#)

approval procedure varies among countries and can involve additional testing. The time required to obtain approval may differ substantially from that required to obtain FDA approval. The regulatory approval process outside the United States generally includes all of the risks associated with obtaining FDA approval. In addition, in many countries outside the United States, it is required that the product be approved for reimbursement before the product can be approved for sale in that country. We may not obtain approvals from regulatory authorities outside the United States on a timely basis, if at all. Approval by the FDA does not ensure approval by regulatory authorities in other countries or jurisdictions, and approval by one regulatory authority outside the United States does not ensure approval by regulatory authorities in other countries or jurisdictions or by the FDA. However, failure to obtain approval in one jurisdiction may impact our ability to obtain approval elsewhere. We may not be able to file for marketing approvals and may not receive necessary approvals to commercialize our products in any market.

A variety of risks associated with marketing our product candidates internationally could affect our business.

We may seek regulatory approval for our product candidates outside of the United States and, accordingly, we expect that we will be subject to additional risks related to operating in foreign countries if we obtain the necessary approvals, including:

- differing regulatory requirements in foreign countries;
- the potential for so-called parallel importing, which is what happens when a local seller, faced with high or higher local prices, opts to import goods from a foreign market with low or lower prices rather than buying them locally;
- unexpected changes in tariffs, trade barriers, price and exchange controls and other regulatory requirements;
- economic weakness, including inflation, or political instability in particular foreign economies and markets;
- compliance with tax, employment, immigration and labor laws for employees living or traveling abroad;
- foreign taxes, including withholding of payroll taxes;
- foreign currency fluctuations, which could result in increased operating expenses and reduced revenues, and other obligations incident to doing business in another country;
- difficulties staffing and managing foreign operations;
- workforce uncertainty in countries where labor unrest is more common than in the United States;
- potential liability under the Foreign Corrupt Practices Act of 1977 or comparable foreign regulations;
- challenges enforcing our contractual and intellectual property rights, especially in those foreign countries that do not respect and protect intellectual property rights to the same extent as the United States;
- production shortages resulting from any events affecting raw material supply or manufacturing capabilities abroad; and
- business interruptions resulting from geo-political actions, including war and terrorism.

These and other risks associated with our international operations may compromise our ability to achieve or maintain profitability.

Any product candidate for which we obtain marketing approval could be subject to post-marketing restrictions or recall or withdrawal from the market, and we may be subject to penalties if we fail to comply with regulatory requirements or if we experience unanticipated problems with our product candidates, when and if any of them are approved.

Any product candidate for which we obtain marketing approval, along with the manufacturing processes, post-approval clinical data, labeling, advertising and promotional activities for such product, will be subject to continual requirements of and review by the FDA and other regulatory authorities. These requirements include submissions of safety and other post-marketing information and reports, registration and listing requirements, cGMP requirements relating to manufacturing, quality control, quality assurance and corresponding maintenance of records and documents, requirements regarding the distribution of samples to physicians and recordkeeping. Even if marketing approval of a product candidate is granted, the approval may be subject to limitations on the indicated uses for which the product may be marketed or to the conditions of approval, including the requirement to implement a risk evaluation and mitigation strategy, which could involve requirements for, among other things, a medication guide, special training for prescribers and dispensers, and patient registries. If any of our product candidates receives marketing approval, the accompanying label may limit its approved uses, which could limit sales of the product.

The FDA may also impose requirements for costly post-marketing studies or clinical trials and surveillance to monitor the safety or efficacy of the product. The FDA closely regulates the post-approval marketing and promotion of products to ensure products are marketed only for the approved indications and in accordance with the provisions of the approved labeling. The FDA imposes stringent restrictions on manufacturers' communications regarding off-label use and if we do not market our drugs for their approved indications, we may be subject to enforcement action for off-label marketing. Violations of the FDCA relating to the promotion of prescription drugs may lead to investigations alleging violations of federal and state healthcare fraud and abuse laws, as well as state consumer protection laws.

In addition, later discovery of previously unknown adverse events or other problems with our drugs, including device malfunctions, manufacturers or manufacturing processes, or failure to comply with regulatory requirements, may have negative consequences, including:

- restrictions on such drugs, manufacturers or manufacturing processes;
- restrictions on the labeling or marketing of a drug;
- restrictions on drug distribution or use;
- requirements to conduct post-marketing studies or clinical trials;
- warning letters;
- recall or withdrawal of the drugs from the market;
- refusal to approve pending applications or supplements to approved applications that we submit;
- clinical holds;
- fines, restitution or disgorgement of profits or revenues;
- suspension or withdrawal of marketing approvals;
- refusal to permit the import or export of our drugs;

[Table of Contents](#)

- product seizure; or
- injunctions or the imposition of civil or criminal penalties.

Non-compliance with European Union requirements regarding safety monitoring, or pharmacovigilance, and with requirements related to the development of drugs for the pediatric population, can also result in significant financial penalties. Similarly, failure to comply with the European Union's requirements regarding the protection of personal information can also lead to significant penalties and sanctions.

Even if we receive orphan drug designation for our most advanced product candidate, CLS-1001, we may not be able to obtain orphan drug marketing exclusivity for this product candidate or any of our future product candidates.

Regulatory authorities in some jurisdictions, including the United States and the European Union, may designate drugs for relatively small patient populations as orphan drugs. Under the Orphan Drug Act, the FDA may designate a drug as an orphan drug if it is intended to treat a rare disease or condition, which is generally defined as a patient population of fewer than 200,000 individuals annually in the United States. We have applied for orphan drug designation from the FDA, and intend to apply for orphan drug designation from the European Medicines Agency, or EMA, for CLS-1001 for the treatment of non-infectious uveitis, and we may seek orphan drug designation for our future drug candidates. However, we have not yet obtained orphan drug designation from either the FDA or the EMA, and there can be no assurance that we will receive this designation from either regulatory authority.

Generally, if a drug with an orphan drug designation subsequently receives the first marketing approval for the indication for which it has such designation, the drug is entitled to a period of marketing exclusivity, which precludes the FDA or the EMA from approving another marketing application for the same drug for the same indication for that time period. The applicable period is seven years in the United States and 10 years in the European Union. The European Union exclusivity period can be reduced to six years if a drug no longer meets the criteria for orphan drug designation or if the drug is sufficiently profitable so that market exclusivity is no longer justified. Orphan drug exclusivity may be lost if the FDA or EMA determines that the request for designation was materially defective or if the manufacturer is unable to assure sufficient quantity of the drug to meet the needs of patients with the rare disease or condition. Even if we obtain orphan drug exclusivity for CLS-1001 for the treatment of non-infectious uveitis, that exclusivity may not effectively protect the drug candidate from competition because different drugs can be approved for the same condition. Even after an orphan drug is approved, the FDA can subsequently approve another drug for the same condition if the FDA concludes that the later drug is clinically superior in that it is shown to be safer, more effective or makes a major contribution to patient care.

Our employees, independent contractors, principal investigators, CROs, consultants, commercial partners and vendors may engage in misconduct or other improper activities, including non-compliance with regulatory standards and requirements.

We are exposed to the risk of employee fraud or other misconduct. Misconduct by employees and independent contractors, such as principal investigators, CROs, consultants, commercial partners and vendors, could include failures to comply with FDA regulations, to provide accurate information to the FDA, to comply with manufacturing standards we have established, to comply with federal and state healthcare fraud and abuse laws, to report financial information or data accurately or to disclose unauthorized activities to us. In particular, sales, marketing and other business arrangements in the

healthcare industry are subject to extensive laws intended to prevent fraud, kickbacks, self-dealing and other abusive practices. These laws may restrict or prohibit a wide range of business activities, including, but not limited to, research, manufacturing, distribution, pricing, discounting, marketing and promotion, sales commission, customer incentive programs and other business arrangements. Employee and independent contractor misconduct could also involve the improper use of individually identifiable information, including, without limitation, information obtained in the course of clinical trials, which could result in regulatory sanctions and serious harm to our reputation. In addition, federal procurement laws impose substantial penalties for misconduct in connection with government contracts and require certain contractors to maintain a code of business ethics and conduct. It is not always possible to identify and deter employee and independent contractor misconduct, and any precautions we take to detect and prevent improper activities may not be effective in controlling unknown or unmanaged risks or losses or in protecting us from governmental investigations or other actions or lawsuits stemming from a failure to be in compliance with such laws. If any such actions are instituted against us, those actions could have a significant impact on our business, including the imposition of civil, criminal and administrative penalties, damages, monetary fines, disgorgement, possible exclusion from participation in Medicare, Medicaid and other federal healthcare programs, contractual damages, reputational harm, diminished profits and future earnings and curtailment or restructuring of our operations, any of which could adversely affect our ability to operate.

Our current and future relationships with healthcare professionals, principal investigators, consultants, customers and third-party payors in the United States and elsewhere may be subject, directly or indirectly, to applicable anti-kickback, fraud and abuse, false claims, physician payment transparency, health information privacy and security and other healthcare laws and regulations, which could expose us to penalties.

Healthcare providers, physicians and third-party payors in the United States and elsewhere will play a primary role in the recommendation and prescription of any product candidates for which we obtain marketing approval. Our current and future arrangements with healthcare professionals, principal investigators, consultants, customers and third-party payors may expose us to broadly applicable fraud and abuse and other healthcare laws, including, without limitation, the federal Anti-Kickback Statute and the federal False Claims Act, that may constrain the business or financial arrangements and relationships through which we sell, market and distribute any product candidates for which we obtain marketing approval. In addition, we may be subject to physician payment transparency laws and patient privacy and security regulation by the federal government and by the states and foreign jurisdictions in which we conduct our business. The applicable federal, state and foreign healthcare laws that may affect our ability to operate include the following:

- the federal Anti-Kickback Statute, which prohibits, among other things, persons from knowingly and willfully soliciting, offering, receiving or providing remuneration, directly or indirectly, in cash or in kind, to induce or reward, or in return for, either the referral of an individual for, or the purchase, lease, order or recommendation of, any good, facility, item or service, for which payment may be made, in whole or in part, under federal and state healthcare programs such as Medicare and Medicaid;
- federal civil and criminal false claims laws and civil monetary penalty laws, including the federal False Claims Act, which impose criminal and civil penalties, including civil whistleblower or *qui tam* actions, against individuals or entities for, among other things, knowingly presenting, or causing to be presented, to the federal government, including the Medicare and Medicaid programs, claims

[Table of Contents](#)

for payment that are false or fraudulent or making a false statement to avoid, decrease or conceal an obligation to pay money to the federal government;

- the civil monetary penalties statute, which imposes penalties against any person or entity who, among other things, is determined to have presented or caused to be presented a claim to a federal health program that the person knows or should know is for an item or service that was not provided as claimed or is false or fraudulent;
- the federal Health Insurance Portability and Accountability Act of 1996, or HIPAA, which created new federal criminal statutes that prohibit knowingly and willfully executing, or attempting to execute, a scheme to defraud any healthcare benefit program or obtain, by means of false or fraudulent pretenses, representations or promises, any of the money or property owned by, or under the custody or control of, any healthcare benefit program, regardless of whether the payor is public or private, knowingly and willfully embezzling or stealing from a health care benefit program, willfully obstructing a criminal investigation of a health care offense and knowingly and willfully falsifying, concealing or covering up by any trick or device a material fact or making any materially false statements in connection with the delivery of, or payment for, healthcare benefits, items or services relating to healthcare matters;
- HIPAA, as amended by the Health Information Technology for Economic and Clinical Health Act of 2009, or HITECH, and their respective implementing regulations, which impose obligations on “covered entities,” including healthcare providers, health plans, and healthcare clearinghouses, as well as their respective “business associates” that create, receive, maintain or transmit individually identifiable health information for or on behalf of a covered entity, with respect to safeguarding the privacy, security and transmission of individually identifiable health information;
- the federal Open Payments program, created under Section 6002 of Patient Protection and Affordable Care Act, as amended by the Health Care and Education Reconciliation Act, or collectively, the ACA, and its implementing regulations, imposed new annual reporting requirements for manufacturers of drugs, devices, biologicals and medical supplies for certain payments and “transfers of value” provided to physicians and teaching hospitals, as well as ownership and investment interests held by physicians and their immediate family members. Failure to submit timely, accurately and completely the required information for all payments, transfers of value and ownership or investment interests may result in civil monetary penalties of up to an aggregate of \$150,000 per year and up to an aggregate of \$1.0 million per year for “knowing failures,” for an aggregate potential annual liability of \$1,150,000; and
- analogous state and foreign laws, such as state anti-kickback and false claims laws, which may apply to sales or marketing arrangements and claims involving healthcare items or services reimbursed by non-governmental third-party payors, including private insurers; state and foreign laws that require pharmaceutical companies to comply with the pharmaceutical industry’s voluntary compliance guidelines and the relevant compliance guidance promulgated by the federal government or to adopt compliance programs as prescribed by state laws and regulations, or that otherwise restrict payments that may be made to healthcare providers; state and foreign laws that require drug manufacturers to report information related to payments and other transfers of value to physicians and other healthcare providers or marketing expenditures; and state and foreign laws governing the privacy and security of health information in certain circumstances, many of which differ from each other in significant ways and often are not preempted by HIPAA, thus complicating compliance efforts.

Further, the ACA, among other things, amended the intent requirement of the federal Anti-Kickback Statute and certain criminal statutes governing healthcare fraud. A person or entity no longer needs to have actual knowledge of the statute or specific intent to violate it. In addition, the ACA provided that the government may assert that a claim including items or services resulting from a violation of the federal Anti-Kickback Statute constitutes a false or fraudulent claim for purposes of the False Claims Act.

Efforts to ensure that our future business arrangements with third parties will comply with applicable healthcare laws and regulations may involve substantial costs. It is possible that governmental authorities will conclude that our business practices may not comply with current or future statutes, regulations or case law involving applicable fraud and abuse or other healthcare laws. If our operations are found to be in violation of any of these laws or any other governmental regulations that may apply to us, we may be subject to significant civil, criminal and administrative penalties, including, without limitation, damages, fines, imprisonment, exclusion from participation in government healthcare programs, such as Medicare and Medicaid, and the curtailment or restructuring of our operations, which could significantly harm our business. If any of the physicians or other healthcare providers or entities with whom we expect to do business, including future collaborators, are found not to be in compliance with applicable laws, they may be subject to criminal, civil or administrative sanctions, including exclusions from participation in government healthcare programs, which could also affect our business.

Recently enacted and future legislation may increase the difficulty and cost for us to obtain marketing approval of and commercialize our product candidates and affect the prices we may obtain.

In the United States and some foreign jurisdictions, there have been a number of legislative and regulatory changes and proposed changes regarding the healthcare system that could prevent or delay marketing approval of our product candidates, restrict or regulate post-approval activities and affect our ability to profitably sell any product candidates for which we obtain marketing approval.

Among policy makers and payors in the United States and elsewhere, there is significant interest in promoting changes in healthcare systems with the stated goals of containing healthcare costs, improving quality and/or expanding access. In the United States, the pharmaceutical industry has been a particular focus of these efforts and has been significantly affected by major legislative initiatives. In March 2010, President Obama signed into law the ACA, a sweeping law intended to broaden access to health insurance, reduce or constrain the growth of healthcare spending, enhance remedies against fraud and abuse, add new transparency requirements for the healthcare and health insurance industries, impose new taxes and fees on the health industry and impose additional health policy reforms.

Among the provisions of the ACA of importance to our potential product candidates are the following:

- an annual, nondeductible fee on any entity that manufactures or imports certain branded prescription drugs and biologic agents, apportioned among these entities according to their market share in certain government healthcare programs;
- a 2.3% medical device excise tax on certain transactions, including many U.S. sales of medical devices, which currently includes, and we expect will continue to include, U.S. sales of drug and device combination products;
- an increase in the statutory minimum rebates a manufacturer must pay under the Medicaid Drug Rebate Program to 23.1% and 13.0% of the average manufacturer price for branded and generic drugs, respectively;

[Table of Contents](#)

- expansion of healthcare fraud and abuse laws, including the False Claims Act and the Anti-Kickback Statute, new government investigative powers and enhanced penalties for non-compliance;
- addressed a new methodology by which rebates owed by manufacturers under the Medicaid Drug Rebate Program are calculated for drugs that are inhaled, infused, instilled, implanted or injected;
- a new Medicare Part D coverage gap discount program, in which manufacturers must agree to offer 50% point-of-sale discounts off negotiated prices of applicable brand drugs to eligible beneficiaries during their coverage gap period, as a condition for the manufacturer's outpatient drugs to be covered under Medicare Part D;
- extension of manufacturers' Medicaid rebate liability to covered drugs dispensed to individuals who are enrolled in Medicaid managed care organizations;
- expansion of eligibility criteria for Medicaid programs by, among other things, allowing states to offer Medicaid coverage to additional individuals and by adding new mandatory eligibility categories for certain individuals with income at or below 133% of the Federal Poverty Level beginning in 2014, thereby potentially increasing manufacturers' Medicaid rebate liability;
- expansion of the entities eligible for discounts under the Public Health Service pharmaceutical pricing program;
- the new requirements under the federal Open Payments program and its implementing regulations;
- a new requirement to annually report drug samples that manufacturers and distributors provide to physicians; and
- a new Patient-Centered Outcomes Research Institute to oversee, identify priorities in, and conduct comparative clinical effectiveness research, along with funding for such research.

In addition, other legislative changes have been proposed and adopted since the ACA was enacted. These changes included aggregate reductions to Medicare payments to providers of up to 2% per fiscal year, which went into effect in April 2013 and will stay in effect through 2024 unless additional Congressional action is taken. In January 2013, President Obama signed into law the American Taxpayer Relief Act of 2012, which, among other things, further reduced Medicare payments to several providers, and increased the statute of limitations period for the government to recover overpayments to providers from three to five years. These new laws may result in additional reductions in Medicare and other healthcare funding, which could negatively impact customers for our product candidates, if approved, and, accordingly, our financial operations.

We expect that the ACA, as well as other healthcare reform measures that may be adopted in the future, may result in more rigorous coverage criteria and in additional downward pressure on the price that we receive for any approved product. Any reduction in reimbursement from Medicare or other government programs may result in a similar reduction in payments from private payors. The implementation of cost containment measures or other healthcare reforms may prevent us from being able to generate revenue, attain profitability, or commercialize our drugs.

Legislative and regulatory proposals have been made to expand post-approval requirements and restrict sales and promotional activities for drugs. We cannot be sure whether additional legislative changes will be enacted, or whether the FDA regulations, guidance or interpretations will be changed, or what the impact of such changes on the marketing approvals of our product candidates, if any, may be. In addition, increased

[Table of Contents](#)

scrutiny by the U.S. Congress of the FDA's approval process may significantly delay or prevent marketing approval, as well as subject us to more stringent product labeling and post-marketing testing and other requirements.

Governments outside the United States tend to impose strict price controls, which may affect our revenue, if any.

In some countries, particularly the countries of the European Union, the pricing of prescription pharmaceuticals is subject to governmental control. In these countries, pricing negotiations with governmental authorities can take considerable time after the receipt of marketing approval for a product. To obtain coverage and reimbursement or pricing approval in some countries, we may be required to conduct a clinical trial that compares the cost-effectiveness of our product candidate to other available therapies. If reimbursement of our drugs is unavailable or limited in scope or amount, or if pricing is set at unsatisfactory levels, our business could be harmed, possibly materially.

If we fail to comply with environmental, health and safety laws and regulations, we could become subject to fines or penalties or incur costs that could harm our business.

We are subject to numerous environmental, health and safety laws and regulations, including those governing laboratory procedures and the handling, use, storage, treatment and disposal of hazardous materials and wastes. Our operations involve the use of hazardous materials, including chemicals and biological materials. Our operations also produce hazardous waste products. We generally contract with third parties for the disposal of these materials and wastes. We cannot eliminate the risk of contamination or injury from these materials. In the event of contamination or injury resulting from our use of hazardous materials, we could be held liable for any resulting damages, and any liability could exceed our resources. We also could incur significant costs associated with civil or criminal fines and penalties for failure to comply with such laws and regulations.

Although we maintain workers' compensation insurance to cover us for costs and expenses we may incur due to injuries to our employees resulting from the use of hazardous materials, this insurance may not provide adequate coverage against potential liabilities. We do not maintain insurance for environmental liability or toxic tort claims that may be asserted against us in connection with our storage or disposal of biological or hazardous materials.

In addition, we may incur substantial costs in order to comply with current or future environmental, health and safety laws and regulations. These current or future laws and regulations may impair our research, development or production efforts. Our failure to comply with these laws and regulations also may result in substantial fines, penalties or other sanctions.

Risks Related to this Offering, Ownership of Our Common Stock and Our Status as a Public Company

An active trading market for our common stock may not develop and you may not be able to resell your shares of our common stock at or above the initial offering price, if at all.

Prior to this offering, there has been no public market for our common stock. The initial public offering price for our common stock will be determined through negotiations with the underwriters and may not be indicative of the price at which our common stock will trade upon completion of this offering. Although we have applied to list our common stock on The NASDAQ Global Market, an active trading market for our

[Table of Contents](#)

shares may never develop or be sustained following this offering. If an active market for our common stock does not develop or is not sustained, it may be difficult for you to sell shares you purchased in this offering at an attractive price or at all.

The trading price of the shares of our common stock may be volatile, and purchasers of our common stock could incur substantial losses.

Our stock price may be volatile. The stock market in general and the market for biopharmaceutical companies in particular have experienced extreme volatility that has often been unrelated to the operating performance of particular companies. As a result of this volatility, investors may not be able to sell their common stock at or above the price paid for the shares. The market price for our common stock may be influenced by many factors, including:

- actual or anticipated variations in our operating results;
- changes in financial estimates by us or by any securities analysts who might cover our stock;
- conditions or trends in our industry;
- stock market price and volume fluctuations of comparable companies and, in particular, those that operate in the biopharmaceutical industry;
- announcements by us or our competitors of significant acquisitions, strategic partnerships or divestitures;
- announcements of investigations or regulatory scrutiny of our operations or lawsuits filed against us;
- capital commitments;
- investors' general perception of us and our business;
- recruitment or departure of key personnel; and
- sales of our common stock, including sales by our directors and officers or specific stockholders.

In addition, in the past, stockholders have initiated class action lawsuits against pharmaceutical and biotechnology companies following periods of volatility in the market prices of these companies' stock. Such litigation, if instituted against us, could cause us to incur substantial costs and divert management's attention and resources from our business.

If equity research analysts do not publish research or reports, or publish unfavorable research or reports, about us, our business or our market, our stock price and trading volume could decline.

The trading market for our common stock will be influenced by the research and reports that equity research analysts publish about us and our business. We do not currently have and may never obtain research coverage by equity research analysts. Equity research analysts may elect not to provide research coverage of our common stock after the completion of this offering, and such lack of research coverage may adversely affect the market price of our common stock. In the event we do have equity research analyst coverage, we will not have any control over the analysts or the content and opinions included in their reports. The price of our stock could decline if one or more equity research analysts downgrade our stock or issue other unfavorable commentary or research. If one or more equity research analysts ceases coverage of us or fails to publish reports on us regularly, demand for our stock could decrease, which in turn could cause our stock price or trading volume to decline.

If you purchase shares of our common stock in this offering, you will suffer immediate dilution of your investment.

We expect the initial public offering price of our common stock to be substantially higher than the net tangible book value per share of our common stock. Therefore, if you purchase shares of our common stock in this offering, you will pay a price per share that substantially exceeds our pro forma as adjusted net tangible book value per share after this offering. Based on an assumed initial public offering price of \$ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, you will experience immediate dilution of \$ per share, representing the difference between our pro forma as adjusted net tangible book value per share after this offering and the assumed initial public offering price. After this offering, we will also have outstanding options and warrants to purchase common stock with exercise prices lower than the initial public offering price. To the extent these outstanding options and warrants are exercised, there will be further dilution to investors in this offering.

A significant portion of our total outstanding shares are restricted from immediate resale but may be sold into the market in the near future. This could cause the market price of our common stock to drop significantly, even if our business is doing well.

Sales of a substantial number of shares of our common stock in the public market could occur at any time. If our stockholders sell, or the market perceives that our stockholders intend to sell, substantial amounts of our common stock in the public market following this offering, the market price of our common stock could decline significantly.

Upon completion of this offering, we will have outstanding shares of common stock, assuming no exercise of outstanding options or warrants. Of these shares, the shares sold in this offering and additional shares will be freely tradable, additional shares of common stock will be eligible for sale in the public market beginning 90 days after the date of this prospectus under Rule 144 and Rule 701, and additional shares of common stock will be available for sale in the public market beginning 180 days after the date of this prospectus following the expiration of lock-up agreements between some of our stockholders and the underwriters. The representatives of the underwriters may release these stockholders from their lock-up agreements with the underwriters at any time and without notice, which would allow for earlier sales of shares in the public market.

In addition, promptly following the completion of this offering, we intend to file one or more registration statements on Form S-8 registering the issuance of approximately shares of common stock subject to options or other equity awards issued or reserved for future issuance under our equity incentive plans. Shares registered under these registration statements on Form S-8 will be available for sale in the public market subject to vesting arrangements and exercise of options, the lock-up agreements described above and, in the case of our affiliates, the restrictions of Rule 144.

Additionally, after this offering, the holders of an aggregate of approximately shares of our common stock and shares of our common stock issuable upon the exercise of outstanding warrants, or their transferees, will have rights, subject to some conditions, to require us to file one or more registration statements covering their shares or to include their shares in registration statements that we may file for ourselves or other stockholders. If we were to register the resale of these shares, they could be freely sold in the public market. If these additional shares are sold, or if it is perceived that they will be sold, in the public market, the trading price of our common stock could decline.

Provisions in our corporate charter documents and under Delaware law may prevent or frustrate attempts by our stockholders to change our management and hinder efforts to acquire a controlling interest in us, and the market price of our common stock may be lower as a result.

There are provisions in our certificate of incorporation and bylaws as they will be in effect following this offering that may make it difficult for a third party to acquire, or attempt to acquire, control of our company, even if a change in control was considered favorable by you and other stockholders. For example, our board of directors will have the authority to issue up to 5,000,000 shares of preferred stock. The board of directors can fix the price, rights, preferences, privileges, and restrictions of the preferred stock without any further vote or action by our stockholders. The issuance of shares of preferred stock may delay or prevent a change in control transaction. As a result, the market price of our common stock and the voting and other rights of our stockholders may be adversely affected. An issuance of shares of preferred stock may result in the loss of voting control to other stockholders.

Our charter documents will also contain other provisions that could have an anti-takeover effect, including:

- only one of our three classes of directors will be elected each year;
- stockholders will not be entitled to remove directors other than by a 66 2/3% vote and only for cause;
- stockholders will not be permitted to take actions by written consent;
- stockholders cannot call a special meeting of stockholders; and
- stockholders must give advance notice to nominate directors or submit proposals for consideration at stockholder meetings.

In addition, we are subject to the anti-takeover provisions of Section 203 of the Delaware General Corporation Law, which regulates corporate acquisitions by prohibiting Delaware corporations from engaging in specified business combinations with particular stockholders of those companies. These provisions could discourage potential acquisition proposals and could delay or prevent a change in control transaction. They could also have the effect of discouraging others from making tender offers for our common stock, including transactions that may be in your best interests. These provisions may also prevent changes in our management or limit the price that investors are willing to pay for our stock.

Concentration of ownership of our common stock among our existing executive officers, directors and principal stockholders may prevent new investors from influencing significant corporate decisions.

Upon completion of this offering, our executive officers, directors and current beneficial owners of 5% or more of our common stock and their respective affiliates will, in the aggregate, beneficially own approximately % of our outstanding common stock. As a result, these persons, acting together, would be able to significantly influence all matters requiring stockholder approval, including the election and removal of directors, any merger, consolidation, sale of all or substantially all of our assets, or other significant corporate transactions.

Some of these persons or entities may have interests different than yours. For example, because many of these stockholders purchased their shares at prices substantially below the price at which shares are being sold in this offering and have held their shares for a longer period, they may be more interested in selling our company to an acquirer than other investors, or they may want us to pursue strategies that deviate from the interests of other stockholders.

We are an “emerging growth company” and as a result of the reduced disclosure and governance requirements applicable to emerging growth companies, our common stock may be less attractive to investors.

We are an “emerging growth company” as defined in the Jumpstart Our Business Startups Act of 2012, or the JOBS Act, and we intend to take advantage of some of the exemptions from reporting requirements that are applicable to other public companies that are not emerging growth companies, including:

- being permitted to provide only two years of audited financial statements, in addition to any required unaudited interim financial statements, with correspondingly reduced “Management’s Discussion and Analysis of Financial Condition and Results of Operations” disclosure;
- not being required to comply with the auditor attestation requirements in the assessment of our internal control over financial reporting;
- not being required to comply with any requirement that may be adopted by the Public Company Accounting Oversight Board regarding mandatory audit firm rotation or a supplement to the auditor’s report providing additional information about the audit and the financial statements;
- reduced disclosure obligations regarding executive compensation; and
- not being required to hold a nonbinding advisory vote on executive compensation and stockholder approval of any golden parachute payments not previously approved.

We cannot predict if investors will find our common stock less attractive because we will rely on these exemptions. If some investors find our common stock less attractive as a result, there may be a less active trading market for our common stock and our stock price may be more volatile. We may take advantage of these reporting exemptions until we are no longer an emerging growth company. We will remain an emerging growth company until the earlier of (1) the last day of the fiscal year (a) following the fifth anniversary of the completion of this offering, (b) in which we have total annual gross revenue of at least \$1.0 billion, or (c) in which we are deemed to be a large accelerated filer, which means the market value of our common stock that is held by non-affiliates exceeds \$700 million as of the prior June 30th, and (2) the date on which we have issued more than \$1.0 billion in non-convertible debt during the prior three-year period.

Under Section 107(b) of the JOBS Act, emerging growth companies can delay adopting new or revised accounting standards until such time as those standards apply to private companies. We have irrevocably elected not to avail ourselves of this exemption from new or revised accounting standards and, therefore, we will be subject to the same new or revised accounting standards as other public companies that are not emerging growth companies.

If we fail to maintain proper and effective internal controls, our ability to produce accurate financial statements on a timely basis could be impaired.

After the completion of this offering, we will be subject to the reporting requirements of the Securities Exchange Act of 1934, the Sarbanes-Oxley Act and the rules and regulations of The NASDAQ Stock Market. The Sarbanes-Oxley Act requires, among other things, that we maintain effective disclosure controls and procedures and internal control over financial reporting. Commencing with our fiscal year ending December 31, 2015, we must perform system and process evaluation and testing of our internal control over financial reporting to allow management to report on the effectiveness of our internal control

[Table of Contents](#)

over financial reporting in our Form 10-K filing for that year, as required by Section 404 of the Sarbanes-Oxley Act. This will require that we incur substantial additional professional fees and internal costs to expand our accounting and finance functions and that we expend significant management efforts. Prior to this offering, we have never been required to test our internal controls within a specified period, and, as a result, we may experience difficulty in meeting these reporting requirements in a timely manner.

We may discover weaknesses in our system of internal financial and accounting controls and procedures that could result in a material misstatement of our financial statements. Our internal control over financial reporting will not prevent or detect all errors and all fraud. A control system, no matter how well designed and operated, can provide only reasonable, not absolute, assurance that the control system's objectives will be met. Because of the inherent limitations in all control systems, no evaluation of controls can provide absolute assurance that misstatements due to error or fraud will not occur or that all control issues and instances of fraud will be detected.

If we are not able to comply with the requirements of Section 404 of the Sarbanes-Oxley Act in a timely manner, or if we are unable to maintain proper and effective internal controls, we may not be able to produce timely and accurate financial statements. If that were to happen, the market price of our stock could decline and we could be subject to sanctions or investigations by The NASDAQ Stock Market, the Securities and Exchange Commission or other regulatory authorities.

We will have broad discretion in the use of proceeds from this offering and may invest or spend the proceeds in ways with which you do not agree and in ways that may not increase the value of your investment.

We will have broad discretion over the use of proceeds from this offering. You may not agree with our decisions, and our use of the proceeds may not yield any return on your investment. We expect to use the net proceeds to us from this offering to advance the development of CLS-1001 into Phase 2 and Phase 3 clinical trials and CLS-1003 into a Phase 2 clinical trial, to fund the research and development of our earlier-stage programs, including drug discovery, and for working capital and general corporate purposes. Our failure to apply the net proceeds of this offering effectively could compromise our ability to pursue our growth strategy and we might not be able to yield a significant return, if any, on our investment of these net proceeds. You will not have the opportunity to influence our decisions on how to use our net proceeds from this offering.

Because we do not anticipate paying any cash dividends on our common stock in the foreseeable future, capital appreciation, if any, will be your sole source of gains and you may never receive a return on your investment.

You should not rely on an investment in our common stock to provide dividend income. We have not declared or paid cash dividends on our common stock to date. We currently intend to retain our future earnings, if any, to fund the development and growth of our business. In addition, the terms of any existing or future debt agreements may preclude us from paying dividends. As a result, capital appreciation, if any, of our common stock will be your sole source of gain for the foreseeable future. Investors seeking cash dividends should not purchase our common stock.

We plan to use potential future operating losses and our federal and state NOL carryforwards to offset future taxable income, if any. However, our ability to use existing NOL carryforwards could be limited as a result of issuances of equity securities.

As of December 31, 2013, we had approximately \$10.4 million of federal and \$12.4 million of state net operating loss, or NOL, carryforwards. If not utilized, these federal NOL carryforwards will begin to expire at various dates beginning in 2031, and these state NOL carryforwards will begin to expire at various dates beginning in 2027. To the extent we generate taxable income in the future, we plan to use our existing NOL carryforwards and future losses to offset income that would otherwise be taxable. However, under Section 382 of the U.S. Internal Revenue Code of 1986, as amended, as interpreted by the U.S. Internal Revenue Service, the amount of benefits from our NOL carryforwards may be impaired or limited if we incur a cumulative ownership change of more than 50% over a three-year period. The completion of this offering, private placements and other transactions that have occurred, and future offerings of our securities may trigger, or may have already triggered, such an ownership change. In addition, since we will need to raise substantial additional funding to finance our operations, we may undergo further ownership changes in the future. We have not conducted an analysis as to whether such a change of ownership has occurred, but if such a change has occurred or occurs in the future, we will be limited regarding the amount of NOL carryforwards that can be utilized annually in the future to offset taxable income. Any such annual limitation may significantly reduce the value of our NOL carryforwards before they expire, which could result in greater tax liabilities than we would incur in the absence of such a limitation.

We will incur increased costs and demands upon management as a result of being a public company.

As a public company listed in the United States, we will incur significant additional legal, accounting and other costs. We estimate the additional costs we will incur as a result of being a public company to be approximately \$1.5 million to \$2.5 million annually. These additional costs could negatively affect our financial results. In addition, changing laws, regulations and standards relating to corporate governance and public disclosure, including regulations implemented by the SEC and The NASDAQ Stock Market, may increase legal and financial compliance costs and make some activities more time consuming. These laws, regulations and standards are subject to varying interpretations and, as a result, their application in practice may evolve over time as new guidance is provided by regulatory and governing bodies. We intend to invest resources to comply with evolving laws, regulations and standards, and this investment may result in increased general and administrative expenses and a diversion of management's time and attention from revenue-generating activities to compliance activities. If, notwithstanding our efforts to comply with new laws, regulations and standards, we fail to comply, regulatory authorities may initiate legal proceedings against us and our business may be harmed.

Failure to comply with these rules might also make it more difficult for us to obtain some types of insurance, including director and officer liability insurance, and we might be forced to accept reduced policy limits and coverage or incur substantially higher costs to obtain the same or similar coverage. The impact of these events could also make it more difficult for us to attract and retain qualified persons to serve on our board of directors, on committees of our board of directors or as members of senior management.

SPECIAL NOTE REGARDING FORWARD-LOOKING STATEMENTS AND INDUSTRY DATA

This prospectus contains forward-looking statements that involve substantial risks and uncertainties. The forward-looking statements are contained principally in the sections entitled “Prospectus Summary,” “Risk Factors,” “Management’s Discussion and Analysis of Financial Condition and Results of Operations” and “Business,” but are also contained elsewhere in this prospectus. In some cases, you can identify forward-looking statements by the words “may,” “might,” “will,” “could,” “would,” “should,” “expect,” “intend,” “plan,” “objective,” “anticipate,” “believe,” “estimate,” “predict,” “project,” “potential,” “continue” and “ongoing,” or the negative of these terms, or other comparable terminology intended to identify statements about the future. These statements involve known and unknown risks, uncertainties and other factors that may cause our actual results, levels of activity, performance or achievements to be materially different from the information expressed or implied by these forward-looking statements. Although we believe that we have a reasonable basis for each forward-looking statement contained in this prospectus, we caution you that these statements are based on a combination of facts and factors currently known by us and our expectations of the future, about which we cannot be certain. Forward-looking statements include statements about:

- our plans to develop and commercialize our product candidates;
- our planned clinical trials for CLS-1001 and CLS-1003;
- the timing of the availability of data from our clinical trials;
- the timing of our selection of a lead drug candidate for our CLS-1002 program;
- the timing of our planned regulatory filings;
- the timing of and our ability to obtain and maintain regulatory approvals for our product candidates;
- our expectation that CLS-1001, if approved, would be the first drug specifically indicated for macular edema associated with non-infectious uveitis;
- the clinical utility of our product candidates;
- our commercialization, marketing and manufacturing capabilities and strategy;
- our intellectual property position;
- our ability to identify additional drug candidates with significant commercial potential that are compatible with SCS administration and which are consistent with our commercial objectives; and
- our estimates regarding future revenues, expenses and needs for additional financing.

You should refer to the “Risk Factors” section of this prospectus for a discussion of important factors that may cause our actual results to differ materially from those expressed or implied by our forward-looking statements. As a result of these factors, we cannot assure you that the forward-looking statements in this prospectus will prove to be accurate. Furthermore, if our forward-looking statements prove to be inaccurate, the inaccuracy may be material. In light of the significant uncertainties in these forward-looking statements, you should not regard these statements as a representation or warranty by us or any other person that we will achieve our objectives and plans in any specified time frame, or at all. We undertake no obligation to publicly update any forward-looking statements, whether as a result of new information, future events or otherwise, except as required by law.

You should read this prospectus and the documents that we reference in this prospectus and have filed as exhibits to the registration statement, of which this prospectus is a part, completely and with the understanding that our actual future results may be materially different from what we expect. We qualify all of our forward-looking statements by these cautionary statements.

[Table of Contents](#)

This prospectus includes statistical and other industry and market data that we obtained from industry publications and research, surveys and studies conducted by third parties. Industry publications and third party research, surveys and studies generally indicate that their information has been obtained from sources believed to be reliable, although they do not guarantee the accuracy or completeness of such information.

USE OF PROCEEDS

We estimate that the net proceeds from our issuance and sale of _____ shares of our common stock in this offering will be approximately \$ _____ million, or approximately \$ _____ million if the underwriters exercise their option to purchase additional shares in full, based upon an assumed initial public offering price of \$ _____ per share, which is the midpoint of the range set forth on the cover page of this prospectus, after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us.

Each \$1.00 increase or decrease in the assumed initial public offering price of \$ _____ per share, which is the midpoint of the range set forth on the cover page of this prospectus, would increase or decrease the net proceeds to us from this offering by \$ _____ million, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same. We may also increase or decrease the number of shares we are offering. Each increase or decrease of 1.0 million in the number of shares we are offering at the assumed initial public offering price would increase or decrease the net proceeds to us from this offering by \$ _____ million.

We currently estimate that we will use the net proceeds from this offering, together with our existing cash and cash equivalents, as follows:

- approximately \$ _____ million to complete our planned Phase 2 and Phase 3 clinical trials of CLS-1001 in uveitis patients;
- approximately \$ _____ million to complete our planned Phase 2 clinical trial of CLS-1003 in RVO patients; and
- the remainder to fund further development of CLS-1003 and continued research and development of our earlier-stage programs, including drug discovery for potential new applications for our SCS microinjection technology, and for working capital and other general corporate purposes.

In the ordinary course of our business, we expect to from time to time evaluate the acquisition of, investment in or in-license of complementary products, technologies or businesses, and we could use a portion of the net proceeds from this offering for such activities. We currently do not have any agreements, arrangements or commitments with respect to any potential acquisition, investment or license.

This expected use of net proceeds from this offering represents our intentions based upon our current plans and business conditions, which could change in the future as our plans and business conditions evolve. The amounts and timing of our actual expenditures may vary significantly depending on numerous factors, including the progress of our development, the status of and results from clinical trials, as well as any collaborations that we may enter into with third parties for our product candidates, and any unforeseen cash needs.

Our management will have broad discretion in the application of the net proceeds from this offering, and investors will be relying on the judgment of our management regarding the application of the net proceeds of this offering. The timing and amount of our actual expenditures will be based on many factors, including cash flows from operations and the anticipated growth of our business. Pending these uses, we plan to invest these net proceeds in short-term, interest bearing obligations, investment-grade instruments, certificates of deposit or direct or guaranteed obligations of the United States. The goal with respect to the investment of these net proceeds is capital preservation and liquidity so that such funds are readily available to fund our operations.

DIVIDEND POLICY

We have never declared or paid any dividends on our common stock. We anticipate that we will retain all of our future earnings, if any, for use in the operation and expansion of our business and do not anticipate paying cash dividends in the foreseeable future.

CAPITALIZATION

The following table sets forth our cash and cash equivalents and our capitalization as of June 30, 2014:

- on an actual basis;
- on a pro forma basis to give effect to:
 - our issuance of an aggregate of 6,009,202 shares of Series B convertible preferred stock in August 2014 at a purchase price of \$2.70 per share, which included the conversion of principal and interest under outstanding convertible promissory notes with a principal balance of \$3.1 million into shares of Series B convertible preferred stock, and our receipt of \$13.0 million in net cash proceeds therefrom;
 - the conversion of all outstanding shares of our convertible preferred stock, including the 6,009,202 shares of Series B convertible preferred stock issued in August 2014, into an aggregate of 15,564,959 shares of our common stock, which will occur automatically upon the closing of this offering;
 - the filing of our amended and restated certificate of incorporation upon the closing of this offering; and
 - the exercise of all outstanding warrants to purchase shares of common stock that will otherwise expire upon the closing of this offering for an aggregate of 248,175 shares of common stock; and
- on a pro forma as adjusted basis to give further effect to our sale of shares of common stock in this offering at an assumed initial public offering price of \$ per share, which is the midpoint of the range set forth on the cover page of this prospectus, after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us.

[Table of Contents](#)

The following information is illustrative only of our cash and cash equivalents and capitalization following the completion of this offering and will change based on the actual initial public offering price and other terms of this offering determined at pricing. You should read this table together with “Management’s Discussion and Analysis of Financial Condition and Results of Operations” and our financial statements and the related notes appearing elsewhere in this prospectus.

	As of June 30, 2014		
	Actual	Pro forma (in thousands)	Pro forma as adjusted
Cash and cash equivalents	\$ 1,453	\$	\$
Long-term debt, including current portion	\$ 2,928	\$	\$
Convertible preferred stock:			
Series A convertible preferred stock, \$0.001 par value; 5,200,000 shares authorized, 5,198,826 shares issued and outstanding, actual; no shares authorized, issued or outstanding, pro forma and pro forma as adjusted	4,046		
Series A-1 convertible preferred stock, \$0.001 par value; 4,800,000 shares authorized, 4,356,931 shares issued and outstanding, actual; no shares authorized, issued or outstanding, pro forma and pro forma as adjusted	7,837		
Series B convertible preferred stock, \$0.001 par value; no shares authorized, issued or outstanding, actual, pro forma or pro forma as adjusted	—		
Stockholders’ equity (deficit):			
Common stock, \$0.001 par value; 17,000,000 shares authorized, 3,950,620 shares issued and outstanding, actual; shares authorized, shares issued and outstanding, pro forma; shares authorized, shares issued and outstanding, pro forma as adjusted	4		
Preferred stock, \$0.001 par value; no shares authorized, issued or outstanding, actual; shares authorized, no shares issued or outstanding, pro forma or pro forma as adjusted	—		
Additional paid-in-capital	1,279		
Deficit accumulated during the development stage	(15,926)		
Total stockholders’ equity (deficit)	(14,643)		
Total capitalization	\$ 168	\$	\$

Each \$1.00 increase or decrease in the assumed initial public offering price of \$ per share, which is the midpoint of the range set forth on the cover page of this prospectus, would increase or decrease the pro forma as adjusted amount of each of cash and cash equivalents, additional paid-in capital, total stockholders’ equity and total capitalization by \$ million, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same. Each increase or decrease of 1.0 million in the number of shares we are offering at the assumed initial public offering price would increase or decrease the pro forma as adjusted amount of each of cash and cash equivalents, additional paid-in capital, total stockholders’ equity and total capitalization by \$ million.

[Table of Contents](#)

The number of shares of common stock outstanding in the table above does not include:

- 1,619,235 shares of our common stock issuable upon the exercise of stock options outstanding under our 2011 stock incentive plan as of June 30, 2014, at a weighted average exercise price of \$0.16 per share;
- 16,550 shares of our common stock issuable upon exercise of a warrant outstanding as of June 30, 2014, at an exercise price of \$1.81 per share; and
- shares of our common stock reserved for future issuance under our equity incentive plans following this offering.

DILUTION

If you invest in our common stock in this offering, your interest will be diluted to the extent of the difference between the initial public offering price per share and the pro forma as adjusted net tangible book value per share of our common stock immediately after this offering. Net tangible book value per share is determined by dividing our total tangible assets less total liabilities and convertible preferred stock by the number of outstanding shares of our common stock.

As of June 30, 2014, we had a net tangible book deficit of \$(14.6) million, or \$(3.65) per share of common stock. On a pro forma basis, after giving effect to our issuance of an aggregate of 6,009,202 shares of Series B convertible preferred stock in August 2014, which included the conversion of principal and interest under outstanding convertible promissory notes into shares of Series B convertible preferred stock, and our receipt of \$13.0 million in net cash proceeds therefrom, as well as the conversion of the outstanding shares of our convertible preferred stock, which includes the 6,009,202 shares of our Series B convertible preferred stock issued in August 2014, into 15,564,959 shares of our common stock upon the completion of this offering, and the exercise of all outstanding warrants to purchase shares of common stock that will otherwise expire upon the closing of this offering for 248,175 shares of common stock, our pro forma net tangible book value would have been \$ million, or \$ per share of common stock.

After giving effect to the issuance and sale of shares of our common stock in this offering at an assumed initial public offering price of \$ per share, which is the midpoint of the range set forth on the cover page of this prospectus, and after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us, our pro forma as adjusted net tangible book value as of June 30, 2014 would have been \$ million, or \$ per share of common stock. This represents an immediate increase in the pro forma net tangible book value of \$ per share to existing stockholders, and an immediate dilution in the pro forma net tangible book value of \$ per share to investors purchasing shares of our common stock in this offering. The following table illustrates this per share dilution:

Assumed initial public offering price per share	\$
Actual net tangible book deficit per share as of June 30, 2014	\$(3.65)
Increase per share attributable to issuance of Series B convertible preferred stock, conversion of convertible promissory notes, conversion of convertible preferred stock and exercise of warrants	_____
Pro forma net tangible book value per share before this offering	
Increase in pro forma net tangible book value per share attributable to this offering	_____
Pro forma as adjusted net tangible book value per share after this offering	
Dilution per share to investors participating in this offering	\$ _____

The dilution information discussed above is illustrative only and will change based on the actual initial public offering price and other terms of this offering determined at pricing. Each \$1.00 increase or decrease in the assumed initial public offering price of \$ per share, which is the midpoint of the range set forth on the cover page of this prospectus, would increase or decrease our pro forma as adjusted net tangible book value by \$ million, or \$ per share, and the dilution per share to investors participating in this offering by \$ per share, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same. Each increase or decrease of 1.0 million shares in the number of shares offered by us at the assumed initial public offering price would increase or decrease our pro forma as adjusted net tangible book value per share after this offering by \$ per share and decrease or increase the dilution to investors participating in this offering by \$ per share.

[Table of Contents](#)

If the underwriters exercise their option in full to purchase _____ additional shares of common stock in this offering, the pro forma as adjusted net tangible book value per share after the offering would be \$ _____ per share, the increase in the pro forma net tangible book value per share to existing stockholders would be \$ _____ per share and the dilution to new investors purchasing common stock in this offering would be \$ _____ per share.

The following table sets forth as of June 30, 2014, on the pro forma basis described above, the differences between the number of shares of common stock purchased from us, the total consideration paid and the weighted average price per share paid by existing stockholders and by investors purchasing shares of our common stock in this offering at an assumed initial public offering price of \$ _____ per share, which is the midpoint of the range set forth on the cover page on this prospectus, before deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us:

	<u>Shares purchased</u>		<u>Total consideration</u>		<u>Weighted average</u>
	<u>Number</u>	<u>Percent</u>	<u>Amount</u>	<u>Percent</u>	<u>price per share</u>
Existing stockholders		%	\$	%	\$
New investors					
Total		100%	\$	100%	

Each \$1.00 increase or decrease in the assumed initial public offering price of \$ _____ per share, which is the midpoint of the range set forth on the cover page of this prospectus, would increase or decrease the total consideration paid by new investors by \$ _____ million, and increase or decrease the percent of total consideration paid by new investors by _____ percentage points, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same.

The table above also excludes:

- 1,619,235 shares of our common stock issuable upon the exercise of stock options outstanding under our 2011 stock incentive plan as of June 30, 2014, at a weighted average exercise price of \$0.16 per share;
- 16,550 shares of our common stock issuable upon exercise of a warrant outstanding as of June 30, 2014, at an exercise price of \$1.81 per share; and
- _____ shares of our common stock reserved for future issuance under our equity incentive plans following this offering.

The shares of our common stock reserved for future issuance under our equity benefit plans may be subject to automatic annual increases in accordance with the terms of the plans. To the extent that options or warrants are exercised, new options or other equity awards are issued under our equity incentive plans, or we issue additional shares of common stock in the future, there will be further dilution to investors participating in this offering. In addition, we may choose to raise additional capital because of market conditions or strategic considerations, even if we believe that we have sufficient funds for our current or future operating plans. If we raise additional capital through the sale of equity or convertible debt securities, the issuance of these securities could result in further dilution to our stockholders.

SELECTED FINANCIAL DATA

The following tables set forth our selected financial data for the periods indicated. The following selected statement of operations data for the years ended December 31, 2012 and 2013 and the selected balance sheet data as of December 31, 2012 and 2013 are derived from our audited financial statements appearing elsewhere in this prospectus. The selected statement of operations data for the six-month periods ended June 30, 2013 and 2014 and for the period from May 26, 2011 (date of inception) through June 30, 2014 and the selected balance sheet data as of June 30, 2014 are derived from unaudited condensed financial statements appearing elsewhere in this prospectus. The data should be read together with “Management’s Discussion and Analysis of Financial Condition and Results of Operations” and in conjunction with the financial statements, related notes, and other financial information included elsewhere in this prospectus.

The unaudited condensed financial statements include all adjustments, consisting of normal recurring accruals, which management considers necessary for a fair presentation of the financial position and the results of operations for these periods. Our historical results are not necessarily indicative of the results to be expected in the future, and our operating results for the six months ended June 30, 2014 are not necessarily indicative of the results that may be expected for the entire year ending December 31, 2014.

	Year Ended December 31,		Six Months Ended June 30,		Period From May 26, 2011 (Date of Inception) to June 30, 2014
	2012	2013	2013	2014	
(in thousands, except share and per share data)					
Statement of Operations Data:					
Costs and expenses:					
Research and development	\$ 2,354	\$ 5,045	\$ 2,204	\$ 2,562	\$ 10,168
General and administrative	1,575	2,193	1,101	1,731	5,640
Total costs and expenses	<u>3,929</u>	<u>7,238</u>	<u>3,305</u>	<u>4,293</u>	<u>15,808</u>
Loss from operations	(3,929)	(7,238)	(3,305)	(4,293)	(15,808)
Other income (expense):					
Interest expense	(3)	(23)	(6)	(99)	(127)
Interest income	<u>1</u>	<u>7</u>	<u>5</u>	<u>1</u>	<u>9</u>
Total other expense	(2)	(16)	(1)	(98)	(118)
Net loss	<u>\$ (3,931)</u>	<u>\$ (7,254)</u>	<u>\$ (3,306)</u>	<u>\$ (4,391)</u>	<u>\$ (15,926)</u>
Net loss per share of common stock — basic and diluted	<u>\$ (2.12)</u>	<u>\$ (2.45)</u>	<u>\$ (1.21)</u>	<u>\$ (1.20)</u>	<u>\$ (6.78)</u>
Weighted average shares outstanding, basic and diluted	1,853,423	2,956,285	2,728,321	3,673,629	2,349,838
Pro forma net loss per share — basic and diluted		<u>\$ (0.58)</u>		<u>\$ (0.33)</u>	
Pro forma weighted average shares outstanding — basic and diluted		12,512,042		13,229,386	

[Table of Contents](#)

	<u>As of December 31,</u>		<u>As of</u>
	<u>2012</u>	<u>2013</u>	<u>June 30,</u>
	<u>(in thousands)</u>		
Balance Sheet Data:			
Cash and cash equivalents	\$ 856	\$ 1,909	\$ 1,453
Total assets	948	2,137	2,862
Long-term debt, including current portion	150	268	2,928
Total liabilities	709	1,004	5,622
Total convertible preferred stock	4,029	11,871	11,883
Total stockholders' deficit	(3,790)	(10,738)	(14,643)

MANAGEMENT'S DISCUSSION AND ANALYSIS OF FINANCIAL CONDITION AND RESULTS OF OPERATIONS

You should read the following discussion and analysis of our financial condition and results of operations together with our financial statements and the related notes and other financial information included elsewhere in this prospectus. Some of the information contained in this discussion and analysis or set forth elsewhere in this prospectus, including information with respect to our plans and strategy for our business, includes forward-looking statements that involve risks and uncertainties. You should review the "Risk Factors" section of this prospectus for a discussion of important factors that could cause actual results to differ materially from the results described in or implied by the forward-looking statements contained in the following discussion and analysis.

Overview

We are a clinical-stage biopharmaceutical company developing first-in-class drug therapies to treat chronic, blinding diseases of the eye. Our current product candidates focus on diseases affecting the retina and choroid, especially diseases associated with macular edema, and are injected non-surgically into the suprachoroidal space, or SCS, using our proprietary microinjector. With the SCS injection procedure, our product candidates are more directly administered to the retina and choroid as compared to currently used intravitreal injections, which we believe may provide a number of benefits, including lower frequency of necessary administration and faster onset of therapeutic effect. We hold the exclusive rights to develop and commercialize drugs for non-surgical administration into the SCS. Our most advanced product candidates, CLS-1001 and CLS-1003, are based on commonly used ophthalmic therapeutic agents, which we believe will allow us to more efficiently and predictably pursue the regulatory approval of these product candidates under Section 505(b)(2) of the Federal Food, Drug, and Cosmetic Act.

CLS-1001 is in an ongoing Phase 1/2 clinical trial of patients with non-infectious uveitis for which we expect final results in the first half of 2015. In the second half of 2014, we plan to initiate a Phase 2 clinical trial in approximately 30 patients with macular edema associated with non-infectious uveitis to evaluate the safety and efficacy of our CLS-1001 administered into the SCS. In the first half of 2015, we also expect to initiate a single pivotal Phase 3 clinical trial with our CLS-1001 administered into the SCS in approximately 150 patients with macular edema associated with non-infectious uveitis, with the goal of being able to submit a Section 505(b)(2) NDA to the FDA in the first half of 2017. Under our CLS-1003 program, we are planning to initiate a Phase 2 clinical trial before the end of 2014 in approximately 40 patients with macular edema associated with RVO, a sight-threatening disorder resulting from the blockage of a retinal vein. Under our CLS-1002 program, we are evaluating a number of compounds for the treatment of wet AMD by SCS injection, including compounds with anti-VEGF activity and compounds with dual anti-VEGF and anti-PDGF activity. We plan to file an IND for this CLS-1002 program by the end of 2015. We also intend to evaluate treatments for DME. If our product candidates are approved, we plan to commercialize them with a specialty sales force of 30 to 40 representatives to target retinal specialists in the United States, and we may also pursue collaborations with third parties to commercialize any of our drugs approved for marketing outside the United States.

We are a development-stage company and have incurred net losses since our inception in May 2011. Our operations to date have been limited to organizing and staffing our company, raising capital, undertaking preclinical studies and other research and development initiatives and, beginning in 2013, conducting a clinical trial of our most advanced product candidate. To date, we have not generated any revenue and have primarily financed our operations through the private placement of our equity securities.

[Table of Contents](#)

and issuance of convertible promissory notes. We have raised net cash proceeds of \$11.7 million from the sale of convertible preferred stock and \$3.1 million from the sale of convertible promissory notes through June 30, 2014, and we raised an additional \$13.0 million of net cash proceeds through a private placement of convertible preferred stock in August 2014. As of June 30, 2014, we had a deficit accumulated during the development stage of \$15.9 million. We recorded net losses of \$3.9 million and \$7.3 million for the years ended December 31, 2012 and 2013, respectively, and \$3.3 million and \$4.4 million for the six months ended June 30, 2013 and 2014, respectively. We anticipate that a substantial portion of our capital resources and efforts in the foreseeable future will be focused on completing the development and obtaining regulatory approval and preparing for potential commercialization of our product candidates.

We expect to continue to incur significant and increasing operating losses at least for the next several years. We do not expect to generate product revenue unless and until we successfully complete development and obtain regulatory approval for one or more of our product candidates. Our net losses may fluctuate significantly from quarter to quarter and year to year, depending on the timing of our clinical trials and our expenditures on other research and development activities. We anticipate that our expenses will increase substantially as we:

- initiate our planned Phase 2 clinical trial of CLS-1001 in the second half of 2014 and our planned pivotal Phase 3 clinical trial of CLS-1001 in the first half of 2015;
- initiate our planned Phase 2 clinical trial of CLS-1003 before the end of 2014;
- continue the research and development of our other product candidates;
- seek to evaluate and develop additional product candidates;
- seek regulatory approvals for any product candidates that successfully complete clinical trials;
- ultimately establish a sales, marketing and distribution infrastructure and scale up external manufacturing capabilities to commercialize any product candidates for which we may obtain regulatory approval;
- maintain, expand and protect our intellectual property portfolio;
- hire additional clinical, manufacturing and scientific personnel;
- add operational, financial and management information systems and personnel, including personnel to support our drug development and potential future commercialization efforts; and
- begin to operate as a public company.

We anticipate that we will use approximately \$ million of the net proceeds from this offering for clinical and non-clinical costs associated with the initiation and completion of planned Phase 2 and Phase 3 clinical trials for CLS-1001 and approximately \$ million for clinical and non-clinical costs associated with the initiation and completion of the planned Phase 2 clinical trial for CLS-1003. We intend to use the remainder of the proceeds of this offering to fund the research and development of our earlier-stage programs, including drug discovery and for working capital and general corporate purposes. We expect that these funds will not be sufficient to enable us to complete all necessary development and commercially launch both of these product candidates. Accordingly, we will be required to obtain further funding through other public or private offerings of our capital stock, debt financing, collaboration and licensing arrangements or other sources. Adequate additional funding may not be available to us on acceptable terms, or at all. If we are unable to raise capital when needed or on attractive terms, we would be forced to delay, reduce or altogether cease our

research and development programs or future commercialization efforts. Although it is difficult to predict future liquidity requirements, we believe that the net proceeds of this offering, together with our existing cash and cash equivalents, will be sufficient to fund our operations for at least the next 12 months. However, our ability to successfully transition to profitability will be dependent upon achieving a level of revenues adequate to support our cost structure. We cannot assure you that we will ever be profitable or generate positive cash flow from our operating activities.

Components of Operating Results

Revenue

We have not generated any revenue from the sale of any drugs, and we do not expect to generate any revenue unless or until we obtain regulatory approval of and commercialize our product candidates. Over time, we may also seek to earn revenue by out-licensing our proprietary microinjection technology for SCS drug administration to third-party strategic collaborators.

Research and Development

Since our inception, we have focused on our development programs. Research and development expenses consist primarily of costs incurred for the research and development of our preclinical and clinical product candidates, which include:

- employee-related expenses, including salaries, benefits, travel and stock-based compensation expense for research and development personnel;
- expenses incurred under agreements with contract research organizations, or CROs, as well as contract manufacturing organizations and consultants that conduct clinical trials and preclinical studies;
- costs associated with preclinical activities and development activities;
- costs associated with technology and intellectual property licenses;
- costs for our research and development facility; and
- depreciation expense for assets used in research and development activities.

We expense research and development costs to operations as incurred. The costs for some of our development activities, such as clinical trials, are recognized based on the terms of underlying agreements, as well as an evaluation of the progress to completion of specific tasks using data such as patient enrollment, clinical site activations and additional information provided to us by our vendors about their actual costs occurred.

Expenses related to activities, such as manufacturing and stability and toxicology studies, that are supportive of a product candidate itself, are classified as direct non-clinical costs. Expenses related to clinical trials and similar activities, including costs associated with CROs, are classified as direct clinical costs. Expenses related to activities that support more than one development program or activity, such as salaries, stock-based compensation and depreciation, are not classified as direct clinical costs or non-clinical costs and are separately classified as unallocated.

For the year ended December 31, 2012, substantially all of our research and development expenses related to the non-clinical development of CLS-1001. For the year ended December 31, 2013 and the six months ended June 30, 2013 and 2014, substantially all of our research and development expenses related to

[Table of Contents](#)

the non-clinical and clinical development of CLS-1001. From inception through June 30, 2014, we have incurred \$10.2 million in research and development expenses, of which only \$0.2 million was expended prior to January 1, 2012. Prior to January 1, 2012, we did not allocate any research and development expenses to any specific product candidate.

The following table shows our research and development expenses by type of activity for the six months ended June 30, 2013 and 2014, the years ended December 31, 2012 and 2013 and the period from May 26, 2011 (date of inception) to June 30, 2014.

	Six Months Ended June 30,		Year Ended December 31,		Period from May 26, 2011 (Date of Inception) to June 30, 2014
	2013	2014	2012	2013	
	(in thousands)				
CLS-1001:					
Direct non-clinical	\$ 635	\$ 675	\$ 926	\$1,326	\$ 2,927
Direct clinical	30	91	—	173	264
Total	665	766	926	1,499	3,191
CLS-1002:					
Direct non-clinical	50	146	—	153	299
Direct clinical	38	6	123	44	173
Total	88	152	123	197	472
CLS-1003:					
Direct non-clinical	—	85	—	45	130
Unallocated	1,451	1,559	1,305	3,304	6,375
Total research and development expense	<u>\$ 2,204</u>	<u>\$ 2,562</u>	<u>\$ 2,354</u>	<u>\$ 5,045</u>	<u>\$ 10,168</u>

Our expenses related to clinical trials are based on estimates of patient enrollment and related expenses at clinical investigator sites as well as estimates for the services received and efforts expended under contracts with research institutions, consultants and CROs that conduct and manage clinical trials on our behalf. We generally accrue expenses related to clinical trials based on contracted amounts applied to the level of patient enrollment and activity according to the protocol. If future timelines or contracts are modified based upon changes in the clinical trial protocol or scope of work to be performed, we would modify our estimates of accrued expenses accordingly on a prospective basis. Historically, any such modifications have not been material.

Research and development activities are central to our business model. Product candidates in later stages of clinical development generally have higher development costs than those in earlier stages of clinical development, primarily due to the increased size and duration of later-stage clinical trials. We expect our research and development expenses to increase over the next several years as we progress CLS-1001 and our other product candidates through clinical development. However, it is difficult to determine with certainty the duration and completion costs of our current or future preclinical programs and clinical trials of our product candidates, or if, when or to what extent we will generate revenues from the commercialization and sale of any of our product candidates that obtain regulatory approval. We may never succeed in achieving regulatory approval for any of our product candidates.

The duration, costs and timing of clinical trials and development of our product candidates will depend on a variety of factors that include, but are not limited to, the following:

- number of trials required for approval and any requirement for extension trials;

Table of Contents

- per patient trial costs;
- the number of patients that participate in the trials;
- the number of sites included in the trials;
- the countries in which the trial is conducted;
- the length of time required to enroll eligible patients;
- the number of doses that patients receive;
- the drop-out or discontinuation rates of patients;
- potential additional safety monitoring or other studies requested by regulatory agencies;
- the duration of patient follow-up; and
- the efficacy and safety profile of the product candidate.

In addition, the probability of success for each product candidate will depend on numerous factors, including competition, manufacturing capability and commercial viability. We will determine which programs to pursue and how much to fund each program in response to the scientific and clinical success of each product candidate, as well as an assessment of each candidate's commercial potential.

General and Administrative

General and administrative expenses consist primarily of salaries and other related costs, including stock-based compensation, for personnel in executive, finance, accounting, business development, and human resources functions. Other significant costs include facility costs not otherwise included in research and development expenses, legal fees relating to intellectual property and corporate matters and fees for accounting and consulting services.

We anticipate that our general and administrative expenses will increase in the future to support our continued research and development activities, potential commercialization of our product candidates and increased costs of operating as a public company. These increases will likely include increased costs related to the hiring of additional personnel and fees to outside consultants, lawyers and accountants, among other expenses. Additionally, we anticipate increased costs associated with being a public company including expenses related to services associated with maintaining compliance with NASDAQ listing rules and SEC requirements, insurance, and investor relations costs. In addition, if CLS-1001 or any of our other product candidates obtains regulatory approval for marketing, we expect to incur expenses associated with building a sales and marketing team. However, we do not expect to receive any such regulatory approval for at least the next several years.

Other Income (Expense)

Other income consists of interest income earned on our cash and cash equivalents. Interest income is not considered significant to our financial statements, but we expect our interest income to increase following this offering as we invest the net proceeds from this offering pending their use in operations.

Other expense consists of interest accrued under promissory notes and amortization of debt discounts arising from the preferred stock purchase warrant described in the footnotes to our financial statements appearing elsewhere in this prospectus.

Critical Accounting Policies and Significant Judgments and Estimates

Our management's discussion and analysis of our financial condition and results of operations is based on our financial statements, which have been prepared in accordance with accounting principles generally accepted in the United States of America, or GAAP. The preparation of these financial statements requires us to make estimates, judgments and assumptions that affect the reported amounts of assets and liabilities, disclosure of contingent assets and liabilities as of the dates of the balance sheets and the reported amounts of expenses during the reporting periods. In accordance with GAAP, we evaluate our estimates and judgments on an ongoing basis. Significant estimates include assumptions used in the determination of stock-based compensation and some of our research and development expenses. We base our estimates on historical experience and on various other factors that we believe are reasonable under the circumstances, the results of which form the basis for making judgments about the carrying value of assets and liabilities that are not readily apparent from other sources. Actual results may differ from these estimates under different assumptions or conditions.

We define our critical accounting policies as those accounting principles generally accepted in the United States of America that require us to make subjective estimates and judgments about matters that are uncertain and are likely to have a material impact on our financial condition and results of operations, as well as the specific manner in which we apply those principles. While our significant accounting policies are more fully described in Note 2 to our financial statements appearing elsewhere in this prospectus, we believe the following are the critical accounting policies used in the preparation of our financial statements that require significant estimates and judgments.

Accrued Expenses

As part of the process of preparing our financial statements, we are required to estimate accrued expenses. This process involves reviewing open contracts and purchase orders, communicating with applicable vendor personnel to identify services that have been performed on our behalf and estimating the level of service performed and the associated cost incurred for the service when we have not yet been invoiced or otherwise notified of actual cost. We make estimates of our accrued expenses as of each balance sheet date in our financial statements based on facts and circumstances known to us. We periodically confirm the accuracy of our estimates with the service providers and make adjustments if necessary. Examples of estimated accrued expenses include fees paid to CROs and investigative sites in connection with clinical trials.

We accrue our expenses related to clinical trials based on our estimates of the services received and efforts expended pursuant to contracts with multiple research institutions and CROs that conduct research activities or manage clinical trials on our behalf. The financial terms of these agreements are subject to negotiation, vary from contract to contract and may result in uneven payment flows. Payments under some of these contracts depend on factors such as the successful enrollment of patients and the completion of clinical trial milestones. In accruing service fees, we estimate the time period over which services will be performed and the level of effort to be expended in each period. If the level of effort varies from our estimate, we will adjust the accrual accordingly. If we underestimate or overestimate the level of services performed or the costs of these services, our actual expenses could differ from our estimates. Although we do not currently anticipate the future settlement of existing accruals to differ materially from our estimates, our understanding of the status and timing of services performed relative to the actual status and timing of services performed may vary and could result in us reporting amounts that are too high or too low for any period. There have been no material changes in estimates for the periods presented in our financial statements.

Fair Value Measurements

We record some of our financial assets and liabilities at fair value in accordance with the provisions of Accounting Standards Codification, or ASC, Topic 820 on fair value measurements. As defined in the guidance, fair value, defined as an exit price, represents the amount that would be received to sell an asset or pay to transfer a liability in an orderly transaction between market participants. As a result, fair value is a market-based approach that should be determined based on assumptions that market participants would use in pricing an asset or a liability. As a basis for considering these assumptions, the guidance defines a three-tier value hierarchy that prioritizes the inputs used in the valuation methodologies in measuring fair value.

- Level 1 — Unadjusted quoted prices in active, accessible markets for identical assets or liabilities.
- Level 2 — Other inputs that are directly or indirectly observable in the marketplace.
- Level 3 — Unobservable inputs that are supported by little or no market activity.

The fair value hierarchy also requires an entity to maximize the use of observable inputs and minimize the use of unobservable inputs when measuring fair value.

Our material financial instruments at December 31, 2012, December 31, 2013 and June 30, 2014 consisted primarily of cash and cash equivalents, long-term debt and stock purchase warrant liabilities. The fair value of cash and cash equivalents and long-term debt approximate their respective carrying values due to the short-term nature of these instruments. We have determined the preferred stock purchase warrant, the value of which is recorded as an adjustment to long-term debt, to be valued under Level 3.

Stock-Based Compensation

We recognize compensation costs related to stock options and restricted stock granted to employees, directors and consultants ratably over the requisite service period, which in most cases is the vesting period of the award for employees, based on the estimated fair value of the awards on the date of grant. Compensation expense for options granted to non-employees is determined as the fair value of consideration received or the fair value of the equity instruments issued, whichever is more reliably measured. The fair value of the awards granted to non-employees is re-measured each period until the related service is complete.

Stock-based compensation expense was \$0.3 million for each of the years ended December 31, 2012 and 2013 and \$0.1 million and \$0.2 million for the six months ended June 30, 2013 and 2014, respectively.

Significant Factors, Assumptions and Methodologies Used in Determining Fair Value

Determining the appropriate fair value measurement of stock-based awards requires the use of subjective assumptions. In the absence of a public trading market for our common stock, and in preparation for this offering, we conducted retrospective assessments of the valuation of our common stock as of December 31, 2012 and 2013 and a contemporaneous valuation as of June 30, 2014, in each case as described below in “— Common Stock Valuations.” The determination of the fair value measurement of options using the Black-Scholes option pricing model is affected by our estimated common stock fair values as well as assumptions regarding a number of other subjective variables. These other variables include the expected term of the options, our expected stock price volatility over the expected term of the options, stock option exercise and cancellation behaviors, risk-free interest rates and expected dividends.

[Table of Contents](#)

We estimated the fair value of stock options at the grant date using the following assumptions:

- *Fair Value of our Common Stock.* Since no public market exists for our stock, we must estimate its fair value, as discussed in “— Common Stock Valuations” below.
- *Volatility.* As we do not have a trading history for our common stock, we utilize data from a representative group of publicly traded companies to estimate expected stock price volatility. We selected representative companies from the pharmaceutical industry with similar characteristics to us, including stage of product development and therapeutic focus.
- *Expected Term.* We used the simplified method as prescribed by the SEC Staff Accounting Bulletin No. 107, *Share-Based Payment*, as we do not have sufficient historical exercise and post-vesting termination data to provide a reasonable basis upon which to estimate the expected term of stock options granted to employees.
- *Risk-Free Rate.* The risk-free interest rate is based on the yields of U.S. Treasury securities with maturities similar to the expected time to liquidity.
- *Forfeitures.* Forfeitures are estimated such that we only recognize expense for the shares expected to vest, and adjustments are made if actual forfeitures differ from those estimates. We estimated our annual forfeiture rates to be zero for 2012 and 2013.
- *Dividend Yield.* We have never declared or paid any cash dividends and do not presently plan to pay cash dividends in the foreseeable future.

The table below lists the weighted average assumptions utilized in the Black-Scholes option pricing model for the years ended December 31, 2012 and 2013 and for the six months ended June 30, 2013. We did not grant any stock options during the six months ended June 30, 2014, although as described below we did grant stock options subsequent to June 30, 2014.

	Year Ended December 31,		Six Months Ended June 30,
	2012	2013	2013
Expected term (years)	7.00	7.00	7.00
Expected stock price volatility	156.84%	97.02%	100.06%
Risk-free interest rate	1.07%	1.69%	1.28%
Dividend yield	0.00%	0.00%	0.00%

The estimation of the number of stock awards that will ultimately vest requires judgment, and to the extent actual results or updated estimates differ from our current estimates, such amounts will be recorded as a cumulative adjustment in the period in which estimates are revised.

Common Stock Valuations

We are a private company with no active public market for our common stock. Our common stock valuations are determined by our board of directors in its sole discretion based on recommendations from management and, beginning in 2014, taking into account advice and assistance provided by a third-party valuation consultant engaged to assist us in connection with such valuations. The valuations of our common stock were determined utilizing guidelines outlined in the American Institute of Certified Public Accountants Practice Aid, *Valuation of Privately-Held-Company Equity Securities Issued as Compensation*, referred to as the Practice Aid. The methodologies used to determine fair value of our common stock included estimating the fair value of the enterprise and then allocating this value to all classes of equity securities using a combination of the option pricing method and, beginning with our retrospective valuation

[Table of Contents](#)

as of December 31, 2013, the initial public offering scenario within the probability-weighted expected return method, described in more detail below.

The assumptions used in the valuation models that ultimately determine the fair value of our common stock as of the valuation date are based on numerous objective and subjective factors combined with management judgment, including the following:

- our results of operations, financial position and the status of our research and development efforts;
- the composition of, and changes to, our management team and board of directors;
- the lack of liquidity of our common stock as a private company;
- our stage of development and business strategy and the material risks related to our business and industry;
- our discounted future cash flows, based on our projected operating results;
- the potential impact on our common stock of liquidation preference rights of our convertible preferred stock;
- the valuation of publicly traded companies in the life sciences and pharmaceutical industry sectors, as well as recently completed mergers and acquisitions of peer companies;
- any external market conditions affecting the life sciences and pharmaceutical industry sectors;
- the likelihood of achieving a liquidity event for the holders of our common stock and stock options, such as an initial public offering, or IPO, or a sale of our company, given prevailing market conditions;
- the state of the IPO market for similarly situated privately held biotechnology companies; and
- any recent contemporaneous valuations prepared in accordance with methodologies outlined in the Practice Aid.

The following table presents the grant dates and related exercise prices of stock options that we granted from January 1, 2013 through the date of this prospectus, along with the corresponding exercise price for each option grant and the fair values per share utilized to calculate stock-based compensation expense. Refer to “Estimate of Fair Value for Option Grants During Six Months Ended June 30, 2013,” “Retrospective Valuation as of December 31, 2013” and “Contemporaneous Valuation as of June 30, 2014” for additional information.

Date of Grant	Number of Shares Underlying Options	Exercise Price Per Option	Common Stock Fair Value Per Share on Grant Date
3/1/2013	585,500	\$ 0.18	\$ 0.50(1)
5/21/2013	5,000	0.18	0.50(1)
6/17/2013	100,000	0.18	0.50(1)
8/7/2013	165,000	0.18	0.66(1)
10/16/2013	50,000	0.18	0.66(1)
11/25/2013	500,000	0.18	0.66(1)
12/17/2013	30,000	0.18	0.66(1)
8/12/2014	70,000	1.40	1.40

(1) We assessed the fair value of our common stock subsequent to the grant date of these awards, as described below.

[Table of Contents](#)

Based on an assumed initial public offering price of \$ per share, the midpoint of the price range set forth on the cover page of this prospectus, the intrinsic value of all stock options outstanding as of June 30, 2014 was \$ million, of which \$ million and \$ million related to stock options that were vested and unvested, respectively, at that date.

Common Stock Valuation Methodologies

The Practice Aid describes market, income and cost approaches to valuing equity securities, each of which approaches is summarized below.

Market Approach. The market approach uses similar companies or transactions in the marketplace. When using the guideline company method of the market approach in determining the fair value of common stock, a company identifies companies similar to its business and uses these guideline companies to develop relevant market multiples and ratios, which are then applied to its financial forecasts to create an indication of total equity value. When using the similar transaction methodology of the market approach in determining the fair value of common stock, a company uses publicly disclosed data from arm's-length transactions involving similar companies to develop relationships or value measures between the prices paid for the target companies and the underlying financial performance of those companies. These value measures are then applied to a company's applicable operating data to create an indication of total equity value.

Income Approach. For the income approach, a company uses the discounted free cash flow method, which is based on the premise that equity value as of the respective valuation date is equal to the projected future free cash flows and expected terminal value of the business, discounted by a required rate of return that investors would demand given the risks of ownership and the risks associated with achieving the stream of projected future free cash flows.

Cost Approach. The cost approach involves identifying a company's significant tangible assets, estimating the individual current market values of each and then totaling them to derive the value of the business as a whole. A company can use the cost approach to value its adjusted net assets available to common stockholders if it were forced to liquidate its assets if its business model failed and the company was unable to raise additional financing.

Methods Used to Allocate Our Enterprise Value to Classes of Securities

In accordance with the Practice Aid, we considered the various methods for allocating the enterprise value across our classes and series of capital stock to determine the fair value of our common stock at the valuation date. The methods we considered consisted of the following:

Current Value Method. Under the current value method, once the fair value of the enterprise is established, the value is allocated to the various series of preferred and common stock based on their respective seniority, liquidation preferences or conversion values, whichever is greatest.

Option Pricing Method, or OPM. Under the option pricing method, shares are valued by creating a series of call options with exercise prices based on the liquidation preferences and conversion terms of each equity class. The values of the preferred and common stock are inferred by analyzing these options.

Probability-Weighted Expected Return Method, or PWERM. The PWERM is a scenario-based analysis that estimates the value per share based on the probability-weighted present value of expected future investment returns, considering each of the possible outcomes available to us, as well as the economic and control rights of each share class.

[Table of Contents](#)

Retrospective Valuation as of December 31, 2012

In view of our relatively early stage of development as of December 31, 2012, as well as the state of the market for biopharmaceutical initial public offerings at that time, we conducted a retrospective valuation of our common stock as of December 31, 2012 using the OPM. In conducting this valuation, we used a market approach to backsolve for an estimate of our enterprise value based on our Series A-1 convertible preferred stock financing, which occurred shortly thereafter in January 2013. Although we did not rely on the guideline company or similar transaction methodologies to determine our enterprise value as part of the market approach, we did review them to gain a deeper insight into the companies in the ophthalmic therapeutics market and to evaluate prevailing market conditions.

Once we determined an estimate of our enterprise value, we then allocated the estimated enterprise value to the various classes of our equity securities using the OPM, which analyzes the rights of the common stock relative to those of the preferred stock by assessing the “break points,” or the points at which it is economically viable for the holders of preferred stock to convert their preferred stock into common stock. At the December 31, 2012 valuation date, the model incorporated an assumed time to a liquidity event of 2.5 years. After applying a 40% discount for lack of marketability, the fair value per share of common stock on a non-marketable interest basis under the OPM was \$0.30 as of December 31, 2012. We used basic put option and Finnerty put option analyses to estimate the discount for lack of marketability. These models are commonly used for estimating illiquidity discounts for securities.

Estimate of Fair Value for Option Grants During Six Months Ended June 30, 2013

In January 2013, we completed our Series A-1 preferred stock financing, raising gross proceeds of \$7.9 million, which we believe resulted in an increased value of our company and our common stock. Also in January 2013, we received approval of our IND from the FDA to initiate our Phase 1/2 clinical trial in our CLS-1001 program. Therefore, for purposes of our retrospective assessment of the fair value of our common stock as of dates after January 2013, we believed that it was appropriate to increase our estimate of the fair value of our common stock from the December 31, 2012 estimated fair value of \$0.30 per share. In deriving this estimate, we interpolated between \$0.30 per share and the \$0.66 per share estimated fair value that we utilized for our grants made in the second half of 2013, as described below. Based on this interpolation, we estimated the fair value per share of common stock to be \$0.50 per share, which we have applied retrospectively to all of our stock option grants made during the first half of 2013.

Retrospective Valuation as of December 31, 2013

By the end of 2013, based on our board of directors’ review of overall market conditions, the improving market for biopharmaceutical initial public offerings and the progress of our clinical and preclinical development, our board of directors determined that a shift was occurring with respect to the potential valuation of our common stock in a liquidity event scenario, and we therefore performed a valuation for our common stock as of December 31, 2013.

In conducting this valuation, we used a market approach to backsolve for an estimate of our enterprise value based upon anticipated transactions involving convertible preferred stock. Specifically, based on our capital plan at the valuation date, we estimated that we would need additional funding during 2014. At the valuation date, we expected that such a new investment, a Series B preferred stock financing, would be completed at the same price as our prior Series A-1 convertible preferred stock, issued in January and February 2013, but would be senior to it in terms of liquidation preference. Although we did not rely on the

[Table of Contents](#)

guideline company or similar transaction methodologies to determine our enterprise value as part of the market approach, we did review them to gain a deeper insight into the companies in the ophthalmic therapeutics market and to evaluate prevailing market conditions.

Once we determined an estimate of our enterprise value, we then allocated the estimated enterprise value to the various classes of our equity securities using the OPM, which analyzes the rights of the common stock relative to those of the preferred stock by assessing the “break points,” or the points at which it is economically viable for the holders of preferred stock to convert their preferred stock into common stock. At the December 31, 2013 valuation date, the model incorporated an assumed time to a liquidity event of two years. After applying a 30% discount for lack of marketability, the fair value per share of common stock on a non-marketable interest basis under the OPM was \$0.32 as of December 31, 2013. We used basic put option and Finnerty put option analyses to estimate the discount for lack of marketability. These models are commonly used for estimating illiquidity discounts for securities.

As of December 31, 2013, our board of directors had authorized management to begin preparing for a potential initial public offering, or IPO, of our common stock, although we did not select underwriters for this offering until April 2014. Because an IPO scenario was considered to be a possible liquidity event, we also used the PWERM to estimate our enterprise value under this scenario and to estimate the fair value of our common stock in such a scenario. For this scenario, we assumed an IPO at the end of 2014. We reviewed a number of IPOs completed by life sciences and biopharmaceutical companies during 2013 and estimated an enterprise value for our company that would have been somewhat below the first quartile of the enterprise values of recent IPOs, after taking into account our stage of development as compared to the reviewed companies. After applying a 20% discount rate to the estimated future enterprise value at the time of an IPO, and after applying a 20% discount for lack of marketability using basic put option and Finnerty put option analyses, the fair value per share of common stock on a non-marketable basis under the PWERM was \$2.01 as of December 31, 2013.

Based upon our estimates of the probabilities of the future outcomes, we then attributed an 80% weighting to the OPM and a 20% weighting to the IPO scenario within the PWERM. This resulted in a weighted per share value of \$0.66 for our common stock as of December 31, 2013. For financial reporting purposes in determining the estimated fair value of the stock options, this value has been applied retrospectively to all of our stock option grants made during the six months ended December 31, 2013.

Contemporaneous Valuation as of June 30, 2014

We deemed it appropriate to obtain a valuation of our common stock as of June 30, 2014. In this valuation, we used the same methodology as we had used for our December 31, 2013 retrospective valuation.

Once we determined an estimate of our enterprise value, we allocated the estimated enterprise value to the various classes of our equity securities using the OPM. At the June 30, 2014 valuation date, the model incorporated an assumed time to a liquidity event of two years. After applying a 30% discount for lack of marketability using basic put option and Finnerty put option analyses, the fair value per share of common stock on a non-marketable interest basis under the OPM was \$0.15 as of June 30, 2014.

Because an IPO scenario was considered to be a possible liquidity event as of June 30, 2014, we also used the PWERM to estimate our enterprise value under this scenario and to estimate the fair value of our common stock in such a scenario. For this scenario, we assumed an IPO at the end of 2014. We reviewed a number of IPOs completed by life sciences and biopharmaceutical companies during 2013 and 2014 and

[Table of Contents](#)

estimated an enterprise value for our company that would have been somewhat below the first quartile of the enterprise values of recent IPOs, after taking into account our stage of development as compared to the reviewed companies. After applying a 20% discount rate to the estimated future enterprise value at the time of an IPO, and after applying a 10% discount for lack of marketability using basic put option and Finnerty put option analyses, the fair value per share of common stock on a non-marketable basis under the PWERM was \$3.89 as of June 30, 2014.

Based upon our estimates of the probabilities of the future outcomes, we then attributed a 67% weighting to the OPM and a 33% weighting to the IPO scenario within the PWERM. This resulted in a weighted per share value of \$1.40 for our common stock as of June 30, 2014.

2014 Option Grants

Our board of directors granted options to purchase common stock on August 12, 2014, with each option having an exercise price of \$1.40 per share. In establishing this exercise price, our board of directors considered input from management, giving substantial weight to the valuation we conducted of our common stock as of June 30, 2014. Our board of directors concluded that there were no events or circumstances that occurred between June 30, 2014 and August 12, 2014 that were indicative of a change in the fair value of our common stock and therefore determined that the fair value of our common stock on that grant date was \$1.40 per share.

Determination of Estimated Offering Price

In April 2014, we selected underwriters for this offering. The midpoint of the preliminary range for the initial public offering as determined by us and the underwriters was \$ per share. In comparison, our estimate of the fair value of our common stock was \$1.40 per share as of the June 30, 2014 valuation. We note that, as is typical in IPOs, the preliminary range was not derived using a formal determination of fair value, but was determined based upon discussions between us and the underwriters. Among the factors that were considered in setting this range were our prospects and the history of and prospects for our industry, the general condition of the securities markets and the recent market prices of, and the demand for, publicly traded common stock of generally comparable companies. We believe that the difference between the fair value of our common stock as of June 30, 2014 and the midpoint of the price range for this offering is the result of these factors as well as the fact that the estimated IPO price range necessarily assumes that the initial public offering has occurred, a public market for our common stock has been created and that our preferred stock converted into common stock in connection with the IPO, and therefore excludes any discount for lack of marketability of our common stock, which was factored into the June 30, 2014 valuation.

Results of Valuation Models May Vary

Valuation models require the input of highly subjective assumptions and estimates. These judgments and estimates include assumptions regarding our future operating performance, the time to completing an IPO or other liquidity event and the determinations of the appropriate valuation methods. Because our common stock has characteristics significantly different from that of publicly traded common stock and because changes in the subjective input assumptions can materially affect our fair value estimates, the existing models do not necessarily provide a reliable, single measure of the fair value of our common stock. If we had made different assumptions, our stock-based compensation expense, our net loss and net loss per share could have been

[Table of Contents](#)

significantly different. The foregoing valuation methodologies are not the only valuation methodologies available and will not be used to value our common stock once this offering is complete. We cannot make assurances as to any particular valuation for our stock. Accordingly, investors are cautioned not to place undue reliance on the foregoing valuation methodologies as an indicator of future stock prices.

Tax Valuation Allowance

We recorded deferred tax assets of \$4.3 million as of December 31, 2013, which have been fully offset by a valuation allowance due to uncertainties surrounding our ability to realize these tax benefits. We record a valuation allowance if it is more likely than not that a deferred tax asset will not be realized. We provided a full valuation allowance on our deferred tax assets that primarily consist of cumulative net operating losses, or NOLs, of \$10.4 million for the period from our inception on May 26, 2011 to December 31, 2013. We incurred a net loss of \$4.4 million for the six months ended June 30, 2014. Due to our three-year cumulative loss position, history of operating losses and losses expected to be incurred in the foreseeable future, we considered it necessary to provide for a full valuation allowance against our net deferred tax assets.

Our deferred tax assets are primarily composed of federal and state tax NOL carryforwards. As of December 31, 2013, we had federal NOL carryforwards of \$10.4 million and state NOL carryforwards of \$12.4 million available to reduce future taxable income, if any. These federal NOL carryforwards will begin to expire at various dates starting in 2031. The state NOL carryforwards will begin to expire at various dates starting in 2027. In general, if we experience a greater than 50 percentage point aggregate change in ownership of specified significant stockholders over a three-year period, utilization of our pre-change NOL carryforwards will be subject to an annual limitation under Section 382 of the U.S. Internal Revenue Code of 1986, as amended, or the Code, and similar state laws. Such limitations may result in expiration of a portion of the NOL carryforwards before utilization and may be substantial. If we experience a Section 382 ownership change in connection with this offering or as a result of future changes in our stock ownership, some of which changes are outside our control, the tax benefits related to the NOL carryforwards may be further limited or lost.

Results of Operations for the Six Months Ended June 30, 2013 and 2014

The following table sets forth our results of operations for the six months ended June 30, 2013 and 2014.

	Six Months Ended		Period-to-Period Change
	June 30,		
	2013	2014	
	(in thousands)		
Costs and expenses:			
Research and development	\$ 2,204	\$ 2,562	\$ 358
General and administrative	1,101	1,731	630
Total costs and expenses	<u>3,305</u>	<u>4,293</u>	<u>988</u>
Loss from operations	(3,305)	(4,293)	(988)
Other income (expense):			
Interest expense	(6)	(99)	(93)
Interest income	5	1	(4)
Total other expense	<u>(1)</u>	<u>(98)</u>	<u>(97)</u>
Net loss	<u><u>\$ (3,306)</u></u>	<u><u>\$ (4,391)</u></u>	<u><u>\$ (1,085)</u></u>

[Table of Contents](#)

Research and development. Research and development expense increased by \$0.4 million, from \$2.2 million for the six months ended June 30, 2013 to \$2.6 million for the six months ended June 30, 2014, an increase of 16%. The increase was primarily attributable to a \$0.2 million increase in costs related to the design, testing and manufacture of our microinjector, and \$0.2 million in non-clinical expenses during the six months ended June 30, 2014 related to the other product candidates in our pipeline, including CLS-1003, for which we are planning to begin a Phase 2 clinical trial in RVO patients before the end of 2014. We also incurred increased personnel costs during the six months ended June 30, 2014 as compared to the prior year, offset by lower costs of pre-clinical studies in our CLS-1001 program.

General and administrative. General and administrative expense increased by \$0.6 million, from \$1.1 million for the six months ended June 30, 2013 to \$1.7 million for the six months ended June 30, 2014, an increase of 57%. The increase was primarily attributable to increases in personnel costs, including stock-based compensation and fees related to the audit of our financial statements in preparation for this offering.

Results of Operations for the Years Ended December 31, 2012 and 2013

The following table sets forth our results of operations for the years ended December 31, 2012 and 2013.

	Year Ended December 31,		Period-to-Period Change
	2012	2013	
	(in thousands)		
Costs and expenses:			
Research and development	\$ 2,354	\$ 5,045	\$ 2,691
General and administrative	1,575	2,193	618
Total costs and expenses	3,929	7,238	3,309
Loss from operations	(3,929)	(7,238)	(3,309)
Other income (expense):			
Interest expense	(3)	(23)	(20)
Interest income	1	7	6
Total other expense	(2)	(16)	(14)
Net loss	<u>\$(3,931)</u>	<u>\$(7,254)</u>	<u>\$ (3,323)</u>

Research and development. Research and development expense increased by \$2.6 million, from \$2.4 million for the year ended December 31, 2012 to \$5.0 million for the year ended December 31, 2013, an increase of 114%. The increase was primarily attributable to higher compensation, travel and facilities costs of \$1.4 million resulting from new hires in 2013, as well as an increase of \$0.6 million in direct clinical expenses for our CLS-1001 program, which began clinical development in the second half of 2013, and an increase of \$0.7 million in preclinical expenses related to our other programs, including our earlier-stage RVO and wet AMD programs, during the year ended December 31, 2013.

General and administrative. General and administrative expense increased by \$0.6 million, from \$1.6 million for the year ended December 31, 2012 to \$2.2 million for the year ended December 31, 2013, an increase of 39%. The increase was primarily attributable to an increase of \$0.5 million in costs related to new hires in our general and administrative functions, including compensation and travel costs. Additionally, we had an increase of \$0.1 million in professional fees as compared to the prior year.

Liquidity and Capital Resources

Sources of Liquidity

We have incurred losses and experienced negative operating cash flows since our inception and anticipate that we will continue to incur losses for at least the next several years. We expect that our research and development and general and administrative expenses will increase over historical levels and, as a result, we will need additional capital to fund our operations, which we may obtain from additional public offerings, debt financing, collaboration and licensing arrangements or other sources.

For the period from our inception on May 26, 2011 to June 30, 2014, we have cumulative net cash used by operating activities of \$13.4 million and cumulative net losses of \$15.9 million. Since our inception, we have funded operations primarily through the sale of convertible preferred stock and the issuance of convertible promissory notes. We have raised net cash proceeds of \$11.7 million from the sale of convertible preferred stock and \$3.1 million from the sale of convertible promissory notes through June 30, 2014, and we raised an additional \$13.0 million of net cash proceeds through a private placement of convertible preferred stock in August 2014. As of December 31, 2013 and June 30, 2014, we had cash and cash equivalents of \$1.9 million and \$1.5 million, respectively. We invest any cash in excess of our immediate requirements primarily with a view to liquidity and capital preservation. As of December 31, 2013 and June 30, 2014, our funds were held in cash and money market funds.

In April 2014, we authorized the sale of convertible promissory notes, or the bridge notes, to our existing stockholders, including two of our executive officers and one of our directors in their individual capacities, in the aggregate principal amount of \$6.0 million. In April 2014, we issued \$3.0 million in aggregate principal amount of bridge notes. We do not expect to issue the remaining \$3.0 million in aggregate principal amount of bridge notes. The outstanding notes accrued interest at an annual rate of 7%, with principal plus interest due upon maturity in April 2015 unless earlier converted. The bridge notes were convertible upon the occurrence of a qualified financing. Our August 2014 Series B convertible preferred stock financing was a qualified financing as contemplated by the bridge notes, and the principal and interest under all of the bridge notes was converted automatically into an aggregate of 1,137,644 shares of Series B convertible preferred stock in connection with this financing. These shares of Series B convertible preferred stock are convertible into shares of our common stock upon the completion of this offering. In connection with the issuance of the bridge notes, we also issued warrants to the lenders to purchase an aggregate of 248,175 shares of our common stock at an exercise price of \$0.01 per share. These warrants expire upon the closing of this offering, and therefore we expect that these warrants will be exercised in connection with the completion of this offering and that we will issue 248,175 shares of our common stock upon their exercise.

In April 2013, we entered into a loan agreement with an entity affiliated with the State of North Carolina under which we borrowed an aggregate of \$125,000. The outstanding balance bore interest at an annual rate of 4.25%. All unpaid principal, together with unpaid and accrued interest, were due and payable in February 2016 or, if earlier, upon the occurrence of specified events. We repaid this note in full in May 2014. In connection with the initial loan, we also issued this lender a warrant to purchase 16,550 shares of our Series A-1 convertible preferred stock, which will become a warrant to purchase 16,550 shares of our common stock following the completion of this offering.

In December 2012, we entered into a \$150,000 unsecured promissory note with a lender, which bore interest at an annual rate of 5%. All unpaid principal, together with unpaid and accrued interest converted into an aggregate of 60,291 shares of Series B convertible preferred stock in connection with our August 2014 Series B convertible preferred stock financing.

[Table of Contents](#)

Between June 2011 and December 2011, we borrowed an aggregate of \$100,000 from Daniel White, our chief executive officer. These advanced amounts, plus accrued interest of approximately \$5,000, were converted into shares of our Series A convertible preferred stock during the year ended December 31, 2012.

Funding Requirements

Our primary uses of capital are, and we expect will continue to be, compensation and related expenses, third-party clinical research and development services, laboratory and related supplies, clinical costs, legal and other regulatory expenses and general overhead costs.

The successful development of our product candidates is highly uncertain. As such, at this time, we cannot reasonably estimate or know the nature, timing and estimated costs of the efforts that will be necessary to complete the remainder of the development of CLS-1001, CLS-1002, CLS-1003 or any of our other product candidates. We are also unable to predict when, if ever, material net cash inflows will commence from product sales. This is due to the numerous risks and uncertainties associated with developing drugs, including the uncertainty of:

- successful enrollment in, and completion of clinical trials;
- receipt of marketing approvals from applicable regulatory authorities;
- establishing commercial manufacturing capabilities or making arrangements with third-party manufacturers;
- obtaining and maintaining patent and trade secret protection and regulatory exclusivity for our product candidates; and
- launching commercial sales of the products, if and when approved, whether alone or in collaboration with others.

A change in the outcome of any of these variables with respect to the development of any of our product candidates would significantly change the costs and timing associated with the development of that candidate.

Until such time, if ever, as we can generate substantial product revenue, we expect to finance our cash needs through a combination of equity offerings, debt financings and potential collaboration, license and development agreements. We do not currently have any committed external source of funds. To the extent that we raise additional capital through the sale of equity or convertible debt securities, your ownership interest will be diluted, and the terms of these securities may include liquidation or other preferences that adversely affect your rights as a common stockholder. Debt financing and preferred equity financing, if available, may involve agreements that include covenants limiting or restricting our ability to take specific actions, such as incurring additional debt, making capital expenditures or declaring dividends.

If we raise additional funds through collaborations, strategic alliances or marketing, distribution or licensing arrangements with third parties, we may be required to relinquish valuable rights to our technologies, future revenue streams, research programs or product candidates or to grant licenses on terms that may not be favorable to us. If we are unable to raise additional funds through equity or debt financings when needed, we may be required to delay, limit, reduce or terminate our drug development or future commercialization efforts or grant rights to develop and market product candidates that we would otherwise prefer to develop and market ourselves.

[Table of Contents](#)

We will also incur costs as a public company that we have not previously incurred or have previously incurred at lower rates, including but not limited to, increased costs and expenses for fees to members of our board of directors, increased personnel costs, increased directors and officers insurance premiums, audit and legal fees, investor relations fees and expenses for compliance with reporting requirements under the Exchange Act and rules implemented by the SEC and NASDAQ. We estimate that we will annually incur approximately \$1.5 million to \$2.5 million in expenses to comply with these requirements.

Outlook

Based on our research and development plans and our timing expectations related to the progress of our programs, we expect that the net proceeds from this offering, together with our existing cash and cash equivalents, will enable us to fund our operating expenses and capital expenditure requirements for at least the next 12 months. We have based this estimate on assumptions that may prove to be wrong, and we could use our capital resources sooner than we expect. Additionally, the process of testing product candidates in clinical trials is costly, and the timing of progress in these trials is uncertain.

Cash Flows

During the six months ended June 30, 2013 and 2014, our operating activities used net cash of \$3.1 million and \$3.3 million, respectively. The use of net cash in each period primarily resulted from our net losses. The increase in net loss for the six months ended June 30, 2014 as compared to the six months ended June 30, 2013 was primarily attributable to higher research and development expenses. The other changes from operating activities were caused primarily by changes in our accounts payable, stock-based compensation, deferred offering costs and other accrued liabilities. During the six months ended June 30, 2013 and 2014, we did not engage in any material investing activities. The net cash provided by financing activities during the six months ended June 30, 2013 related to \$7.8 million from the sale of our Series A-1 convertible preferred stock, net of issuance costs, while the net cash provided by financing activities during the six months ended June 30, 2014 related primarily to \$3.0 million received from the issuance of the bridge notes.

During the years ended December 31, 2012 and 2013, our operating activities used net cash of \$3.2 million and \$6.8 million, respectively. The use of net cash in each year primarily resulted from our net losses. The increase in net loss for the year ended December 31, 2013 as compared to the year ended December 31, 2012 was due primarily to the increase in research and development expenses and, to a lesser extent, an increase in general and administrative expenses. The other changes from operating activities were caused primarily by changes in our accounts payable and other accrued liabilities. During the years ended December 31, 2012 and 2013, our investing activities included primarily purchases of office furnishings and equipment to facilitate our increased research and development activities and headcount. During the years ended December 31, 2012 and 2013, our financing activities provided net cash of \$4.1 million and \$8.0 million, respectively. The net cash provided by financing activities during the year ended December 31, 2012 was primarily the result of \$3.9 million in net proceeds from the sale of our Series A convertible preferred stock and \$150,000 from the issuance of debt, while net cash provided by financing activities during the year ended December 31, 2013 was the result of \$7.8 million from the sale of our Series A-1 convertible preferred stock as well as \$125,000 from the issuance of debt.

[Table of Contents](#)

Contractual Obligations

The following table summarizes our significant contractual obligations as of December 31, 2013, all of which consisted of obligations under leases for our corporate headquarters in Alpharetta, Georgia and our research facility in Durham, North Carolina.

	Payment due by period (in thousands)				
	Total	Less than 1 year	1-3 years	3-5 years	More than 5 years
Operating lease obligations	\$547	\$ 254	\$ 270	\$ 23	\$ —
Long-term debt obligations	275	—	125	150	—
Total	<u>\$822</u>	<u>\$ 254</u>	<u>\$ 395</u>	<u>\$ 173</u>	<u>\$ —</u>

We have no material non-cancelable purchase commitments with contract manufacturers or service providers, as we have generally contracted on a cancelable purchase order basis.

Off-Balance Sheet Arrangements

We did not have during the periods presented, and we do not currently have, any off-balance sheet arrangements, as defined under SEC rules.

Recent Accounting Pronouncements

We have considered all recently issued accounting pronouncements and do not believe the adoption of such pronouncements will have a material impact on our financial statements.

JOBS Act

In April 2012, the Jumpstart Our Business Startups Act of 2012, or the JOBS Act, was enacted. Section 107(b) of the JOBS Act provides that an emerging growth company can take advantage of an extended transition period for complying with new or revised accounting standards. Thus, an emerging growth company can delay the adoption of certain accounting standards until those standards would otherwise apply to private companies. We have irrevocably elected not to avail ourselves of this extended transition period, and, as a result, we will adopt new or revised accounting standards on the relevant dates on which adoption of such standards is required for other public companies.

Quantitative and Qualitative Disclosures about Market Risk

The market risk inherent in our financial instruments and in our financial position represents the potential loss arising from adverse changes in interest rates. As of December 31, 2012 and 2013 and June 30, 2014, we had cash and cash equivalents of \$0.9 million, \$1.9 million and \$1.5 million, respectively. We generally hold our cash in interest-bearing money market accounts. Our primary exposure to market risk is interest rate sensitivity, which is affected by changes in the general level of U.S. interest rates. Due to the short-term maturities of our cash equivalents and the low risk profile of our investments, an immediate 100 basis point change in interest rates would not have a material effect on the fair market value of our cash equivalents.

We do not engage in any hedging activities against changes in interest rates. Our outstanding debt instruments carry a fixed interest rate and, as such, are not subject to interest rate risk.

We do not have any foreign currency or other derivative financial instruments.

BUSINESS

Overview

We are a clinical-stage biopharmaceutical company developing first-in-class drug therapies to treat chronic, blinding diseases of the eye. Our current product candidates focus on diseases affecting the retina and choroid, especially diseases associated with macular edema, and are injected non-surgically into the suprachoroidal space, or SCS, adjacent to the choroid, using our proprietary microinjector. With the SCS injection procedure, our product candidates are more directly administered to the retina and choroid as compared to currently used intravitreal injections, which we believe may provide a number of benefits, including lower frequency of necessary administration and faster onset of therapeutic effect. We hold the exclusive rights to develop and commercialize drugs for non-surgical administration into the SCS. Our most advanced product candidates, CLS-1001 and CLS-1003, are based on commonly used ophthalmic therapeutic agents, which we believe will allow us to more efficiently and predictably pursue the regulatory approval of these product candidates under Section 505(b)(2) of the Federal Food, Drug, and Cosmetic Act. Under Section 505(b)(2), if we can establish that reliance on the FDA's previous findings of safety or effectiveness or both for an approved product is scientifically appropriate, or there is scientific literature to support the product's safety or effectiveness, it may eliminate the need for us to conduct some of the preclinical studies or clinical trials for the new product candidate that might otherwise have been required. CLS-1001, for macular edema associated with non-infectious uveitis, is in an ongoing Phase 1/2 clinical trial, from which we expect final results in the first half of 2015. We expect to be able to conduct a single pivotal Phase 3 clinical trial in approximately 150 patients beginning in the first half of 2015, the results of which we believe, based on a meeting with the FDA and subsequent written correspondence, will be sufficient to support the filing of a New Drug Application, or NDA, to the FDA in the first half of 2017. For CLS-1003, for macular edema associated with retinal vein occlusion, or RVO, we plan to initiate a Phase 2 clinical trial before the end of 2014, with data expected in the second half of 2015. In our CLS-1002 program, for wet age-related macular degeneration, or wet AMD, we are evaluating drug compounds with anti-VEGF activity or dual anti-VEGF and anti-PDGF activity for SCS injection, and plan to file an investigational new drug application, or IND, by the end of 2015. We are also considering a development program for drug compounds that may be able to treat diabetic macular edema, or DME. We estimate that there are nearly five million people in the United States diagnosed with our target indications and that worldwide annual sales of drugs to treat these indications were approximately \$6 billion in 2013.

Our current and planned development programs aim to restore or improve visual function primarily by reducing the macular edema associated with a number of diseases affecting the retina, the tissue that lines the inside of the eye and is the part of the eye primarily responsible for vision, and the choroid, the layer adjacent to the retina that supplies the retina with blood, oxygen and nourishment. These diseases include uveitis, RVO, wet AMD and DME. Macular edema is the build-up of fluid that can cause abnormal swelling of the macula, the portion of the retina responsible for central vision and color perception. This swelling can rapidly result in deterioration of vision and can eventually lead to blindness.

The most common treatments for diseases of the retina and choroid are corticosteroids and drugs that inhibit vascular endothelial growth factor, or anti-VEGF drugs. These corticosteroids and anti-VEGF drugs are injected into the vitreous, a jelly-like substance that occupies the central portion of the eye. Intravitreal injections rely on drug to diffuse outward from the vitreous to the retina and choroid, which diffusion can result in lower concentrations of drug in these targeted areas. At the same time, this diffusion can result in drug spreading to unintended parts of the eye, where, for some drugs, it can cause significant side effects.

[Table of Contents](#)

By contrast, with our approach, drug is injected into and spreads within the SCS, which is the space located between the choroid and the outer protective layer of the eye known as the sclera, to reach the back of the eye. We believe SCS administration may provide improved bioavailability to the targeted retina and choroid, with drug remaining more localized and away from other parts of the eye, where the drug could cause side effects. Corticosteroids are effective both at treating the inflammatory aspect of ocular disease and at reducing macular edema, but when delivered intravitreally, they have been associated with significant side effects, such as cataracts and elevated intraocular pressure, or IOP, which can lead to glaucoma. Anti-VEGF drugs, while highly effective in reducing macular edema, require frequent administration, in some cases as often as monthly. We believe that our differentiated approach of injecting ocular drugs into the SCS may result in fewer side effects and improved bioavailability of drug at the diseased retina and choroid, as compared to intravitreal administration, a faster onset of therapeutic effect, a similar efficacy profile with lower drug amounts required, and longer duration of therapeutic effect with less frequent required injections. The procedure for SCS injection with our microinjector is conducted in an in-office setting and is similar to the procedure for intravitreal injection, and we therefore expect our products, if they achieve regulatory approval, to be incorporated into retinal specialists' standard medical practice.

Under our CLS-1001 program, we are conducting a Phase 1/2 clinical trial in patients with non-infectious uveitis, a condition characterized by inflammation of the choroid and surrounding tissues. We intend to seek orphan drug designation and exclusivity for CLS-1001 for this indication. The most common treatment for non-infectious uveitis involves the use of corticosteroids, such as triamcinolone acetonide, or TA. CLS-1001 consists of an SCS injection of CLS-TA, our proprietary, preservative-free formulation of TA, specifically designed to be administered through our SCS microinjector. In this ongoing Phase 1/2 clinical trial, for which we have filed an IND with the FDA, we are evaluating the safety of SCS injection of Triesence, a TA formulation that is similar to CLS-TA and that has been approved by the U.S. Food and Drug Administration, or FDA, to treat non-infectious uveitis. While the trial is primarily a safety and tolerability study, we are also assessing efficacy measures. We intended to enroll approximately 10 patients in this trial. We completed enrollment after dosing the eighth patient. Interim data from the eight patients for whom data is available indicate that SCS injection of the drug has generally been well tolerated, with none of the eight patients having developed cataracts or experienced elevated IOP. At eight weeks after a single SCS injection, the visual acuity of all seven patients who currently have reached week 8 had improved by five or more letters on a standard eye chart over baseline, a clinically meaningful improvement, and the visual acuity of five of these seven patients improved by at least 15 letters. Because CLS-1001 is based on our formulation of TA, and TA is known to be effective in treating uveitis, we believe that our product candidate will be at least as effective as commonly used formulations of TA in treating the disease, including the associated macular edema. Our preclinical studies suggest that SCS injection of TA may provide improved bioavailability of drug at the diseased retina and choroid, which may result in faster onset of therapeutic effect and a similar efficacy profile with lower drug amounts required. Our early clinical trial results suggest the potential for CLS-1001 to effectively treat uveitis for at least 90 days following a single SCS injection. Currently used ocular injections of TA are typically effective for approximately 90 days. In the second half of 2014, we plan to initiate a Phase 2 clinical trial in approximately 30 patients with macular edema associated with non-infectious uveitis to evaluate the safety and efficacy of our CLS-TA administered into the SCS. In the first half of 2015, we also expect to initiate a single pivotal Phase 3 clinical trial with our CLS-TA administered into the SCS in approximately 150 patients with macular edema associated with non-infectious uveitis, with the goal of being able to submit an NDA to the FDA in the first half of 2017. Based on our discussions with the FDA, which included both a

[Table of Contents](#)

pre-IND meeting in 2012 and subsequent written correspondence in 2013, we believe this single pivotal clinical trial will be sufficient to support a 505(b)(2) NDA filing for CLS-1001 for macular edema associated with non-infectious uveitis. If approved, CLS-1001 would be the first drug specifically indicated for macular edema associated with non-infectious uveitis.

Under our CLS-1003 program, we are planning to initiate a Phase 2 clinical trial before the end of 2014 in approximately 40 patients with macular edema associated with RVO, a sight-threatening disorder resulting from the blockage of a retinal vein. We have not yet discussed our clinical development plans for this program with the FDA, however. The current standard of care for RVO involves monthly intravitreal injections of an anti-VEGF drug, such as Lucentis (ranibizumab), Eylea (aflibercept) or Avastin (bevacizumab). Although studies have shown that corticosteroids effectively address the inflammatory aspect of RVO, they are not used as first-line therapy because they are generally not considered to be as effective as anti-VEGF drugs and also have an unattractive side effect profile when injected intravitreally. In our Phase 2 clinical trial, we will evaluate the safety and efficacy of the injection of our CLS-TA into the SCS together with Eylea injected into the vitreous, as compared to an intravitreal injection of Eylea alone. We believe that SCS administration of our CLS-TA, by addressing the inflammatory aspect of RVO, in combination with intravitreal anti-VEGF treatment, addressing the vascular aspect of RVO, may result in faster onset of therapeutic effect compared to currently used intravitreal anti-VEGF treatment alone, while also potentially reducing the frequency of necessary anti-VEGF treatments from once every 30 days to once every 90 days. In addition, we believe that the side effect profile of CLS-1003 will be similar to that of current intravitreal anti-VEGF treatments alone, as SCS injections of CLS-TA should allow the corticosteroid to remain substantially localized in the retina and choroid without introducing the side effects that are commonly observed with other routes of administration of corticosteroids. Our planned clinical trials will evaluate this side effect profile.

Under our CLS-1002 program, we are evaluating a number of compounds as potential therapies for the treatment of wet AMD by SCS injection, including compounds with anti-VEGF activity and compounds with dual anti-VEGF and anti-platelet-derived growth factor, or anti-PDGF, activity. Wet AMD is a condition involving the abnormal formation and growth of new blood vessels in the choroid, their intrusion into the outer layers of the retina and the leakage of blood and fluid into the retina. The current standard of care for the treatment of wet AMD is intravitreal injection of anti-VEGF drugs as a standalone therapy, as often as seven times per year. We have completed a Phase 1 clinical trial evaluating the safety and efficacy of SCS injection of Avastin, an anti-VEGF drug, in four patients with wet AMD. In this trial, SCS injection of Avastin was observed to be well tolerated, with no treatment-related serious adverse events and no unexpected adverse events. In addition, the four patients showed an average improvement in visual acuity of nine letters on a standard eye chart at the end of the eight-week post-treatment observation period. Two of the four patients showed an improvement in visual acuity of at least 13 letters. Because wet AMD manifests in the retina and choroid, we believe that SCS injection of drug more directly to the retina and choroid may result in faster onset of therapeutic effect as compared to intravitreal injection, and could reduce the necessary frequency of treatment to once every 90 days. We plan to further study this effect in any future clinical trials that we conduct as part of our CLS-1002 program.

Our drug candidates, microinjector, and method of non-surgical drug delivery into the SCS are protected by four issued U.S. patents and over 20 patent applications filed in the United States and major international markets, including the European Union, Canada, India, Japan, China and Australia. Our issued patents are broadly directed to methods of administering drugs of any type into the SCS by injection and are not scheduled to expire until between 2027 and 2029. Our patent applications relate to SCS delivery

[Table of Contents](#)

technology and methods, our current therapeutic drug candidates and formulations, various therapeutic uses, including treatment of specific indications such as macular edema associated with non-infectious uveitis and RVO, as well as wet AMD and DME, and administration of specific classes of drugs to the SCS, including anti-inflammatory drugs and anti-VEGF drugs. Our pending patent applications are not projected to expire until between 2027 and 2034.

If our product candidates are approved, we plan to commercialize them with a specialty sales force of 30 to 40 representatives to target retinal specialists in the United States, and we may also pursue collaborations with third parties to commercialize any of our drugs approved for marketing outside the United States.

The members of our management team have held senior leadership positions at a number of pharmaceutical and biotechnology companies with approved ophthalmic drugs, including Alcon, Alimera Sciences, CIBA Vision and ISTA Pharmaceuticals. Members of our team have contributed to the development and commercialization of several retinal products, such as Triesence, Iluvien, Nevanac, Visudyne and Xibrom.

Strategy

Our goal is to be a leading biopharmaceutical company focused on the development and commercialization of therapeutics to treat chronic, blinding diseases of the eye, with a particular emphasis on diseases affecting the choroid and the retina. The key elements of our strategy are:

- **Advancing CLS-1001 and CLS-1003 to FDA approval under the Section 505(b)(2) NDA regulatory pathway.** Our most advanced product candidates, CLS-1001 and CLS-1003, utilize CLS-TA, our proprietary formulation of TA, a corticosteroid that has been used for decades to treat ocular inflammatory conditions. We believe this approach will allow us to more efficiently and predictably pursue the regulatory approval of these product candidates under Section 505(b)(2) of the Federal Food, Drug and Cosmetic Act. We plan to advance CLS-1001 into a single pivotal Phase 3 clinical trial for macular edema associated with non-infectious uveitis and, based on our pre-IND meeting and subsequent written correspondence with the FDA, we expect this single trial to be sufficient to support a 505(b)(2) NDA filing. Our goal is to submit the 505(b)(2) NDA for CLS-1001 in the first half of 2017. Separately, we expect to initiate a Phase 2 clinical trial for CLS-1003 for macular edema associated with RVO before the end of 2014, with data expected in the second half of 2015.
- **Maximizing the commercial potential of our product candidates.** If either CLS-1001 or CLS-1003 is approved, we plan to build a specialized sales force of approximately 30 to 40 representatives to target the approximately 1,700 retinal specialists in the United States. We expect that this sales force would be able to effectively sell more than one of our products, in the event that multiple product candidates are approved for marketing. We have retained the worldwide commercialization rights to all of our product candidates, except we have granted a third party the exclusive right to commercialize our product candidates in Russia and specified adjacent territories pursuant to a license agreement, as described later in this prospectus. We may pursue additional collaborations with third parties to commercialize any of our drugs approved for marketing outside the United States, either through territorial licenses or distributor relationships.

[Table of Contents](#)

- **Developing therapies for other back of the eye diseases that can be treated more effectively with SCS injection.** We believe that SCS injection of ocular therapies could have benefits in a variety of other back of the eye indications. Our initial areas of focus for development include:
 - **Advancing our CLS-1002 wet AMD development program.** We are investigating opportunities to improve the treatment of wet AMD through SCS injection of compounds that are commonly used for this disease, such as anti-VEGF drugs, or have shown clinical promise, such as anti-VEGF drugs in combination with anti-PDGF drugs. These therapies are commonly administered by intravitreal injection. We have completed an exploratory Phase 1 clinical trial using an anti-VEGF drug, Avastin, administered into the SCS, rather than intravitreally, in four patients with wet AMD, from which we observed encouraging safety and efficacy results. We are conducting preclinical studies evaluating other compounds with dual anti-VEGF and anti-PDGF activity, with the goal of selecting a lead drug candidate under this program for IND submission in 2015.
 - **Developing a product candidate to treat DME.** DME, like uveitis and RVO, is characterized by an inflammatory aspect. Once the data from the CLS-1003 Phase 2 clinical trial have been analyzed, we intend to initiate clinical trials to develop a product candidate to treat DME modeled after our approach for addressing macular edema associated with RVO.
 - **Developing additional therapies through collaborations with third parties.** We plan to explore collaborations with third parties to develop SCS-administered treatments for back of the eye diseases. These collaborations may take the form of in-licenses of the third parties' drugs or drug candidates for our own development for SCS injection, or out-licenses for third parties to use our intellectual property covering SCS injection as part of the development of their own drugs. We are currently studying several compounds under a research collaboration with Santen Pharmaceutical Co., Ltd.
- **Maintaining and strengthening our intellectual property portfolio.** Our intellectual property strategy aims to control the commercialization of SCS access, with patents and patent applications covering non-surgical SCS delivery methods, novel formulations of drugs, and microinjectors used to access the SCS. Our current patents and patent applications provide us with exclusivity in the United States and major international markets, and are not projected to expire until between 2027 and 2034. We will seek to broaden the scope of and increase the geographic reach of our patent protection throughout the world.

The Clearside Approach

We are developing drug candidates for back of the eye diseases to be administered to the retina and choroid through the SCS using our proprietary microinjector. We believe that our novel, patented approach of SCS drug administration may provide a number of potential benefits, including:

- **Improved bioavailability at the site of disease and faster onset of therapeutic effect.** In preclinical studies, we observed higher amounts of drug present in the retina and choroid following SCS injection, as compared to the distribution of the same drug following intravitreal injection. We believe this suggests that SCS injection may have similar or better efficacy with a faster onset of therapeutic effect than intravitreal injection in diseases of the retina and choroid like uveitis, RVO, wet AMD and DME.

- **Similar efficacy profile with lower drug amounts required.** In a preclinical study in a model of uveitis, SCS injection of TA achieved similar efficacy results with only 10% of the drug amount commonly administered through intravitreal injection. We believe this suggests our approach may allow us to develop ocular therapies with similar efficacy, but with lower amounts of active drug, compared to current therapies.
- **Less frequent injections.** Current treatments for RVO require monthly intravitreal injections of anti-VEGF drugs, while current treatments for wet AMD require such injections as often as seven times per year. For RVO patients, we believe that a combination of a standard intravitreal injection of an anti-VEGF drug that addresses the vascular aspect of the disease, together with an SCS injection of CLS-TA, which addresses the inflammatory aspect of RVO, may have similar or better efficacy with a reduction in the frequency of required anti-VEGF treatments from once every 30 days to once every 90 days. In wet AMD, we believe that more direct application of drugs with anti-VEGF or dual anti-VEGF and anti-PDGF activity to the neovascularization in the choroid through SCS injection may block the process of additional new vascular growth within the choroid before the vessels break into the retina and create further damage through leakage. We believe that such SCS injection of treatment therefore has the potential to delay vision loss for a longer duration of time than the current standard intravitreal administration of anti-VEGF drugs and, accordingly, could require less frequent administration.
- **Enhanced safety profile.** Intravitreal injections result in drugs diffusing throughout the eye, including into the lens, iris and ciliary body at the front of the eye, which for some drugs, has been associated with safety issues, such as cataracts and elevated IOP levels. Intravitreal administration of TA has been associated with cataracts and increases in IOP levels in 20% to 60% of patients. Because SCS injection of drugs appears to result in drug remaining localized in the retina and choroid without substantial diffusion to the vitreous or the front portion of the eye, we believe SCS injection has the potential to reduce the incidence of these side effects. Based on interim results from our ongoing Phase 1/2 clinical trial, none of the eight patients for whom data is available have experienced cataracts, abnormally high levels of IOP or other treatment-related serious adverse events over the course of 26 weeks following a single SCS injection of TA.
- **Incorporated into standard medical practice.** If approved for marketing, our drugs will be packaged together with our SCS microinjector for one-time use by retinal specialists in their offices. Our product candidates are designed to be administered using a similar procedure as is used for intravitreal injections, and no capital equipment will be required. Accordingly, we expect our products, if approved, will be easily incorporated into retinal specialists' standard medical practice.

[Table of Contents](#)

The current development status of our product candidates is summarized in the chart below:

Development Program	Indication	Route of Administration and Active Ingredient	Status and Upcoming Milestones
CLS-1001	Macular edema associated with non-infectious uveitis	SCS injection of CLS-TA	<ul style="list-style-type: none">• Final results from ongoing Phase 1/2 clinical trial with Trience expected 1H 2015• Initiate ~30-patient Phase 2 clinical trial in 2H 2014 with CLS-TA, with data expected 1H 2015• Initiate ~150-patient single pivotal Phase 3 clinical trial in 1H 2015 with CLS-TA, with 505(b)(2) NDA filing expected 1H 2017
CLS-1003	Macular edema associated with RVO	SCS injection of CLS-TA together with intravitreal injection of anti-VEGF compound	<ul style="list-style-type: none">• Initiate ~40-patient Phase 2 clinical trial before end of 2014, with data expected 2H 2015
CLS-1002	Wet AMD	SCS injection of anti-VEGF compound or compound with dual anti-VEGF and anti-PDGF activity	<ul style="list-style-type: none">• Exploratory Phase 1 clinical trial with Avastin completed outside the U.S.• Selection of lead drug candidate for IND submission expected in 2015

We have discussed our proposed clinical development program with the FDA for CLS-1001, but have not yet done so for our planned CLS-1003 and CLS-1002 development programs.

CLS-1001 Program Targeting Macular Edema Associated with Non-infectious Uveitis

We are developing CLS-1001 for treatment of macular edema associated with non-infectious uveitis, a condition characterized by inflammation of the choroid and surrounding tissues. CLS-1001 consists of an SCS injection of CLS-TA, our proprietary formulation of TA specifically designed to be administered through our SCS microinjector. We expect final results from our ongoing Phase 1/2 clinical trial in the first half of 2015, plan to initiate a Phase 2 clinical trial with CLS-TA in approximately 30 patients in the second half of 2014, with expected data in the first half of 2015, and plan to initiate a single pivotal Phase 3 clinical trial in approximately 150 patients in the first half of 2015, with a Section 505(b)(2) NDA filing expected in the first half of 2017. Based on our discussions with the FDA, which included both a pre-IND meeting as well as subsequent written correspondence, we believe we will be able to initiate our single pivotal Phase 3 clinical trial of CLS-1001 before we have the final results from our planned Phase 2 clinical trial. We believe that CLS-1001 will be at least as effective in treating uveitis, including the associated macular edema, as commonly used treatments with corticosteroids, but has the potential to provide improved bioavailability of drug at the diseased retina and choroid, which may result in faster onset of therapeutic effect, a similar efficacy profile with lower drug amounts required, and longer duration of therapeutic effect, potentially resulting in a reduced frequency of necessary injections. We also believe that CLS-1001 may result in fewer side effects compared to commonly used corticosteroid treatments, and we are testing this potential side effect profile in clinical trials.

Market Opportunity for Treatment of Macular Edema Associated with Non-infectious Uveitis

Uveitis is one of the most frequent causes of blindness in the developed world. Based on prevalence data published in the journal *Ophthalmology* in 2004 and United States census data for 2010, we estimate approximately 350,000 individuals in the United States suffer from some form of uveitis. Typically diagnosed in individuals between the ages of 20 and 50, uveitis can occur in one or both eyes and accounts for approximately 10% of cases of blindness in the United States, according to a study published in *Journal of Ophthalmology*. Uveitis can be either infectious or non-infectious. Non-infectious uveitis accounts for approximately 80% of all uveitis cases. Macular edema related to uveitis is the predominant cause of blindness or visual impairment among patients with uveitis, accounting for approximately 30% of cases of blindness in uveitis patients. Because uveitis can become chronic or recurrent if not adequately treated, some patients may become refractory, or unresponsive, to treatment, leading to irreversible blindness.

Limitations of Currently Available Therapies for Macular Edema Associated with Non-Infectious Uveitis

Corticosteroids are the most commonly used treatments for non-infectious uveitis. Corticosteroids administered through eye drops, although used to treat some uveitis patients, are generally ineffective in treating patients with macular edema associated with non-infectious uveitis. Corticosteroids administered orally or systemically can be effective in treating patients with macular edema associated with non-infectious uveitis, but their long-term use is associated with significant side effects. As a result of the shortcomings associated with these methods of administering corticosteroids, the most common method for treating macular edema associated with non-infectious uveitis is intravitreal injection or implant of a corticosteroid, such as TA, dexamethasone or fluocinolone acetonide. Examples of intravitreal corticosteroid treatments include Ozurdex®, Retisert®, Kenalog® and Trience. Ozurdex is a biodegradable dexamethasone implant that has been approved by the FDA as a treatment for non-infectious uveitis, but has been associated with increased IOP in 25% of patients, conjunctival hemorrhages in 22% of patients and cataracts in 5% of patients. Retisert, a non-biodegradable fluocinolone acetonide implant that requires surgical administration, has also been approved by the FDA as a treatment of non-infectious uveitis, but more than 75% of patients receiving Retisert require additional medications within three years of receiving Retisert therapy to effectively treat their uveitis, more than 35% of patients require surgical intervention in order to control increase in IOP levels, and nearly all patients develop cataracts and require cataract surgery following treatment.

Trience and Kenalog are the main injectable formulations of TA that are used for the treatment of intraocular inflammatory conditions. Prescription of Kenalog is off-label because it has not been approved by the FDA to treat intraocular conditions. TA has been associated with increases in IOP or cataract progression in 20% to 60% of patients when injected intravitreally.

Immunosuppressive agents are also sometimes used to control the inflammation associated with non-infectious uveitis, but due to their ability to systemically impair the body's immune system and their association with additional side effects, physicians are cautious in using these agents. Anti-VEGF therapies are also used to treat macular edema in a variety of diseases, but are generally not used in uveitis patients because they do not treat the inflammation inherent in uveitis or most of the other complications associated with uveitis.

Potential Benefits of CLS-1001

If approved by the FDA, CLS-1001 would be the first treatment specifically indicated for macular edema associated with non-infectious uveitis. Because CLS-1001 is based on our CLS-TA formulation of TA, a

corticosteroid known to be effective in treating uveitis, we believe that our product candidate will be at least as effective as TA in treating the disease, including the associated macular edema. Our clinical development program for CLS-1001 has been designed to test this hypothesis. Our preclinical studies suggest that SCS injection of TA may provide improved bioavailability of drug at the retina and choroid, which could result in faster onset of therapeutic effect and a similar efficacy profile with lower drug amounts required. Our early clinical trial results suggest the potential for CLS-1001 to be effective at treating uveitis for at least 90 days following a single SCS injection. Currently used ocular injections of TA are typically effective for approximately 90 days. In contrast to intravitreal injections, based on our preclinical studies, we believe that SCS injection will allow the drug to remain more localized in the diseased retina and choroid. As a result of this localization, we believe that CLS-1001 may be effective in treating non-infectious uveitis, including the associated macular edema, without significant drug exposure to other eye tissues, thereby potentially reducing the incidence of side effects associated with commonly used corticosteroid treatments. We will evaluate this result in our future clinical trials for CLS-1001.

Initial Clinical Research on Surgical Administration of TA into the SCS

In 2011, two separate studies were published by researchers evaluating administration of TA together with the anti-VEGF drug Avastin in a total of 27 patients with severe retinal disease. In one study, six patients had RVO or DME, and in the other study, 21 patients had wet AMD. In both studies, a surgical procedure was used to cut open the eye to expose the choroidal tissue and insert a small tube into the SCS, through which both TA and Avastin were administered. The six RVO and DME patients were followed for one year and the 21 wet AMD patients were followed for six months. Of the 27 patients, only one patient had developed cataracts, and one other patient was in the initial stages of cataract formation at the end of the observation period. In addition, only one patient showed elevated IOP, which was controlled with a topical glaucoma medication. We believe these studies suggest that surgical administration of TA into the SCS might present a lower risk of increased IOP and cataracts than intravitreal injection of TA, which has been associated with increases in IOP or cataract progression in 20% to 60% of patients.

Our Clinical and Preclinical Development of CLS-1001 for Non-Surgical Administration

Based on our consultation with the FDA, we have conducted or are conducting the following clinical trial and preclinical studies, in each case using TA injected non-surgically using a prototype of our microinjector, as part of our CLS-1001 development program:

- an ongoing Phase 1/2 clinical trial in non-infectious uveitis patients evaluating SCS injections of Triesence, a commercially available TA formulation, with our microinjector, from which we expect final safety and efficacy data in the first half of 2015;
- a completed preclinical study comparing the pharmacokinetic effects of SCS and intravitreal injection of Triesence in rabbits;
- a completed preclinical study comparing the pharmacokinetic effects of SCS injection of CLS-TA and Triesence in rabbits;
- a completed preclinical study evaluating the pharmacodynamic effects of SCS and intravitreal injection of Triesence in a pig model of uveitis; and
- completed preclinical studies evaluating the toxicology of SCS injection of CLS-TA and Triesence in rabbits.

Details of this trial and these studies are summarized below.

Ongoing Phase 1/2 Clinical Trial

In July 2013, we initiated a Phase 1/2 multi-center, open-label clinical trial designed to evaluate the safety and tolerability of a single injection of Triesence into the SCS in patients diagnosed with non-infectious uveitis. The trial is being conducted under an IND we submitted to the FDA in December 2012 for the potential treatment of sympathetic ophthalmia, temporal arteritis, uveitis and ocular inflammatory conditions unresponsive to topical corticosteroids. The IND was amended in March 2014 to add macular edema following non-infectious uveitis as a treatment indication. The primary purpose of this trial is to demonstrate the overall safety of treating uveitis patients by administering a commercially available formulation of TA into the SCS, rather than intravitreally. We expect to use CLS-TA for all future clinical trials intended to support a 505(b)(2) NDA submission for CLS-1001.

Clinical Trial Design. We plan to enroll approximately 10 patients at three centers in this trial. Eight patients have been dosed in the trial to date. Eligibility criteria include adult patients with non-infectious uveitis experiencing either macular edema or vitreous haze, another common complication of uveitis. We are enrolling patients with either of these complications of uveitis in order to assess the safety and tolerability of SCS injection of TA and to determine whether SCS injection of TA can improve patient vision by reducing the effects of either of these conditions. For inclusion in the trial, patients must have IOP of no greater than 22 millimeters of mercury, or mmHg, a common measure of pressure. Normal IOP levels are generally between 10 and 12 mmHg at the low end and 20 and 22 mmHg at the high end. IOP levels above 22 mmHg are considered to be elevated, increasing the patient's risk of developing glaucoma.

Each patient enrolled receives a single SCS microinjection of 4.0 mg of Triesence, a common intravitreal dose of TA, at a location at the front of the eye similar to the location typically used for intravitreal injection. Patients return for a follow-up examination on the day after the injection and then for eight additional evaluations at weeks 1, 2, 4, 8, 12, 16, 20 and 26 following the treatment. Patients may receive other treatment for non-infectious uveitis at any time during the trial with any accepted therapy, if the patient's condition deteriorates or if the treating physician otherwise determines it to be advisable. In the event a patient receives other treatment, we continue to follow the patient for the duration of the trial for safety purposes, but we thereafter no longer evaluate efficacy measures.

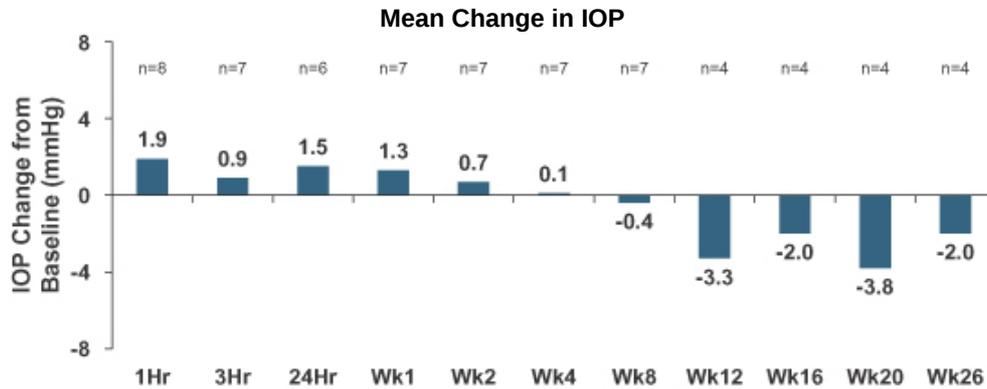
This Phase 1/2 clinical trial is not powered to show efficacy results with statistical significance. Statistical significance means that an effect is unlikely to have occurred by chance. Clinical trial results are considered statistically significant when the probability of the results occurring by chance, rather than from the efficacy of the product candidate, is sufficiently low. Since the trial is not powered to show results with statistical significance, the results from the trial may be attributable to chance and not the clinical efficacy of TA. This trial design is typical of Phase 1 and some Phase 2 clinical trials, the principal purpose of which is to provide the basis for the design of larger, definitive trials with the inclusion of more patients to show statistical significance.

Endpoints. Our Phase 1/2 clinical trial is primarily a safety and tolerability study, although we are also assessing efficacy measures. The main safety endpoint relates to changes from baseline in IOP. We are also assessing efficacy endpoints relating to changes in best-corrected visual acuity, or BCVA, and changes in retinal thickness, which is a common measurement of macular edema. BCVA is a common measurement of a patient's ability to see at distances and is measured as the number of letters that the patient can read on the Early Treatment Diabetic Retinopathy Study, or ETDRS, eye chart. The ETDRS eye chart is a well-established standardized method of testing visual acuity.

Table of Contents

Interim Safety Results. The chart below shows the mean change in IOP for the eight patients treated in the trial for whom initial data is available, as measured at different time points post-treatment. Four of these eight patients have completed the full 26-week observation period. No patient has experienced a clinically meaningful increase in IOP, defined as a 10 mmHg increase in IOP from baseline, or an absolute increase in IOP to a level above 25 mmHg, which symptoms are typically seen at four to 12 weeks after administration. In contrast, the prescribing information for Ozurdex notes that in clinical trials of over 300 patients, 28% of patients receiving Ozurdex experienced at least a 10 mmHg increase in IOP from baseline at a follow-up visit, and 33% of patients treated with Ozurdex experienced an increase in IOP to a level above 25 mmHg. Additionally, no patient has required medication to lower increased IOP during the course of the trial.

The number of patients included in the results for each time point in the chart below, indicated with the notation “n=”, varies because, at some time points up to week 4, one or two patients either missed a follow-up visit or a measurement was inadvertently skipped, resulting in only six or seven of the eight patients being measured.



In addition to these IOP observations, the drug has been generally well tolerated in the trial to date. One patient with a history of pulmonary embolisms was hospitalized following a pulmonary embolism 10 weeks after treatment. The principal investigator considered this serious adverse event to be unrelated to the treatment, an assessment that was reviewed and approved by an independent medical monitor. The condition resolved after three days.

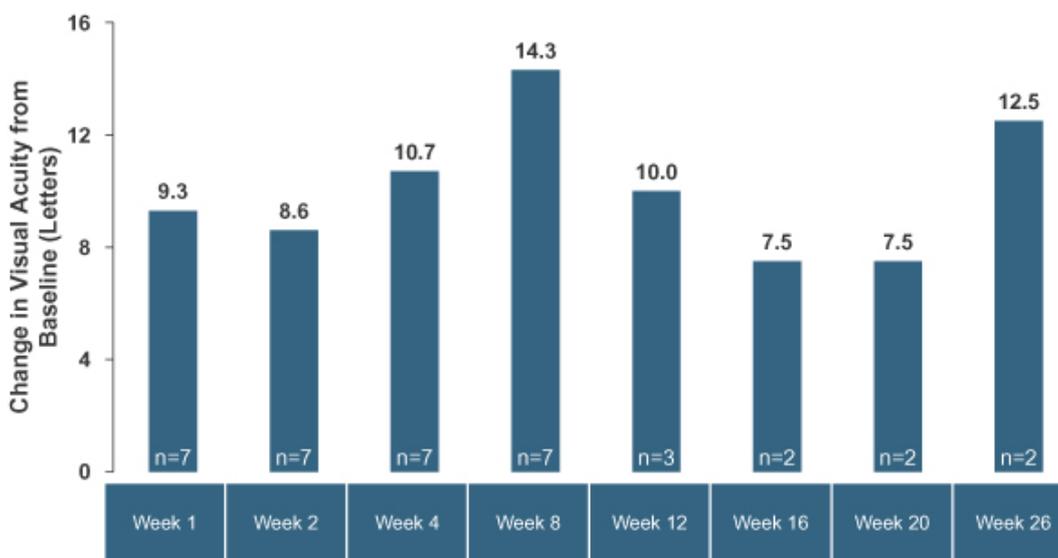
Interim Efficacy Results — Visual Acuity. Data are available at week 8 with respect to BCVA for seven of the eight patients who have received treatment. At each follow-up examination, the change in BCVA was measured as the difference from the patient’s baseline in the number of letters read on the ETDRS eye chart. Of the seven patients for whom data is available at or beyond week 8, two patients have completed the entire 26-week observation period without receiving other treatment for non-infectious uveitis. One patient received other treatment for non-infectious uveitis at week 8 and one patient received other treatment for non-infectious uveitis at week 16.

The chart below summarizes the mean improvement in BCVA observed to date in the trial for the eight patients for whom data is available. At eight weeks after a single SCS injection of Triesence, the visual acuity of all seven patients who currently have reached week 8 had improved by five or more letters on a

standard eye chart over baseline, a clinically meaningful improvement. Five of the seven patients who have reached week 8, or approximately 70% of these patients, had improved BCVA by at least 15 letters at week 8, compared to Ozurdex, for which approximately 40% of patients had improved BCVA by at least 15 letters at week 8.

In addition to the number of patients varying at each time point as a result of missing follow-up visits or inadvertently skipped measurements, as described above under “—Interim Safety Results,” the number of patients included in the results for the time points in the chart below also varies because efficacy observations for the two patients who received other treatment for their non-infectious uveitis during the follow-up observation period of the trial are only included up to the point that these patients received such other treatment.

Improvement in Best Corrected Visual Acuity (Number of Letters Read on ETDRS Chart)



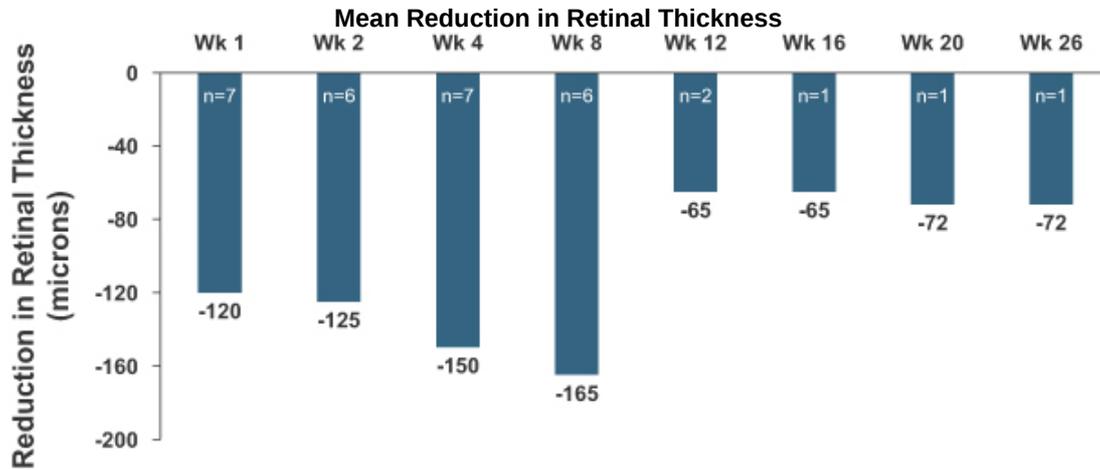
We believe these interim safety results are generally comparable to BCVA improvement observed in clinical trials of Ozurdex, in which patients with non-infectious uveitis achieved an average improvement of 13 letters at week 8 after treatment and 10 letters at week 26.

Interim Efficacy Results — Retinal Thickness. Of the eight patients treated in the trial, seven were experiencing macular edema at the time of treatment and were therefore also evaluated for change in retinal thickness after the injection. A reduction in retinal thickness in patients with macular edema occurs with the removal of excess fluid from the retina, reflecting a reduction of the swelling of the macula and other parts of the retina, affected by the edema. The eighth patient was experiencing vitreous haze associated with uveitis, an inflammatory symptom of uveitis where inflammatory cells cloud the vitreous, but in this case, the patient did not have macular edema, and change in retinal thickness is therefore not a relevant measurement for this patient.

The chart below summarizes the mean change in retinal thickness observed to date in the trial in the seven evaluated patients. Data are available at week 8 with respect to retinal thickness for six of the seven

evaluated patients. Each of these six patients achieved meaningful reductions in retinal thickness of approximately 50 microns from their respective baselines by week 8 following treatment. In addition, by week 8, four of these six evaluated patients experienced a reduction in retinal thickness to near or below 310 microns, which represents the maximum retinal thickness for 95% of the population with normal retinas. One micron is equal to one-thousandth of one millimeter. The reduction in retinal thickness experienced by one of the other two patients was over 20% from the patient’s baseline. Both of these measures are considered clinically meaningful improvements. For the three patients experiencing macular edema who have completed the study, their levels of reduction were generally sustained throughout the remainder of their measurement periods of 8, 12 and 26 weeks, respectively.

In addition to the number of patients varying at each time point as a result of missing follow-up visits or skipped measurements, as described above under “—Interim Safety Results,” of the seven patients that were experiencing macular edema, the number of patients included in the results for the time points in the chart below also varies because efficacy observations for the two patients who received other treatment for their non-infectious uveitis during the follow-up observation period of the trial are only included up to the point that these patients received such other treatment.



We believe these interim results are encouraging because we observed a rapid and persistent effect in the back of the eye after the SCS injection procedure at a location at the front of the eye similar to the location typically used for intravitreal injection. It is important to note that the improvements in BCVA and reductions in retinal thickness were achieved only in a small patient population, with data from only eight patients to date, in an open-label setting, were not statistically significant and might not be achieved by any additional patients observed in this trial or replicated in larger-scale trials that we intend to conduct.

Preclinical Study Comparing the Pharmacokinetic Effects of SCS and Intravitreal Injection of Triesence in Rabbits

We conducted a preclinical study in rabbits to compare the pharmacokinetic results of SCS injections and intravitreal injections of Triesence, a commercially available TA formulation. Pharmacokinetics refers to the process by which a drug is distributed and metabolized in the body, which provides information on drug levels in specific tissues and how these levels change over time. To compare these results, we observed the relative distributions of Triesence in the different tissues and areas of the eye for each method of administration.

In this study, 50 rabbits received a single dose of 4.0 mg of Triesence on day 1 of the study injected either intravitreally or into the SCS. The rabbits were then observed for periods of up to 90 days and the concentration of TA in various parts of the eye was measured at days 14, 28, 56 and 91.

The tables below illustrate the results of this study. For each tissue type or area of the eye, the values shown represent the average concentration of drug observed in the rabbits at the particular measurement points during the study, when comparing the two routes of injection. For clarity of presentation, the tables below include varying concentration scales.

Concentration of TA Following Either SCS or Intravitreal (IVT) Injection

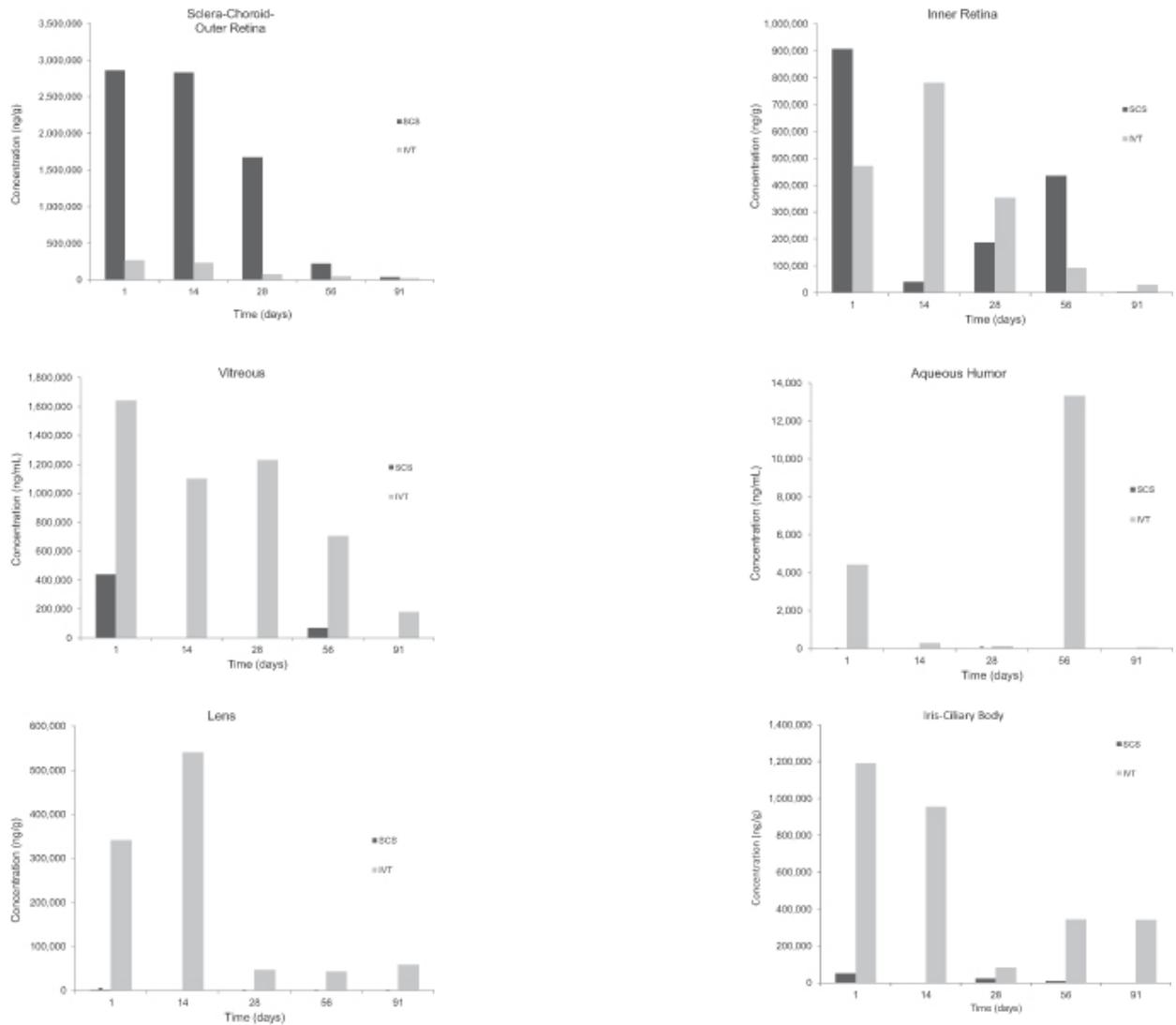


Table of Contents

In the inner sclera, choroid and outer retina, significantly higher concentrations of TA administered into the SCS were present throughout the 91-day period as compared to TA administered intravitreally. The opposite was the case in the iris, ciliary body, lens and aqueous humor, all of which are located at the front of the eye, as well as in the vitreous, with each of these tissues showing higher levels of TA throughout the 91-day period when administered intravitreally as compared to its administration into the SCS. Only minimal levels of TA were present in the iris, ciliary body, lens and aqueous humor when administered into the SCS. Although there was variability in the relative concentrations in the inner retina, which is the part of the retina adjacent to the vitreous, at different time points, the concentrations were generally comparable across both administration methods. In this study, we also compared the drug levels in the blood plasma for each method of administration. Minimal levels of drug in the blood plasma were detected following each method of administration.

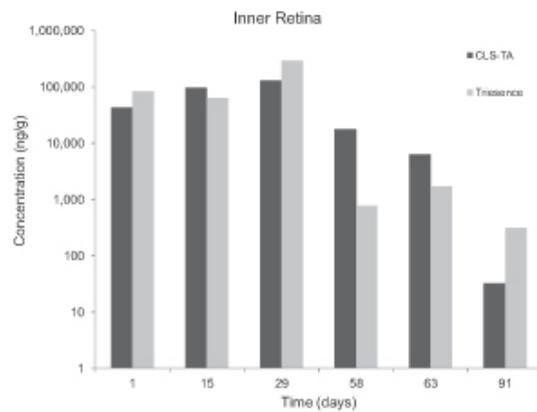
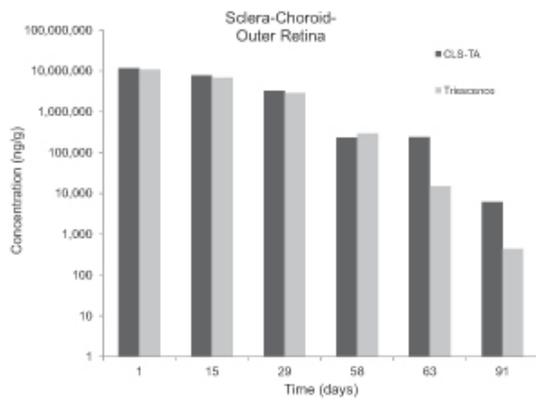
We believe these preclinical study results suggest that drugs administered through the SCS can remain substantially localized within the targeted retina and choroid, and away from other parts of the eye, where they could result in a higher incidence of side effects. We also believe these results suggest that SCS injection provides substantially better bioavailability for at least 90 days in the targeted retina and choroid than intravitreal injection. We intend to further evaluate this bioavailability data in future clinical trials before any potential submission of our marketing application to the FDA.

Preclinical Study Comparing the Pharmacokinetic Effects of SCS Injection of CLS-TA and Triesence in Rabbits

In order to establish that our CLS-TA behaves in the same manner as Triesence despite the differences in formulation, we conducted a preclinical pharmacokinetic study in rabbits, comparing the pharmacokinetic profile of our CLS-TA with the profile of Triesence, each injected into the SCS. In the study, 25 rabbits received a single dose of 4.0 mg of either our CLS-TA or Triesence injected into the SCS on day 1. The rabbits were then observed for periods of up to 90 days and the resulting concentrations of each of the two TA formulations in various parts of the eye was measured at days 15, 29, 58, 63 and 91.

In this study, our CLS-TA and Triesence had comparable distributions throughout the eye over the 90-day period. As shown in the graphs below, both our CLS-TA and Triesence, administered through the SCS, remained present in the retina and choroid throughout the 90-day period following injection.

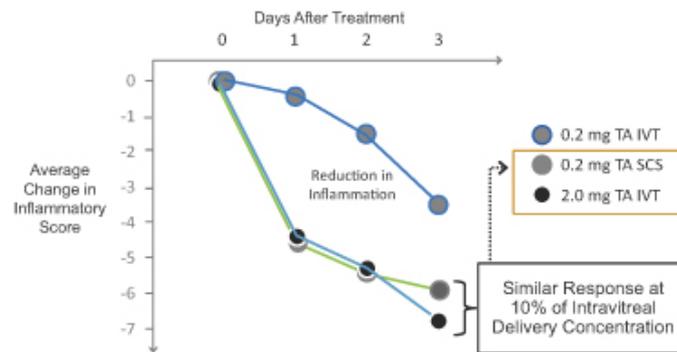
Concentration of Either CLS-TA or Triesence Following SCS Injection



Preclinical Study Evaluating the Pharmacodynamic Effects of SCS and Intravitreal Injection of Triesence in a Pig Model of Uveitis

In this preclinical study, the results of which were published in the journal *Investigative Ophthalmology and Vision Science*, we studied the pharmacodynamics of Triesence injected into the SCS and injected intravitreally in a pig model of uveitis. Pharmacodynamics refers to the biochemical and physiological effects of a drug on the body. In this study, 20 pigs were assigned to one of three drug treatment arms: two intravitreal and one SCS. The intravitreal injections were with doses of either 0.2 mg or 2.0 mg of Triesence and the SCS injection was with a dose of 0.2 mg of Triesence. The pigs were observed for three days after injection to measure their average change in inflammatory score, as measured by an established scale for measuring inflammation in the eye, the modified McDonald-Shadduck grading system. After three days, both the 2.0 mg intravitreal and the 0.2 mg SCS doses were observed to be more effective at reducing uveitic inflammation than the 0.2 mg intravitreal injection. Additionally, the 0.2 mg SCS dose and 2.0 mg intravitreal dose achieved similar reductions in inflammatory scores each day. Therefore, SCS injection of Triesence had a similar pharmacodynamic response to intravitreal administration at only 10% of the dose. The graph below illustrates these results.

Reduction in Inflammation Following Either SCS or Intravitreal Injection of Triesence in a Pig Model of Uveitis



Preclinical Studies Evaluating the Toxicology of SCS Injection of CLS-TA and Triesence in Rabbits

We have conducted toxicology studies in rabbits in which both our CLS-TA and Triesence were well tolerated when injected into the SCS. In one study, 110 rabbits received a single SCS injection of either 3.2 mg or 5.2 mg of Triesence or of a vehicle control without active drug. The rabbits were then evaluated for the following 17 weeks. In the other study, 48 rabbits received an initial SCS injection of either 4.0 mg of our CLS-TA or of vehicle control and were then evaluated for the following 13 weeks. A subgroup of the 48 rabbits received a second SCS injection of our CLS-TA after the first 13 weeks and this subgroup was evaluated for an additional 13 weeks, resulting in a total observation period of 26 weeks. In both studies, the rabbits were observed for tolerability assessments as well as mortality and clinical observations, body weight and food and water consumption.

In these studies, all doses of our CLS-TA and Triesence were well-tolerated. In the study in which rabbits received our CLS-TA, a slight increase in IOP was observed. There were no other treatment-related or administration-related side effects, and in each case localization of TA in the back of the eye was observed, with minimal exposure in the front of the eye.

Planned Phase 2 Clinical Trial

Based on our preclinical data and on our consultation with the FDA, we plan to conduct a Phase 2 randomized, controlled, multi-center clinical trial in the United States in approximately 30 patients with macular edema associated with non-infectious uveitis. We currently expect to initiate this trial in the second half of 2014 after we have completed dosing in the ongoing Phase 1/2 clinical trial. Of the approximately 30 patients to be enrolled, approximately equal numbers of patients will receive either a 4.0 mg dose of our CLS-TA, which is the standard dose of TA when administered intravitreally, or a lower dose of 0.8 mg of our CLS-TA, in both cases administered through the SCS with our microinjector. The trial will provide safety and efficacy information on SCS injection of our CLS-TA using the standard dose. We intend to explore the lower dose in this trial to assess whether efficacy may be observed at the lower dose. If the results of the Phase 2 clinical trial suggest that a lower dose is effective, we may seek to explore, and potentially seek approval of, the lower dose subsequent to the completion of our planned Phase 3 clinical trial using the standard 4.0 mg dose.

The primary efficacy endpoint will be the mean change from baseline in retinal thickness at two months after treatment. Secondary efficacy endpoints will include visual acuity improvements at one and two months post-treatment, measured by the mean change in BCVA from baseline. Safety measures will be monitored over the two-month observation period and will include the incidence of adverse events and serious adverse events, including cataracts and increases in IOP.

Planned Pivotal Phase 3 Clinical Trial

We intend to conduct a single pivotal Phase 3 randomized, controlled, multi-center clinical trial in patients with macular edema associated with non-infectious uveitis to support a Section 505(b)(2) NDA for this indication, which we expect to be able to file in the first half of 2017. We expect to commence this trial in the first half of 2015.

We held a pre-IND meeting with the FDA in September 2012, at which we discussed our proposed clinical development plan for CLS-1001. We then scheduled a meeting with the FDA for August 2013 and submitted questions related to our planned clinical trials. Based on the written responses that we received from the FDA, we elected not to proceed with the requested meeting, and the responses from the FDA became the final minutes for the meeting. Based on these discussions and this correspondence with the FDA, we do not expect to be required to wait for findings from our Phase 2 clinical trial to begin the single pivotal Phase 3 clinical trial and we expect that the single pivotal trial will be sufficient to support a potential Section 505(b)(2) NDA filing. We expect to enroll approximately 150 patients with macular edema associated with non-infectious uveitis, randomized 1:1 either to a treatment arm or to a sham injection arm. We expect to use a sham injection as a comparator for CLS-1001, as opposed to an active drug, because there are no approved therapies for macular edema associated with non-infectious uveitis against which to compare CLS-1001. All of the approximately 75 patients in the treatment arm will receive a 4.0 mg dose of our CLS-TA injected into the SCS using our microinjector. In order to simulate an injection to maintain masking, the sham injection procedure will include all steps involved in the SCS injection procedure, except that an empty microinjector without a needle will be used to apply pressure to the eye. We anticipate that the treatment arm will receive an initial SCS injection of our CLS-TA at the beginning of the trial and a second SCS injection of our CLS-TA at week 12.

The primary efficacy endpoint of the trial will be mean change in BCVA from baseline at week 26. Secondary efficacy endpoints will include additional measures of change in visual acuity and reductions in

[Table of Contents](#)

retinal thickness from baseline. Safety measures will be monitored over the 26-week observation period and will include the incidence of adverse events and serious adverse events, including cataracts and increases in IOP.

Regulatory Approval Pathway of CLS-1001

If our single pivotal Phase 3 clinical trial is successful, we intend to seek regulatory approval of CLS-1001 by utilizing Section 505(b)(2) of the FDCA. As part of our NDA submission under Section 505(b)(2), we intend to rely on the results from our ongoing Phase 1/2 clinical trial, our planned Phase 2 and single pivotal Phase 3 clinical trials conducted under our existing IND for CLS-TA, as well as the FDA's previous findings of safety and efficacy for TA and an analysis of available data from clinical literature.

Based on our pre-IND meeting and subsequent written correspondence with the FDA, we believe this single pivotal Phase 3 clinical trial will be sufficient to support our NDA filing for macular edema associated with non-infectious uveitis pursuant to Section 505(b)(2). We also intend to base any foreign marketing applications, in part, on data obtained through these trials.

Orphan Drug Application for CLS-1001

We have applied for orphan drug designation from the FDA in the United States, and intend to apply for orphan drug designation from the European Medicines Agency in the European Union for CLS-1001 for the treatment of non-infectious uveitis. Orphan drug designation for this indication would make us eligible for seven years of marketing exclusivity in the United States and 10 years of marketing exclusivity in the European Union, if we receive the first marketing approval of TA for macular edema associated with non-infectious uveitis. Orphan drug designation would also provide us with certain tax incentives and user fee waivers. However, even if we are able to obtain orphan drug designation, it will not convey any advantage in, or shorten the duration of, the regulatory review and approval process for our marketing application. In addition, even if we ultimately obtain marketing exclusivity, competition is still possible if, for example, different drugs are approved for the same indication, a similar drug is shown to be clinically superior to ours, doctors prescribe a competitor's products for off-label use, or a patent to which we do not have rights covers our drug product, in each case undermining our orphan drug exclusivity and our opportunity to market our drug successfully, if at all.

CLS-1003 Program Targeting Macular Edema Associated with Retinal Vein Occlusion

We are developing CLS-1003 for treatment of macular edema associated with RVO, a sight-threatening disorder resulting from the blockage of a retinal vein. CLS-1003 consists of an SCS injection of CLS-TA with our microinjector, for administration together with an intravitreal injection of an anti-VEGF drug. We plan to initiate a Phase 2 clinical trial in approximately 40 patients before the end of 2014, with data expected in the second half of 2015. We believe that CLS-1003 may provide faster onset of therapeutic effect compared to currently used intravitreal anti-VEGF injections alone, while also reducing the frequency of required intravitreal anti-VEGF treatments from once every 30 days to once every 90 days. In addition, we believe that the side effect profile of CLS-1003 will be similar to that of current intravitreal anti-VEGF injections alone.

Market Opportunity for Treatment of Macular Edema Associated with RVO

RVO is a sight-threatening disorder resulting from a blockage of one of the veins carrying blood out of the retina. RVO is estimated to affect more than 16 million adults worldwide, according to a 2010 study

[Table of Contents](#)

published in the journal *Ophthalmology*, and we estimate RVO affects 2.2 million individuals in the United States. In RVO, the blockage of a retinal vein can lead to poor blood circulation, low oxygen, and sometimes inflammation. A blocked vein will leak its contents of blood and fluid. Bleeding within the retina and swelling from fluid can create macular edema.

Limitations of Currently Available Therapies for Macular Edema Associated with RVO

There are three primary treatments currently used for macular edema caused by RVO: intravitreal injections of anti-VEGF medications, intravitreal corticosteroid injections and laser surgery. The current standard treatment for macular edema associated with RVO involves intravitreal injections of anti-VEGF drugs, such as Lucentis, Eylea or Avastin. Lucentis and Eylea are indicated for monthly administration and Avastin is used in a similar fashion, but off-label. These monthly treatments are required throughout the course of the disease, which could last for years. Anti-VEGF drugs are effective in significantly improving vision in approximately 30% to 40% of patients with macular edema associated with RVO, and have limited side effects.

Corticosteroid treatment, specifically TA injected intravitreally, was previously evaluated in a series of randomized, multi-center clinical trials, called the SCORE studies, conducted by the National Eye Institute. In these studies, approximately 20% of patients with macular edema associated with RVO had meaningful gains in visual acuity after one year with a dosing frequency of one intravitreal injection of TA every four months. However, several side effects were observed, including cataracts and elevated IOP. Because of these side effects, intravitreal administration of corticosteroids, such as TA, is not the preferred therapy, even though it has been observed to be effective in reducing macular edema associated with RVO. Another therapy, Ozurdex, an extended release corticosteroid implanted in the vitreous, has also been approved by the FDA for the treatment of RVO, but has also been associated with increased IOP in 25% of patients, conjunctival hemorrhages in 22% of patients and cataracts in 5% of patients.

Before the introduction of anti-VEGF drugs, laser surgery was the leading therapy for treating macular edema associated with RVO. Laser surgery can be used to help control bleeding and swelling, which can reduce the loss of sight, but cannot improve vision. Laser surgery, however, cannot be used long term because it permanently damages the retina.

Potential Benefits of CLS-1003

We believe that SCS administration of CLS-TA, by addressing the inflammatory aspect of RVO, in combination with intravitreal anti-VEGF treatment, addressing the vascular aspect of RVO, may result in faster onset of therapeutic effect compared to currently used intravitreal anti-VEGF treatment alone. Based on the SCORE studies described above and our findings in preclinical studies, we believe that each SCS injection of CLS-TA may provide a duration of effect longer than anti-VEGF therapy alone, while also potentially reducing the frequency of necessary anti-VEGF treatments from once every 30 days to once every 90 days, due to the benefits of both corticosteroid and anti-VEGF treatment. In addition, we believe that the side effect profile of CLS-1003 will be similar to that of current intravitreal anti-VEGF treatments alone, as SCS injections of CLS-TA should allow the corticosteroid to remain substantially localized in the retina and choroid without introducing the side effects that are commonly observed with other routes of administration of corticosteroids.

Planned Clinical Trials

In support of our planned clinical development program for CLS-1003, we are in the process of completing a toxicology study in rabbits with the SCS injection of CLS-TA together with the intravitreal injection of the anti-VEGF drug Eylea. Depending on the outcome of this study, we intend to initiate a Phase 2 randomized, multi-center, double-masked clinical trial of CLS-1003 before the end of 2014.

Based on our ongoing toxicology study and our clinical and preclinical experience with SCS administration of TA in our CLS-1001 program, we anticipate being able to proceed directly into a Phase 2 clinical trial for CLS-1003. The goal of the Phase 2 clinical trial will be to demonstrate that patients treated with CLS-TA administered into the SCS together with Eylea administered intravitreally may require less frequent treatments than monotherapy of Eylea administered intravitreally.

We expect to enroll approximately 40 patients at 10 sites in the United States for the Phase 2 clinical trial. All patients will start by receiving one intravitreal injection of 2.0 mg of Eylea, in a total volume of 50 microliters, and will then be randomized on a 1:1 basis to receive an SCS injection of either 4.0 mg of our CLS-TA, in a total volume of 100 microliters, or a sham procedure in the same visit. After randomization, patients will be seen in the clinic once per month for three months. Patients in the CLS-TA treatment arm will receive additional intravitreal injections of Eylea at the subsequent visits one and two months after the initial injection if they continue to experience increases in macular edema or reductions in visual acuity. If they do not experience increases in macular edema or reductions in visual acuity at these subsequent visits, they will not receive additional intravitreal Eylea treatment. Patients in the control arm will receive monthly intravitreal Eylea injections, as specified on the Eylea label.

The primary objective of the trial will be to evaluate the safety and efficacy of a single SCS injection of CLS-TA together with the initial intravitreal injection of Eylea, compared to monthly intravitreal Eylea injections alone. The primary efficacy endpoints in the trial will include determining the number of required Eylea treatments in each arm, which we believe will provide an indication of whether concomitant therapy provides any advantage to the patient in reducing the number of required Eylea treatments. Secondary efficacy endpoints will include measures of change in visual acuity and reductions in retinal thickness from baseline. The safety endpoints will be the incidence of adverse events and serious adverse events, including increases in IOP.

Regulatory Approval Pathway

To date, we have not met formally with the FDA to discuss our clinical development plan for the CLS-1003 program. However, we intend to meet with the FDA following the completion of our Phase 2 clinical trial to discuss the details of the regulatory approval pathway for CLS-1003. If the results of our planned Phase 2 clinical trial are positive, we expect to be able to follow a 505(b)(2) NDA regulatory approval pathway and to conduct one or two pivotal Phase 3 clinical trials in order to support an NDA submission for CLS-1003 in macular edema associated with RVO. In pursuing the 505(b)(2) regulatory pathway, we intend to rely on the results from our planned CLS-1003 clinical trials, the FDA's previous findings of safety and efficacy for TA, and peer-reviewed literature.

CLS-1002 Program Targeting Wet Age-Related Macular Degeneration

Under our CLS-1002 program, we intend to develop a treatment for wet AMD that, through SCS administration with our microinjector, could potentially reduce the frequency of necessary treatments compared to the current standard of care. Wet AMD is a condition involving the abnormal formation and

growth of new blood vessels in the choroid, their intrusion into the outer layers of the retina and leakage of blood and fluid into the retina. We have completed a Phase 1 clinical trial evaluating the safety and efficacy of SCS injection of Avastin, an anti-VEGF drug, in four patients with wet AMD. We are evaluating a number of compounds as SCS-injected therapies for the treatment of wet AMD, including compounds with anti-VEGF activity, and compounds with dual anti-VEGF and anti-PDGF activity, and expect to select a lead drug candidate for IND submission in 2015. We believe that CLS-1002 may show faster onset of therapeutic effect as compared to intravitreal injection, while potentially reducing the frequency of necessary treatment. We plan to further study this effect in any future clinical trials that we conduct as part of our CLS-1002 program.

Market Opportunity for Treatment of Wet AMD

Wet AMD is a leading cause of severe vision loss and blindness in people over the age of 50 in the developed world. If untreated, the blood vessel growth and leakage associated with wet AMD can eventually lead to blindness. The majority of patients with wet AMD experience severe vision loss in the affected eye within approximately two years after diagnosis of the disease. According to a study published in 2004 in the journal *Ophthalmology*, approximately 1.2 million people in the United States suffer from wet AMD. Based on estimates from AMD Alliance International, a non-profit organization focused on AMD awareness, and census growth data, we estimate there are approximately 230,000 new cases of wet AMD each year in the United States. Because the risk of developing wet AMD increases with age, we expect that the overall aging of the population will result in a significant increase in the number of wet AMD cases, both in the United States and worldwide.

Limitations of Currently Available Therapies for Wet AMD

The current standard of care for the treatment of wet AMD is administration by intravitreal injection of anti-VEGF drugs as a standalone therapy. The anti-VEGF drugs most commonly used include Lucentis and Eylea, both of which have been approved by the FDA for the treatment of wet AMD, and Avastin, which is used off-label for the disease.

Because anti-VEGF therapies inhibit abnormal blood vessel growth and leakage, they are effective in reducing the macular edema associated with wet AMD. Anti-VEGF therapies have demonstrated the ability to prevent further visual loss in approximately 90% of treated patients. A retrospective study published in 2012 in the journal *Archives of Ophthalmology* concluded that the prevalence of both legal blindness and moderate visual impairment in patients two years after being diagnosed with wet AMD had decreased substantially following the introduction of anti-VEGF therapy. However, because anti-VEGF drugs do not appear to cause any regression of the underlying neovascularization associated with wet AMD, the current standard of care requires regular injections, typically as often as seven times per year, and therapy may be required indefinitely. Further, while anti-VEGF treatment is effective at preventing vision loss, only 30% of patients improve their visual acuity by 15 letters or more. Anti-VEGF treatments have been associated with subretinal fibrosis, or the formation of excess connective tissue under the retina, as well as retinal scarring in some patients, with 45% of patients experiencing scarring after two years of anti-VEGF therapy according to a 2014 study published by the American Academy of Ophthalmology. Additionally, monthly treatment with anti-VEGF agents have been associated with increased risk of geographic atrophy, according to a 2013 study published by the American Academy of Ophthalmology.

In addition to anti-VEGF therapies, anti-PDGF drugs are being developed as potential treatments for wet AMD. Because PDGF is a major factor in the formation and stabilization of blood vessel growth in the

[Table of Contents](#)

choroid, therapies inhibiting PDGF may also be effective in addressing the abnormal growth of new blood vessels associated with wet AMD. In clinical trials, an intravitreal injection of a PDGF inhibitor together with an intravitreal injection of Lucentis improved visual acuity in approximately 60% of patients with wet AMD compared to approximately 30% of patients receiving Lucentis alone. However, no anti-PDGF therapy has been approved by the FDA for the treatment of wet AMD. Even if approved, we expect that anti-PDGF therapy would also require regular injections indefinitely.

Even though anti-VEGF and anti-PDGF therapies may be effective in treating wet AMD, intravitreal administration of these drugs relies on the process of diffusion of the drug from the vitreous outward to the retina and, eventually, the choroid, which results in lower bioavailability and the need for frequent retreatment.

Potential Benefits of CLS-1002

The compounds we are evaluating have anti-VEGF activity, and we believe these compounds should provide the clinically established advantages that currently used anti-VEGF drugs provide, especially the reduction in macular edema and improvements in visual outcomes. Some of the compounds we are evaluating have dual anti-VEGF and anti-PDGF activity. We believe that the anti-PDGF properties together with anti-VEGF properties may provide superior visual outcomes to standalone anti-VEGF drugs. We believe that the SCS administration of drugs with anti-VEGF, or dual anti-VEGF and anti-PDGF, activity more directly to the neovascularization in the choroid through SCS injection could block the process of additional new vascular growth within the choroid before the vessels break into and create further damage through leakage into the retina. We believe that such SCS injection of treatment may, therefore, provide faster onset of therapeutic effect and delay vision loss for a longer duration of time than the current standard intravitreal administration and will require less frequent administration, potentially only once every 90 days. We will test this hypothesis in any future clinical trials that we may conduct as part of our clinical development program for CLS-1002.

Clinical and Preclinical Development of CLS-1002

We have completed the following clinical trial and preclinical studies as part of our CLS-1002 development program:

- a completed Phase 1 clinical trial in wet AMD patients evaluating the safety and tolerability of SCS injection of Avastin, an anti-VEGF drug, with our microinjector;
- a completed preclinical efficacy study of a compound with dual anti-VEGF and anti-PDGF activity in a rabbit model of wet AMD; and
- a completed preclinical safety study of SCS injection of compounds with dual anti-VEGF and anti-PDGF activity.

Details of the trial and studies are summarized below.

Completed Phase 1 Clinical Trial with Avastin

In order to evaluate the safety and tolerability of SCS administration of Avastin, an anti-VEGF drug, in humans with our microinjector, we conducted a Phase 1, single-center, open-label clinical trial in 2012 at the Hospital of the Association to Prevent Blindness in Mexico. Because of its established safety profile and its prevalence as an off-label treatment for wet AMD, we selected Avastin as the therapy to be evaluated in

this exploratory trial. Four patients with wet AMD were enrolled in the trial, each of whom was between the ages of 63 and 73. Each patient received a 2.5 mg dose of Avastin in a 100 microliter injection in one eye using a prototype of our microinjector. The trial was conducted in accordance with current U.S. good clinical practices.

Trial Design. Eligibility criteria included adult patients with choroidal neovascularization associated with wet AMD that had previously responded to Avastin treatment, had retinal thickness in the study eye of between 275 and 600 microns in the central subfield, and had specified levels of visual acuity. In addition, patients could not have received any anti-VEGF treatment within 60 days of screening or have had any expectation of receiving such treatment during their participation in the study other than the single Avastin injection.

Once eligibility was established, each patient received a single SCS injection of Avastin and then returned for a follow-up examination on the day after the injection. Thereafter, patients returned for five additional evaluations at approximately weeks 1, 2, 3, 4 and 8 following the injection. No additional treatments were administered during the course of the trial.

Endpoints. The trial was primarily a safety and tolerability study. The safety endpoints included incidence, intensity and type of adverse events, as well as changes from baseline in IOP. We also measured efficacy endpoints, including change from baseline in BCVA letter score.

Safety results. SCS injection of Avastin in the trial was observed to be generally well tolerated. No increases in IOP or serious or unexpected adverse events were reported in any of the patients. Two patients experienced hyperemia, or redness in the study eye, which was mild in severity, was determined to be unrelated to treatment in the trial and resolved within a day without other treatment.

Efficacy results. Two of the four patients achieved an improvement in BCVA letter score of at least 13 letters at the end of the eight-week trial following the SCS injection of Avastin. One of these patients improved by 16 letters within one day of treatment and maintained that level of improvement for the remainder of the trial. The second patient achieved an improvement of four letters by one week after treatment, which improved to 10 letters by week 3 after treatment and 13 letters by week 8. The other two patients in the trial maintained their BCVA letter score within four letters of their baseline at each measurement point in the trial. Overall, there was an average gain of nine letters in the four patients at the end of the eight-week observation period following the SCS injection of Avastin.

By comparison, in a trial conducted by the National Eye Institute, the Comparison of AMD Treatment Trial, or CATT, which was a Phase 3 randomized, masked, controlled, multi-center study, the efficacy of Avastin was compared to that of Lucentis, which has been approved by the FDA for the treatment of wet AMD. In the CATT trial, patients in the respective treatment arms received either monthly intravitreal injections of Lucentis or monthly intravitreal injections of Avastin over two years. After one year, patients receiving monthly Lucentis treatment experienced an average improvement in BCVA of 8.5 letters, while those receiving Avastin experienced an average improvement of 8.0 letters. After two years, the average improvement for the patients receiving Lucentis and Avastin was 8.8 letters and 7.8 letters, respectively.

Our Phase 1 trial was not designed to show any efficacy results with statistical significance and was conducted in an open-label, uncontrolled setting. While the data from this exploratory trial suggested potential improvement in visual acuity over eight weeks following SCS injection of Avastin, it is important to note that the results from this trial were observed in only four patients and might not be achieved by any other patient treated with a product candidate to be developed as part of our CLS-1002 program. Any later-

stage trials intended to support an application for regulatory approval of a product candidate for this indication will need to show statistical significance in larger, well-controlled clinical trials.

Preclinical Efficacy Study of a Compound with Dual Anti-VEGF and Anti-PDGF Activity in a Rabbit Model of Wet AMD

We conducted a preclinical study assessing the effect on retinal leakage in a wet AMD rabbit model of a drug candidate with dual anti-VEGF and anti-PDGF activity administered by SCS injection. In this study, the treatment arm of three rabbits received an SCS injection of the drug and the control arm of three rabbits received an SCS injection of a vehicle without the active drug. After 28 days, retinal leakage was induced on the surface vessels of the inner retina in both arms and leakage was measured approximately two hours after this induction. The treatment arm showed 48% less retinal leakage than the vehicle arm. We believe this suggests that this dual anti-VEGF and anti-PDGF drug administered through the SCS may be effective in treating wet AMD by reducing leakage in the retina, and even in vessels in the inner retina, which is closest to the vitreous. We are continuing to evaluate this compound.

Preclinical Safety Study of SCS Injection of Compounds with Dual Anti-VEGF and Anti-PDGF Activity

We conducted a preclinical study assessing the safety of compounds with dual anti-VEGF and anti-PDGF activity administered by SCS injection. In this study, a total of 18 rabbits received an SCS injection of a number of compounds and were evaluated on days 1, 4 and 7. Overall, the compounds were well tolerated with only mild or transient ocular effects observed.

Planned Clinical Development

Based on the results of our Phase 1 clinical trial in four wet AMD patients and our preclinical rabbit study, we believe that wet AMD may be effectively treated through SCS injection, and we will evaluate this hypothesis as part of any clinical development program for CLS-1002. We will continue to evaluate the SCS injection of a number of compounds with anti-VEGF and anti-PDGF activity to assess their potential for clinical development for the treatment of wet AMD. Once we select a lead compound for clinical development, we expect to perform pharmacokinetic and toxicology studies, as well as additional preclinical testing, to support an IND filing by the end of 2015.

Future Potential Product Candidates

We believe that our SCS-focused approach has the potential to become more broadly used for the treatment of other back of the eye diseases, and we intend to develop additional product candidates for SCS injection based on the results of our current and planned clinical trials. We will then seek to secure appropriate regulatory authorizations to begin additional clinical testing for any such product candidates. In addition to uveitis, RVO and wet AMD, there are a number of ophthalmic conditions affecting the retina and choroid with unmet medical need for which SCS injection of therapy may be beneficial, including:

- *Diabetic macular edema*, or DME, a common ocular complication of diabetes that results in swelling in the macula and is the primary cause of vision loss associated with diabetes;
- *Polypoidal choroidal vasculopathy*, or PCV, a disorder resulting from the abnormal growth of new blood vessels in the choroid adjacent to the macula. PCV is characterized by dilated and branching blood vessels in “polyp like” groups in the choroid that could lead to leakage; and

- *Geographic atrophy*, an advanced form of AMD, is characterized by a loss of the layers of cells in the retina next to the choroid, eventually including the photoreceptor cells in the macula, leading to gradual irreversible loss of central vision and eventually blindness.

According to the U.S. Centers for Disease Control and Prevention, or CDC, the number of Americans diagnosed with diabetes has increased from approximately 7.7 million in 1994 to approximately 21.0 million in 2010. All patients with diabetes are at risk of developing some form of diabetic eye disease. DME accounts for a majority of vision loss in patients with diabetic eye disease. According to a study published in the journal *Diabetes Care* in 2003, patients with diabetes have a 10% risk of developing clinically significant DME over a 10-year period. A study published in the journal *Ophthalmology* in 2009 found that over a 25-year period, approximately 17% of diabetics studied were diagnosed with DME. We estimate approximately 1.1 million individuals in the United States suffer from DME.

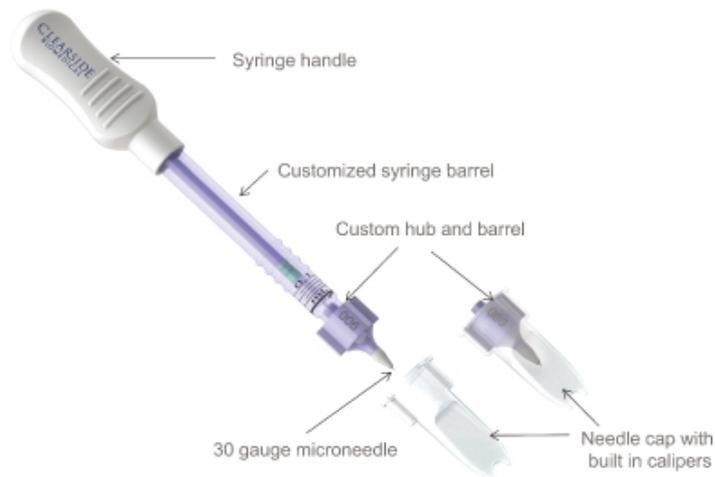
Currently, the only FDA-approved drugs for the treatment of DME are the anti-VEGF drugs Lucentis and Eylea and the intravitreal corticosteroid implant Ozurdex. Some retinal specialists also prescribe other anti-VEGF drugs for off-label use, such as Avastin, as well as corticosteroid therapies, primarily Kenalog and Triescence, and laser photocoagulation to treat DME. Laser photocoagulation is a procedure in which a laser is used to apply a burn or a pattern of burns to cauterize leaky blood vessels to reduce edema in the retina. Studies have shown that both anti-VEGF drugs and corticosteroids are effective in some DME patients. However, both anti-VEGF drugs and corticosteroids are limited by a need for multiple injections to maintain a therapeutic effect. This raises concerns, not only for patients, but also for caregivers who are affected by frequent doctor visits, as well as for healthcare providers who must monitor patients monthly. In addition, these therapies have safety concerns. Corticosteroids administered intravitreally have historically been associated with acceleration of cataract formation and significant increases in IOP, which may increase the risk of glaucoma, and monthly anti-VEGF treatments have been associated with increased risk of geographic atrophy. Laser photocoagulation can also have undesirable side effects, including partial loss of peripheral and night vision.

Our development program for DME will be modeled after our approach for addressing macular edema associated with RVO. We expect that any clinical trials we may conduct for this indication will be designed based on the results observed in our Phase 2 clinical trial of CLS-1003 in RVO patients.

Our Proprietary SCS Microinjector

Our drug candidates have been and will be specifically formulated to be injected with our microinjector into the SCS in order to spread around to the back of the eye. The single-use microinjector is intended to consistently administer drug into the SCS, in volumes similar to the amount of drug commonly used in an intravitreal injection. If approved for marketing by applicable regulatory authorities, the microinjector will be packaged with two 30-gauge hollow microneedles of varying lengths, each less than 1.2 millimeters, to account for varying scleral thicknesses, within a custom-designed hub that optimizes insertion into the eye. Based on our discussions with the FDA to date, we do not anticipate that the FDA will require a separate medical device authorization for the microinjector, but this could change during the course of its review of any marketing application that we may submit.

Our microinjector, shown below, can be used to inject a wide variety of drugs into the SCS.

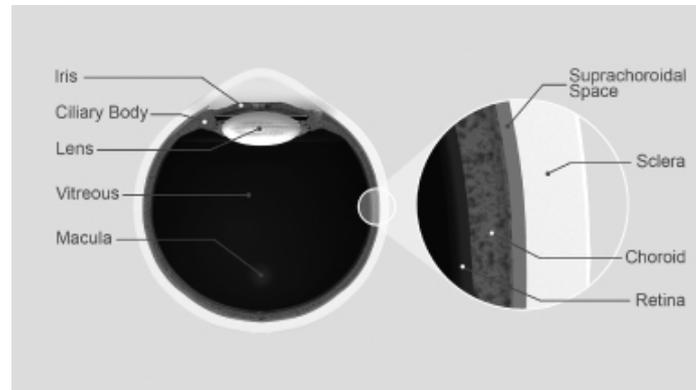


The SCS microinjection is designed to be made perpendicular to the sclera, at a site similar to an intravitreal administration, under local anesthesia. Once the microneedle penetrates the sclera and reaches the SCS, the boundary between the sclera and the choroid, the open bevel of the needle releases the drug between these two layers. Once drug is injected into the SCS, it spreads within the SCS to the back of the eye, where diseases manifest. The preparation and injection will require minimal training for the administering retinal specialist and can be accomplished in an in-office setting.

Current intravitreal injections are performed in a procedure similar to that of SCS injections, except that the hypodermic needles and syringes used in intravitreal injection procedures are designed to inject through all of the layers of the eye and into the vitreous, where the precise spacial location of the needle is not as important as when injecting into the SCS. Intravitreal injections are typically performed using a needle that is approximately five millimeters in length, or four times the length of our microneedle. Using a needle of this length, the physician penetrates past the sclera, choroid and retina until reaching the vitreous and does not receive any tactile feedback as to when the needle had reached one of the layers between the sclera and the vitreous. With our microinjector, however, the needle hub is designed to enable the release of the drug into the SCS only after the needle passes the sclera and to prevent penetration into the layers beyond the SCS.

Background on Eye Disease

The human eye possesses focusing elements in the front, the cornea and lens, and a light-sensing element in the back, the retina. Light falls on the photoreceptors that are part of the retina, called rods and cones, and is converted into electrical energy, which travels via the optic nerve to the brain. The central portion of the retina is the macula, which is the region responsible for seeing color and the acute central vision necessary for activities such as reading, face recognition, watching television and driving. The brain processes the complex signals sent from the retina into vision. The following diagram illustrates the principal elements of the anatomy of a healthy eye, including a detailed cross-section of the back of the eye, which refers to the portion of the eye behind the lens.



Role of the Choroid in Retinal Disease

Eye disease can be caused by many factors and can affect both the front and back of the eye. In its most extreme cases, eye disease can result in either partial or total blindness. In the developed world, the major diseases that result in blindness are those affecting the retina. Millions of people live with varying degrees of irreversible vision loss because they have a degenerative eye disorder that affects the retina. In these conditions, the retina is damaged, affecting the retina's ability to send light signals to the brain. The choroid provides blood, oxygen and nourishment to the outer layers of the retina, but it is also thought to regulate retinal heat, to assist in the control of IOP and to absorb excess light so as to avoid reflection that can damage the retina.

The choroid can be subject to disorders that can lead to retinal damage and vision loss, including inflammatory disorders, vascular disorders and tumors. Our focus is on the first two categories of disorders. When the choroid and surrounding tissue become inflamed from an immune response, it can result in macular edema, or swelling in the macular region of the retina, which often results in distorted vision or even blindness. In addition, in the case of choroidal neovascularization, abnormal blood vessels forming from choroidal blood vessels may rupture and leak into the retina, also leading to macular edema.

Back of the Eye Diseases

Some of the more common diseases that originate or manifest in the retinal and choroidal areas of the eye include:

Uveitis

Uveitis is a group of ocular conditions that are characterized by inflammation that typically affects the choroid and surrounding tissues. Prolonged or severe inflammation in the back of the eye associated with uveitis can result in the breakdown of cells at the interface of the retina and choroid, leading to the leakage and accumulation of fluid in the macular region of the retina. This fluid build-up can cause macular edema, which can rapidly result in distortion of vision, and eventually blindness, given the macula's critical role in vision. Macular edema is the most frequent cause of visual impairment among patients with uveitis.

Uveitis is one of the most frequent causes of blindness in the developed world. Based on prevalence data published in the journal *Ophthalmology* in 2004 and United States census data for 2010, we estimate approximately 350,000 individuals in the United States suffer from some form of uveitis. Uveitis can be either infectious, caused by an immune response to fight an infection inside the eye, or non-infectious. Non-infectious uveitis accounts for approximately 80% of all uveitis cases.

Uveitis is commonly treated with corticosteroids. Corticosteroids administered through eye drops, although used to treat some uveitis patients, are generally ineffective in treating patients with macular edema associated with non-infectious uveitis. Corticosteroids administered orally or systemically can be effective in treating patients with macular edema associated with non-infectious uveitis, but their long-term use is associated with significant side effects. As a result of the shortcomings associated with these methods of administering corticosteroids, the most common method for treating macular edema associated with non-infectious uveitis is intravitreal injection or implant of a corticosteroid, such as TA, dexamethasone or fluocinolone acetonide.

Retinal Vein Occlusion

RVO is a sight-threatening visual disorder resulting from a blockage of one of the veins carrying blood out of the retina. In RVO, the blockage of a retinal vein can lead to poor blood circulation, low oxygen, and sometimes inflammation in the area that is normally drained by the affected vein. A blocked vein will leak its contents of blood and fluid. This leakage within the retina and the consequential swelling from fluid can cause macular edema. Persistent inadequately treated macular edema associated with RVO can blur vision, cause significant loss in visual acuity and eventually lead to blindness.

RVO is estimated to affect more than 16 million adults worldwide, according to a 2010 study published in the journal *Ophthalmology*. We estimate RVO affects 2.2 million individuals in the United States.

RVO can be treated and vision can be restored in most cases when treatment is administered relatively soon after initial diagnosis and aggressive treatment is obtained. One common method of reducing the macular edema associated with RVO is to use an anti-VEGF drug. Anti-VEGF drugs are effective in drying the leaking fluid and, therefore, they are useful in eye conditions where edema is a complication of the disease. An added challenge of RVO is inflammation that arises in response to the vascular damage and further contributes to the swelling by causing even more fluid to accumulate in the retina. Since inflammation also plays a role in vein occlusion, corticosteroids may provide an added advantage in the reduction of macular edema associated with RVO because they block the inflammatory pathways and also act to stabilize membranes.

Age-related Macular Degeneration

AMD is a chronic, progressive disease of the macula that results in the loss of central vision. The most common symptoms are a central blurred or blank spot, distortion of objects or simply blurred vision. Peripheral vision usually remains intact. The disease typically affects patients initially in one eye, with a high likelihood of it occurring in the second eye over time. Because AMD is strongly correlated with aging, the disease may recur, notwithstanding treatment, as the aging process continues.

There are two forms of AMD, “dry” AMD and “wet” AMD. Dry AMD is the most common form, representing approximately 85% to 90% of all AMD cases. However, dry AMD cases can develop into wet AMD, and wet AMD accounts for 90% of the severe vision loss associated with AMD.

Wet AMD occurs when new blood vessels in the choroid intrude into the retinal layers. This is referred to as choroidal neovascularization. Abnormal new blood vessels tend to be fragile and often bleed and leak fluid into the macula. Untreated, blood vessel growth and associated leakage typically lead to retinal distortion and eventual retinal scarring, resulting in irreversible destruction of the macula and loss of vision. This visual loss occurs rapidly with a progressive course.

According to a study published in 2004 in the journal *Ophthalmology*, approximately 1.2 million people in the United States suffer from wet AMD. Based on estimates from AMD Alliance International and census growth data, we estimate there are approximately 230,000 new cases of wet AMD each year in the United States. Because the risk of developing wet AMD increases with age, we expect that the overall aging of the population will result in a significant increase in the number of wet AMD cases, both in the United States and worldwide.

The current standard of care for the treatment of wet AMD is administration by intravitreal injection of anti-VEGF drugs as a standalone therapy. Because anti-VEGF therapies inhibit abnormal blood vessel growth and leakage, they are effective in reducing the macular edema associated with wet AMD. Anti-VEGF treatments have been associated with subretinal fibrosis, or the formation of excess connective tissue under the retina, as well as retinal scarring in some patients. In addition to anti-VEGF therapies, anti-PDGF drugs are being developed as potential treatments for wet AMD. Even though anti-VEGF and anti-PDGF therapies may be effective in treating wet AMD, intravitreal administration of these drugs relies on the process of diffusion of the drug from the vitreous outward to the retina and, eventually, the choroid, which results in lower bioavailability and the need for frequent retreatment. Patients typically receive injections as often as seven times per year to manage this chronic disease.

Diabetic Macular Edema

DME is a complication of diabetic retinopathy, a disease affecting the blood vessels of the retina. Diabetic retinopathy results in multiple abnormalities in the retina, including retinal thickening, hemorrhages, impeded blood flow, excessive leakage of fluid from blood vessels and, in the final stages, abnormal blood vessel growth. When the blood vessel leakage causes swelling in the macula, it is referred to as DME. Poorly controlled blood glucose levels, high blood pressure, abnormal kidney function causing fluid retention, high cholesterol levels and other general systemic factors are risk factors associated with DME. The principal symptom of DME is a severe loss of central vision.

According to the CDC, the number of Americans diagnosed with diabetes has increased from approximately 7.7 million in 1994 to approximately 21.0 million in 2010. According to a study published in the journal *Diabetes Care* in 2003, patients with diabetes have a 10% risk of developing clinically significant DME over a 10-year period. A study published in the journal *Ophthalmology* in 2009 found that

[Table of Contents](#)

over a 25-year period, approximately 17% of diabetics studied were diagnosed with DME. We estimate approximately 1.1 million individuals in the United States suffer from DME.

Currently, the only FDA-approved drugs for the treatment of DME are the anti-VEGF drugs Lucentis and Eylea and the intravitreal corticosteroid implant Ozurdex. Some retinal specialists also prescribe other anti-VEGF drugs for off-label use, such as Avastin, as well as corticosteroid therapies, primarily Kenalog and Triescence, and laser photocoagulation to treat DME. Studies have shown that both anti-VEGF drugs and corticosteroids are effective in some DME patients. However, both anti-VEGF drugs and corticosteroids administered intravitreally are limited by a need for frequent injections to maintain a therapeutic effect. In addition, these therapies have safety concerns. Corticosteroids administered intravitreally have historically been associated with acceleration of cataract formation and significant increases in IOP, which may increase the risk of glaucoma. Laser photocoagulation can also have undesirable side effects, including partial loss of peripheral and night vision.

Challenges of Ophthalmic Drug Administration

Administration of drugs to treat back of the eye diseases is a significant issue in ophthalmology. Due to the effectiveness of the blood/eye barrier, it is difficult for oral or other systemically administered drugs to reach the eye in sufficient quantities to have a beneficial effect without adverse side effects to other parts of the body. It is also difficult to provide localized delivery of drug to the individual tissues of the eye through common eye drop or intravitreal injection techniques.

There is a need for more localized drug delivery to the back of the eye in a manner that is safe, effective and practical for long-term use. While there have been many attempts to deliver drugs to the back of the eye, most do not achieve sufficient and consistent concentrations for the appropriate period of time. Currently, intravitreal injection remains as the standard for delivering drugs to the retina and choroid. We believe that our novel approach using SCS injection can address several of the shortcomings of current therapies.

Manufacturing

We do not have any manufacturing facilities. We utilize contract manufacturers to formulate and produce our drug candidates and to produce our microinjector components used for our clinical trials. We assemble our microinjector ourselves for our clinical trials. We procure the active pharmaceutical ingredient for our drugs from a third-party supplier. We expect to utilize third-party manufacturers to produce commercial quantities of our drugs and microinjector, if approved. We anticipate entering into commercial supply agreements with these or other manufacturers at a later date.

Commercialization

We intend to develop and, if approved by the FDA, to commercialize our product candidates in the United States, and we may enter into distribution or licensing arrangements for commercialization rights for other regions. Our management team, who will lead the commercialization of our lead product candidates, has substantial experience in sales and marketing based on their participation in the commercialization of ophthalmic drugs at large pharmaceutical companies including Alcon, Allergan, CIBA Vision, ISTA and Novartis.

For marketing in the United States, we intend to hire a dedicated field sales force consisting of approximately 30 to 40 sales professionals to target the approximately 1,700 retinal specialists in the United States.

Competition

The biotechnology and pharmaceutical industries are characterized by rapidly advancing technologies, intense competition and a strong emphasis on proprietary drugs. While we believe that our technologies, knowledge, experience and scientific resources provide us with competitive advantages, we face potential competition from many different sources, including major pharmaceutical, specialty pharmaceutical and biotechnology companies, academic institutions and governmental agencies and public and private research institutions. Any product candidate that we successfully develop and commercialize will compete with existing therapies and new therapies that may become available in the future.

With respect to CLS-TA, we face competition from other commercially available forms of TA and other injectable and implantable corticosteroids. Bristol-Myers Squibb markets TA under the brand name Kenalog. Kenalog is indicated only for intramuscular or intraarticular injection, although it is commonly used off-label for intraocular inflammation. There are also a number of generic equivalents of Kenalog. In addition, Alcon's injectable TA, Trience, is approved in the United States for the treatment of uveitis and other inflammatory conditions unresponsive to topical corticosteroids, although it is not indicated for the treatment of macular edema associated with uveitis or RVO or DME. Ozurdex, marketed by Allergan, is a bioerodable extended release implant that delivers the corticosteroid dexamethasone and is approved for the treatment of macular edema following non-infectious uveitis and RVO in both the United States and in the European Union, and recently received approval from the FDA to treat DME in adult patients who have an artificial lens implant or who are scheduled for cataract surgery. Bausch + Lomb markets Retisert, an intravitreal implant of the corticosteroid fluocinolone acetonide for the treatment of non-infectious uveitis. Alimera Sciences is developing Iluvien, an injectable form of fluocinolone acetonide, as a therapy for DME in the United States. Iluvien has been approved in the European Union for the treatment of DME.

Our product candidates will also compete with anti-VEGF drugs, which are the current standard of care for RVO and wet AMD and have also been approved for DME. Genentech's products Lucentis and Avastin are both anti-VEGF antibodies. Lucentis is currently approved in the United States and European Union for the treatment of wet AMD and the treatment of macular edema following RVO and DME. Avastin is an anti-VEGF drug used off-label by retinal specialists in both the United States and in certain countries of the European Union in the treatment of numerous retinal diseases, but it is not indicated for ophthalmic use. In the United States, Regeneron's anti-VEGF product, Eylea, is approved for the treatment of wet AMD, macular edema following central retinal vein occlusion and DME. In the European Union, Eylea is approved for the treatment of wet AMD, DME and macular edema secondary to central retinal vein occlusion, and is being reviewed for the treatment of macular edema following branch retinal vein occlusion.

Ophthotech's drug candidate, Fovista, an anti-PDGF agent, is in Phase 3 clinical trials as a potential treatment for wet AMD to be administered together with an anti-VEGF drug. We are also aware of several other drug candidates in earlier stages of development as potential treatments for the diseases that we intend to target.

Many of our competitors have significantly greater financial resources and expertise in research and development, manufacturing, preclinical testing, conducting clinical trials, obtaining regulatory approvals and marketing approved drugs than we do. These competitors also compete with us in recruiting and retaining qualified scientific and management personnel and establishing clinical trial sites and patient recruitment for clinical trials, as well as in acquiring technologies complementary to, or necessary for, our

programs. Smaller or early stage companies may also prove to be significant competitors, particularly through collaborative arrangements with large and established companies.

Our commercial opportunity could be reduced or eliminated if our competitors develop and commercialize drugs that are safer, more effective, have fewer or less severe side effects, are more convenient or are less expensive than any drugs that we may develop. Our competitors may also obtain FDA or other regulatory approval for their drugs more rapidly than we may obtain approval for ours, which could result in our competitors establishing a strong market position before we are able to enter the market or obtaining FDA market exclusivity, thereby delaying our product approvals. Competitors' patent protection could also potentially delay FDA approval of our product candidates for up to 30 months, and we will still be subject to potential patent litigation that might arise beyond the 30 months. The key competitive factors affecting the success of our product candidates, if approved for marketing, are likely to be their efficacy, safety, method of administration, convenience, price and the availability of coverage and reimbursement from government and other third-party payors.

Intellectual Property

Our success depends in part on our ability to obtain and maintain patent and other intellectual property and proprietary protection for our technologies and methods of accessing the SCS, drug candidates and formulations as well as patent and other intellectual property and proprietary protection for novel applications, uses, and technological innovations related to our drug candidates, proprietary delivery devices and core technologies. We also rely on trade secrets, know-how, and continuing technological innovation to develop and maintain our proprietary position.

Patents and Patent Applications

Our patent estate, on a worldwide basis, includes four granted U.S. patents broadly directed to methods of administering drugs into the SCS by injection. In addition, our patent estate includes five patent applications pending in the United States, two issued foreign patents, four pending international PCT applications and 13 patent applications pending in major international markets, including the European Union, Canada, India, Japan, China and Australia, relating to our SCS delivery technology, as well as the formulations of our current therapeutic drug candidates and delivery device. Of these patents and patent applications, we license the four issued U.S. patents, one pending U.S. application, the two issued foreign patents, one of the pending international PCT applications and the 13 foreign patent applications, each relating to methods of delivering drugs to the eye by microneedle administration, pursuant to a license agreement with Georgia Tech Research Corporation and Emory University that is described below. With respect to the in-licensed international PCT application, according to the terms of the license agreement, we advise Georgia Tech and Emory regarding the countries in which patent applications should be pursued. Subject to payment of required maintenance fees, annuities and other charges, our issued patents are expected to expire between 2027 and 2029, without taking into account possible patent term adjustments or extensions. Applications relating to SCS delivery technology and methods, our current therapeutic drug candidates and formulations, various therapeutic uses, including treatment of specific indications such as macular edema associated with non-infectious uveitis and RVO, wet AMD as well as DME and improvement of specified clinical parameters, are expected to expire between 2027 and 2034, without taking into account possible patent term adjustments or extensions.

The term of individual patents depends on the legal term for patents in the countries in which they are granted. In most countries, including the United States, the patent term is generally 20 years from the

earliest claimed filing date of a non-provisional patent application in the applicable country. In the United States, a patent's term may, in certain cases, be extended by patent term adjustment, which compensates a patentee for administrative delays by the U.S. Patent and Trademark Office in examining and granting a patent, or may be shortened if a patent is terminally disclaimed over a commonly owned patent or a patent naming a common inventor and having an earlier expiration date. The Drug Price Competition and Patent Term Restoration Act of 1984, or the Hatch-Waxman Act, permits a patent term extension of up to five years beyond the expiration date of a U.S. patent as partial compensation for the length of time the drug is under regulatory review.

Our product candidates are protected against genericization through several levels of patents, including the patented approach of non-surgical administration into the SCS. In the case of CLS-TA injected into the SCS, an applicant who files a paragraph 4 ANDA or 505(b)(2) NDA certifying they may circumvent our patents must also demonstrate comparability of the proposed drug, which we believe may require a clinical trial in uveitic macular edema against our product, unless a biowaiver is obtained.

Third-Party Patent Filings

Numerous U.S. and foreign issued patents and patent applications owned by third parties exist in the fields in which we are developing products, including patents and applications related to surgical methods of accessing the SCS and certain formulations of TA. Because patent applications can take many years to issue, there may be applications unknown to us, which may later result in issued patents that our product candidates or proprietary technologies may infringe.

Under U.S. law, a person may be able to patent a discovery of a new way to use a previously known compound, even if such compound itself is patented, provided the newly discovered use is novel and nonobvious. Such a method-of-use patent, however, if valid, only protects the use of a claimed compound for the specified methods claimed in the patent. This type of patent does not prevent persons from using the compound for any previously known use of the compound. Further, this type of patent does not prevent persons from making and marketing the compound for an indication that is outside the scope of the patented method.

License Agreement with Emory and Georgia Tech

In July 2012, we entered into a license agreement with Emory University, or Emory, and the Georgia Tech Research Corporation, or GTRC, under which we received a worldwide exclusive license to specified patents relating to methods and devices for drug delivery using a microinjector. The field of the license covers all ophthalmic uses of the microinjector for mammals and birds. Under this license agreement, we have agreed to direct all prosecution of intellectual property to the licensors' patent counsel and have agreed to pay for all intellectual property expenses associated with the licensed patents.

Under this license agreement, we made an initial \$30,000 upfront payment and a \$35,000 milestone payment upon dosing of the first human patient in a clinical trial. This license agreement requires us to make a \$75,000 milestone payment upon the occurrence of a specified event relating to the commercialization of a drug developed using the licensed patents. Beginning on the third anniversary of this license agreement, we will be obligated to pay an annual license maintenance fee of \$25,000, which will continue until the first commercial sale of a product covered by the licensed patents. Additionally, we will owe a low single-digit royalty on any worldwide net product sales related to the licensed patents, with minimum annual royalties starting at \$15,000 per year after commercialization. The minimum annual

[Table of Contents](#)

royalty increases each year after the first commercial sale, up to a maximum amount of \$100,000 per year in the sixth calendar year and in subsequent years.

We are solely responsible for the development and commercialization of products related to the licensed patents. We are obligated to provide Emory and GTRC a written report detailing our activities related to the development and commercialization of products twice a year.

Royalties are calculated based on net sales of products covered by a patent licensed under the agreement, and are due on each such product sold as long as the patent covering that particular product is valid and unexpired. Unless otherwise terminated pursuant to its termination provisions, this license agreement will expire upon the expiration of the last to expire of the licensed patents. We have the right to terminate this license agreement at any time upon 60 days' written notice to Emory and GTRC. Emory and GTRC may also terminate this license agreement in the event of a material breach by us. This license agreement will immediately terminate if we challenge the validity or enforceability of any of the licensed patents in a court or other governmental agency of competent jurisdiction.

Trademarks, Trade Secrets and Know-How

Our trademark portfolio currently contains seven trademarks and trademark applications, consisting of two trademarks registered in the European Union, three pending U.S. trademark applications and two pending trademark applications in Canada. In addition to patent and trademark protection, we rely upon unpatented trade secrets and know-how and continuing technological innovation to develop and maintain our competitive position. We seek to protect our proprietary information, in part, using confidentiality agreements with our commercial partners, collaborators, employees, and consultants, and employees. These and other agreements, such as invention assignment agreements, grant us ownership of technologies that are developed through a relationship with a third party.

Government Regulation

In the United States, the FDA regulates drug and device products under the FDCA and its implementing regulations. Our product candidates are subject to regulation as combination products, which means that they are composed of both a drug product and device product. If marketed individually, each component would be subject to different regulatory pathways and reviewed by different Centers within the FDA. A combination product, however, is assigned to a Center that will have primary jurisdiction over its regulation based on a determination of the combination product's primary mode of action, which is the single mode of action that provides the most important therapeutic action. In the case of our product candidates, the primary mode of action is attributable to the drug component of the product, which means that the FDA's Center for Drug Evaluation and Research has primary jurisdiction over the premarket development, review and approval of our product candidates. Accordingly, we plan to investigate our products through the IND framework and seek approval through the NDA pathway. Based on our discussions with the FDA to date, we do not anticipate that the FDA will require a separate medical device authorization for the microinjector, but this could change during the course of its review of any marketing application that we may submit.

The process of obtaining regulatory approvals and the subsequent compliance with appropriate federal, state, local and foreign statutes and regulations requires the expenditure of substantial time and financial resources. Failure to comply with the applicable U.S. requirements at any time during the product development process, approval process or after approval may subject an applicant and/or sponsor to a

[Table of Contents](#)

variety of administrative or judicial sanctions, including refusal by FDA to approve applications, withdrawal of an approval, imposition of a clinical hold, import/export delays, issuance of warning letters and other types of letters, product recalls, product seizures, total or partial suspension of production or distribution, injunctions, fines, refusals of government contracts, restitution, disgorgement of profits, or civil or criminal investigations and penalties brought by FDA and the Department of Justice or other governmental entities.

Drugs

The clinical testing, manufacturing, labeling, storage, distribution, record keeping, advertising, promotion, import, export and marketing, among other things, of our product candidates are governed by extensive regulation by governmental authorities in the United States and other countries. The FDA, under the Federal Food, Drug, and Cosmetic Act, or FDCA, regulates pharmaceutical products in the United States. The steps required before a drug may be approved for marketing in the United States generally include:

- preclinical laboratory tests and animal tests conducted under Good Laboratory Practices, or GLP;
- the submission to the FDA of an IND application for human clinical testing, which must become effective before human clinical trials commence;
- adequate and well-controlled human clinical trials to establish the safety and efficacy of the product and conducted in accordance with Good Clinical Practices, or GCP;
- the submission to the FDA of an NDA;
- FDA acceptance, review and approval of the NDA, which might include an Advisory Committee review; and
- satisfactory completion of an FDA inspection of the manufacturing facilities at which the product is made to assess compliance with current Good Manufacturing Practices, or cGMPs.

The testing and approval process requires substantial time, effort and financial resources, and the receipt and timing of any approval is uncertain. The FDA may suspend clinical trials at any time on various grounds, including a finding that the subjects or patients are being exposed to an unacceptable health risk.

Preclinical studies include laboratory evaluations of the product candidate, as well as animal studies to assess the potential safety and efficacy of the product candidate. The results of the preclinical studies, together with manufacturing information and analytical data, are submitted to the FDA as part of the IND, which must become effective before clinical trials may be commenced. The IND will become effective automatically 30 days after receipt by the FDA, unless the FDA raises concerns or questions about the conduct of the trials as outlined in the IND prior to that time. In this case, the IND sponsor and the FDA must resolve any outstanding concerns before clinical trials can proceed. The FDA may nevertheless initiate a clinical hold after the 30 days if, for example, significant public health risks arise.

Each clinical trial must be reviewed and approved by an independent institutional review board, or IRB, at each of the sites at which the trial will be conducted. The IRB will consider, among other things, ethical factors, the safety of human subjects and the possible liability of the institution.

[Table of Contents](#)

Clinical trials are typically conducted in three sequential phases prior to approval, but the phases may overlap. These phases generally include the following:

Phase 1. Phase 1 clinical trials represent the initial introduction of a product candidate into human subjects, frequently healthy volunteers. In Phase 1, the product candidate is usually tested for safety, including adverse effects, dosage tolerance, absorption, distribution, metabolism, excretion and pharmacodynamics.

Phase 2. Phase 2 clinical trials usually involve studies in a limited patient population to (1) evaluate the efficacy of the product candidate for specific indications, (2) determine dosage tolerance and optimal dosage and (3) identify possible adverse effects and safety risks.

Phase 3. If a product candidate is found to be potentially effective and to have an acceptable safety profile in Phase 2 clinical trials, the clinical trial program will be expanded to Phase 3 clinical trials to further demonstrate clinical efficacy, optimal dosage and safety within an expanded patient population at geographically dispersed clinical trial sites.

Phase 4 clinical trials are conducted after approval to gain additional experience from the treatment of patients in the intended therapeutic indication and to document a clinical benefit in the case of drugs approved under accelerated approval regulations, or when otherwise requested by the FDA in the form of post-market requirements or commitments. Failure to promptly conduct any required Phase 4 clinical trials could result in withdrawal of approval.

The results of preclinical studies and clinical trials, together with detailed information on the manufacture, composition and quality of the product, are submitted to the FDA in the form of an NDA, requesting approval to market the product. The application must be accompanied by a significant user fee payment, although waivers may be granted in limited cases. The FDA has substantial discretion in the approval process and may refuse to accept any application or decide that the data is insufficient for approval and require additional preclinical, clinical or other studies.

Review of Application

Once an NDA has been accepted for filing, which occurs, if at all, 60 days after submission, the FDA sets a user fee goal date that informs the applicant of the specific date by which the FDA intends to complete its review. This is typically 10 months from the date of submission for drugs that are not new molecular entities, such as our product candidates. The review process can be extended by FDA requests for additional information or clarification. The FDA reviews NDAs to determine, among other things, whether the proposed product is safe and effective for its intended use, and whether the product is being manufactured in accordance with cGMPs to assure and preserve the product's identity, strength, quality and purity. Before approving an NDA, the FDA may inspect the facilities at which the product is manufactured and will not approve the product unless the manufacturing facility complies with cGMPs and may also inspect clinical trial sites for integrity of the data supporting safety and efficacy. The device component of our products will also be subject to review as part of the NDA review process and its manufacturers subject to inspection for compliance with device cGMPs embodied in the Quality System Regulation, or QSR. During the approval process, the FDA also will determine whether a risk evaluation and mitigation strategy, or REMS, is necessary to assure the safe use of the product. If the FDA concludes a REMS is needed, the sponsor of the application must submit a proposed REMS, and the FDA will not approve the application without an approved REMS, if required. A REMS can substantially increase the costs of obtaining approval.

The FDA could also require a special warning, known as a black box warning, to be included in the product label in order to highlight a particular safety risk. The FDA may also convene an advisory committee of external experts to provide input on certain review issues relating to risk, benefit and interpretation of clinical trial data. The FDA may delay approval of an NDA if applicable regulatory criteria are not satisfied and/or the FDA requires additional testing or information. The FDA may require post-marketing testing and surveillance to monitor safety or efficacy of a product. FDA will issue either an approval of the NDA or a Complete Response Letter, detailing the deficiencies and information required for reconsideration of the application.

Post-Approval Requirements

Approved drugs that are manufactured or distributed in the United States are subject to pervasive and continuing regulation by the FDA, including, among other things, requirements relating to recordkeeping, periodic reporting, product distribution, advertising and promotion and reporting of adverse experiences with the product. After approval, most changes to the approved product, such as adding new indications or other labeling claims and some manufacturing and supplier changes are subject to prior FDA review and approval. There also are continuing, annual user fee requirements for products and the establishments at which such products are manufactured, as well as new application fees for certain supplemental applications.

The FDA may impose a number of post-approval requirements as a condition of approval of an NDA. For example, the FDA may require post-marketing testing, including Phase 4 clinical trials, and surveillance to further assess and monitor the product's safety and effectiveness after commercialization. The FDA may also require a REMS, which could involve requirements for, among other things, a medication guide, special training for prescribers and dispensers, and patient registries.

In addition, entities involved in the manufacture and distribution of approved devices or drugs are required to register their establishments with the FDA and state agencies, and are subject to periodic unannounced inspections by the FDA and these state agencies for compliance with cGMP requirements. The FDA has promulgated specific requirements for drug cGMPs, device cGMPs embodied in the QSR and combination product cGMPs that, for drug/device combination products, specify the requirements within the drug cGMPs and the QSR with which manufacturers must comply. Changes to the manufacturing process are strictly regulated and often require prior FDA approval before being implemented. FDA regulations also require investigation and correction of any deviations from cGMP requirements and impose reporting and documentation requirements upon the sponsor and any third-party manufacturers that the sponsor may decide to use. Accordingly, manufacturers must continue to expend time, money, and effort in the area of production and quality control to maintain cGMP compliance.

Once an approval is granted, the FDA may withdraw the approval if compliance with regulatory requirements and standards is not maintained or if problems occur after the product reaches the market. Later discovery of previously unknown problems with a product, including adverse events of unanticipated severity or frequency, or with manufacturing processes, or failure to comply with regulatory requirements, may result in revisions to the approved labeling to add new safety information; imposition of post-market studies or clinical trials to assess new safety risks; or imposition of distribution or other restrictions under a REMS program. Other potential consequences include, among other things:

- restrictions on the marketing or manufacturing of the product, complete withdrawal of the product from the market or product recalls;

[Table of Contents](#)

- fines, warning letters or holds on post-approval clinical trials;
- refusal of the FDA to approve applications or supplements to approved applications, or suspension or revocation of product approvals;
- product seizure or detention, or refusal to permit the import or export of products; or
- injunctions or the imposition of civil or criminal penalties.

The FDA strictly regulates marketing, labeling, advertising and promotion of products that are placed on the market. Drugs may be promoted only for the approved indications and in accordance with the provisions of the approved label. The FDA and other agencies actively enforce the laws and regulations prohibiting the promotion of off-label uses, and a company that is found to have improperly promoted off-label uses may be subject to significant liability, including investigation by federal and state authorities.

The Hatch-Waxman Amendments

Our regulatory strategy is to pursue development of our drugs for suprachoroidal delivery as Section 505(b)(2) NDAs under the FDCA. As an alternative path to FDA approval for modifications to formulations or uses of drugs previously approved by the FDA, an applicant may submit an NDA under Section 505(b)(2) of the FDCA. Section 505(b)(2) was enacted as part of the Hatch-Waxman Amendments. A Section 505(b)(2) NDA is an application that contains full reports of investigations of safety and effectiveness, but where at least some of the information required for approval comes from studies not conducted by, or for, the applicant and for which the applicant has not obtained a right of reference or use from the person by or for whom the investigations were conducted. This type of application permits reliance for such approvals on literature or on an FDA finding of safety, effectiveness or both for an approved drug product. As such, under Section 505(b)(2), the FDA may rely, for approval of an NDA, on data not developed by the applicant. Therefore, if we can satisfy the conditions required for a Section 505(b)(2) NDA submission, it may eliminate the need for us to conduct some of the preclinical studies or clinical trials for the new product candidate that might otherwise have been required. The FDA may also require companies to perform additional studies or measurements, including clinical trials, to support the change from the approved branded reference drug. The FDA may then approve the new product candidate for all, or some, of the label indications for which the branded reference drug has been approved, as well as for any new indication sought by the 505(b)(2) applicant.

Orange Book Listing

In seeking approval for a drug through an NDA, including a 505(b)(2) NDA, applicants are required to list with the FDA certain patents whose claims cover the applicant's product. Upon approval of an NDA, each of the patents listed in the application for the drug is then published in the FDA's Approved Drug Products with Therapeutic Equivalence Evaluations, known as the Orange Book. Any applicant who files an Abbreviated New Drug Application, or ANDA, seeking approval of a generic equivalent version of a drug listed in the Orange Book or a 505(b)(2) NDA referencing a drug listed in the Orange Book must certify, for each patent listed in the Orange Book for the referenced drug, to the FDA that (1) no patent information on the drug product that is the subject of the application has been submitted to the FDA, (2) such patent has expired, (3) the date on which such patent expires or (4) such patent is invalid or will not be infringed upon by the manufacture, use or sale of the drug product for which the application is submitted. There might also be no relevant patent certification. The fourth certification described above is known as a paragraph IV

[Table of Contents](#)

certification. A notice of the paragraph IV certification must be provided to each owner of the patent that is the subject of the certification and to the holder of the approved NDA to which the ANDA or 505(b)(2) application refers. The applicant may also elect to submit a “section viii” statement certifying that its proposed label does not contain (or carves out) any language regarding the patented method-of-use rather than certify to a listed method-of-use patent. This section viii statement does not require notice to the patent holder or NDA owner.

If the reference NDA holder and patent owners assert a patent challenge directed to one of the Orange Book listed patents within 45 days of the receipt of the paragraph IV certification notice, the FDA is prohibited from approving the application until the earlier of 30 months from the receipt of the paragraph IV certification expiration of the patent, settlement of the lawsuit, or a decision in the infringement case that is favorable to the applicant. Even if the 45 days expire, a patent infringement lawsuit can be brought and could delay market entry, but it would not extend the FDA-related 30-month stay of approval.

The ANDA or 505(b)(2) application also will not be approved until any applicable non-patent exclusivity listed in the Orange Book for the branded reference drug has expired as described in further detail below.

Non-Patent Exclusivity

In addition to patent exclusivity, the holder of the NDA for the listed drug may be entitled to a period of non-patent exclusivity, during which the FDA cannot approve an ANDA or 505(b)(2) application that relies on the listed drug.

A drug, including one approved under a 505(b)(2) application, may obtain a three-year period of non-patent market exclusivity for a particular condition of approval, or change to a marketed product, such as a new formulation for a previously approved product, if one or more new clinical studies (other than bioavailability or bioequivalence studies) was essential to the approval of the application and was conducted/sponsored by the applicant. Should this occur, the FDA would be precluded from approving any ANDA or 505(b)(2) application for the protected modification until after that three-year exclusivity period has run. However, the FDA can accept an application and begin the review process during the three-year exclusivity period. A 505(b)(2) NDA may also be subject to a five-year exclusivity period for a new chemical entity, whereby the FDA will not accept for filing, with limited exception, a product seeking to rely upon the FDA’s findings of safety or effectiveness for such new chemical entity.

Orphan Drugs

Under the Orphan Drug Act, the FDA may grant orphan designation to a drug intended to treat a rare disease or condition affecting fewer than 200,000 individuals in the United States, or in other limited cases. Orphan drug designation does not convey any advantage in or shorten the duration of the regulatory review and approval process, though companies developing orphan drugs are eligible for certain incentives, including tax credits for qualified clinical testing. In addition, an NDA for a product that has received orphan drug designation is not subject to a prescription drug user fee unless the application includes an indication other than the rare disease or condition for which the drug was designated.

If a product that has orphan drug designation subsequently receives the first FDA approval for the disease or condition for which it has such designation, the product is entitled to orphan drug exclusivity, which means that the FDA may not approve any other applications to market the same active moiety for the

same indication for seven years, except in limited circumstances, such as another drug's showing of clinical superiority over the drug with orphan exclusivity. Competitors, however, may receive approval of different active moieties for the same indication or obtain approval for the same active moiety for a different indication. In addition, doctors may prescribe products for off-label uses and undermine our exclusivity. Orphan drug exclusivity could block the approval of one of our products for seven years if a competitor obtains approval for the same active moiety for the same indication before we do, unless we are able to demonstrate that our product is clinically superior.

We intend to seek orphan drug designation and exclusivity for CLS-1001 for the treatment of non-infectious uveitis in the United States and Europe, and we may seek designation for other products in the future, but we cannot guarantee that we will obtain orphan drug designation for any products in any jurisdiction. Even if we are able to obtain orphan drug designation for a product, we cannot be sure that such product will be approved, that we will be able to obtain orphan drug exclusivity upon approval, if ever, or that we will be able to maintain any exclusivity that is granted.

Foreign Regulation

In order to market any product outside of the United States, we would need to comply with numerous and varying regulatory requirements of other countries and jurisdictions regarding quality, safety and efficacy and governing, among other things, clinical trials, marketing authorization, commercial sales and distribution of our products. Whether or not we obtain FDA approval for a product, we would need to obtain the necessary approvals by the comparable foreign regulatory authorities before we can commence clinical trials or marketing of the product in foreign countries and jurisdictions. Although many of the issues discussed above with respect to the United States apply similarly in the context of the European Union and other geographies, the approval process varies between countries and jurisdictions and can involve additional product testing and additional administrative review periods. The time required to obtain approval in other countries and jurisdictions might differ from and be longer than that required to obtain FDA approval. Regulatory approval in one country or jurisdiction does not ensure regulatory approval in another, but a failure or delay in obtaining regulatory approval in one country or jurisdiction may negatively impact the regulatory process in others.

Federal and State Fraud and Abuse, Data Privacy and Security and Transparency Laws

In addition to FDA restrictions on marketing and promotion of drugs and devices, other federal and state laws restrict our business practices. These laws include, without limitation, anti-kickback and false claims laws, data privacy and security laws, as well as transparency laws regarding payments or other items of value provided to healthcare providers.

The federal Anti-Kickback Statute prohibits, among other things, knowingly and willfully offering, paying, soliciting or receiving any remuneration (including any kickback, bribe or rebate), directly or indirectly, overtly or covertly, to induce or in return for purchasing, leasing, ordering or arranging for or recommending the purchase, lease or order of any good, facility, item or service reimbursable, in whole or in part, under Medicare, Medicaid or other federal healthcare programs. The term "remuneration" has been broadly interpreted to include anything of value. Although there are a number of statutory exceptions and regulatory safe harbors protecting some common activities from prosecution, the exceptions and safe harbors are drawn narrowly. Practices that involve remuneration that may be alleged to be intended to induce prescribing, purchases or recommendations may be subject to scrutiny if they do not qualify for an

exception or safe harbor. Failure to meet all of the requirements of a particular applicable statutory exception or regulatory safe harbor does not make the conduct per se illegal under the Anti-Kickback Statute. Instead, the legality of the arrangement will be evaluated on a case-by-case basis based on a cumulative review of all its facts and circumstances. Several courts have interpreted the statute's intent requirement to mean that if any one purpose of an arrangement involving remuneration is to induce referrals of federal healthcare covered business, the Anti-Kickback Statute has been violated.

Additionally, the intent standard under the Anti-Kickback Statute was amended by the Patient Protection and Affordable Care Act of 2010, as amended by the Health Care and Education Reconciliation Act of 2010, collectively the Affordable Care Act, to a stricter standard such that a person or entity no longer needs to have actual knowledge of the statute or specific intent to violate it in order to have committed a violation. In addition, the Affordable Care Act codified case law that a claim including items or services resulting from a violation of the federal Anti-Kickback Statute constitutes a false or fraudulent claim for purposes of the federal civil False Claims Act.

The federal civil False Claims Act prohibits, among other things, any person or entity from knowingly presenting, or causing to be presented, a false or fraudulent claim for payment or approval to the federal government or knowingly making, using or causing to be made or used a false record or statement material to a false or fraudulent claim to the federal government. A claim includes "any request or demand" for money or property presented to the U.S. government. The civil False Claims Act also applies to false submissions that cause the government to be paid less than the amount to which it is entitled, such as a rebate. Intent to deceive is not required to establish liability under the civil False Claims Act. Several pharmaceutical, device and other healthcare companies have been prosecuted under these laws for, among other things, allegedly providing free product to customers with the expectation that the customers would bill federal programs for the product. Other companies have been prosecuted for causing false claims to be submitted because of the companies' marketing of products for unapproved, and thus non-covered, uses.

The government may further prosecute conduct constituting a false claim under the criminal False Claims Act. The criminal False Claims Act prohibits the making or presenting of a claim to the government knowing such claim to be false, fictitious, or fraudulent and, unlike the civil False Claims Act, requires proof of intent to submit a false claim.

The federal Health Insurance Portability and Accountability Act of 1996, or HIPAA, created new federal criminal statutes that prohibit among other actions, knowingly and willfully executing, or attempting to execute, a scheme to defraud any healthcare benefit program, including private third-party payors, knowingly and willfully embezzling or stealing from a healthcare benefit program, willfully obstructing a criminal investigation of a healthcare offense, and knowingly and willfully falsifying, concealing or covering up a material fact or making any materially false, fictitious or fraudulent statement in connection with the delivery of or payment for healthcare benefits, items or services. Like the Anti-Kickback Statute, the Affordable Care Act amended the intent standard for certain healthcare fraud statutes under HIPAA such that a person or entity no longer needs to have actual knowledge of the statute or specific intent to violate it in order to have committed a violation.

The civil monetary penalties statute imposes penalties against any person or entity that, among other things, is determined to have presented or caused to be presented a claim to a federal health program that the person knows or should know is for an item or service that was not provided as claimed or is false or fraudulent.

[Table of Contents](#)

Also, many states have similar fraud and abuse statutes or regulations that may be broader in scope and may apply regardless of payor, in addition to items and services reimbursed under Medicaid and other state programs.

Under HIPAA, the U.S. Department of Health and Human Services, or HHS, has issued regulations to protect the privacy and security of protected health information used or disclosed by covered entities including health care providers, such as us. HIPAA also regulates standardization of data content, codes and formats used in health care transactions and standardization of identifiers for health plans and providers. Penalties for violations of HIPAA regulations include civil and criminal penalties. In addition to federal privacy regulations, there are a number of state laws governing confidentiality and security of health information that are applicable to our business. New laws governing privacy may be adopted in the future as well. Failure to comply with privacy requirements could result in civil or criminal penalties, which could have a materially adverse effect on our business.

Additionally, there has been a recent trend of increased federal and state regulation of payments and transfers of value provided to healthcare professionals and/or entities. On February 8, 2013, the Centers for Medicare & Medicaid Services, or CMS, released its final rule implementing section 6002 of the Affordable Care Act known as the Physician Payment Sunshine Act that imposes new annual reporting requirements on manufacturers of drugs, devices, biologicals and medical supplies for payments and other transfers of value provided by them, directly or indirectly, to physicians and teaching hospitals, as well as ownership and investment interests held by physicians and their family members. A manufacturer's failure to submit timely, accurately and completely the required information for all payments, transfers of value or ownership or investment interests may result in civil monetary penalties of up to an aggregate of \$150,000 per year, and up to an aggregate of \$1 million per year for "knowing failures." Any failure to comply could result in significant fines and penalties. Certain states also mandate implementation of commercial compliance programs, impose restrictions on device manufacturer marketing practices or require tracking and reporting of gifts, compensation and other remuneration to particular types of healthcare professionals and entities.

Because of the breadth of these laws and the narrowness of the statutory exceptions and safe harbors available under such laws, it is possible that some of our current or future business activities, including certain sales and marketing practices and the provision of certain items and services to our customers, could be subject to challenge under one or more of such laws. The heightening compliance environment and the need to build and maintain robust and expandable systems to comply with different compliance and/or reporting requirements in multiple jurisdictions could increase the possibility that a healthcare company may fail to comply fully with one or more of these requirements. If our operations are found to be in violation of any of the health care laws or regulations described above that are applicable to us, or any other laws that apply to us, we may be subject to penalties, including potentially significant criminal and civil and/or administrative penalties, damages, fines, disgorgement, imprisonment, exclusion from participation in government healthcare programs, contractual damages, reputational harm, administrative burdens, diminished profits and future earnings, and the curtailment or restructuring of our operations, any of which could adversely affect our ability to operate our business and our results of operations. To the extent that any of our products are sold in a foreign country, we may be subject to similar foreign laws, which may include, for instance, applicable post-marketing requirements, including safety surveillance, anti-fraud and abuse laws and implementation of corporate compliance programs and reporting of payments or transfers of value to healthcare professionals.

Coverage and Reimbursement

We believe that physicians who use our product candidates, if approved, will be reimbursed by third-party payors for both the SCS injection using our microinjector and for the drug itself. We intend to seek a specific Current Procedural Terminology, or CPT, code as established and maintained by the American Medical Association, or AMA, for the SCS injection of pharmacologic agents using our microinjector, and a separate Healthcare Common Procedure Coding System, or HCPCS, code as maintained by the Centers for Medicare and Medicaid Services, or CMS, for the drug itself. We believe that separate CPT and HCPCS codes will help payors differentiate our drug candidates from other drugs currently on the market administered through intravitreal injections, although we can provide no assurance that the AMA or CMS will approve the creation of such new codes applicable to our products.

Our strategy will include efforts to engage physician societies and encourage third-party payors to establish coverage, coding and payment that will facilitate access to our product candidates and microinjector as we expand our commercialization efforts in the United States. Our success in these efforts depends in part on the extent to which governmental authorities, private health insurers and other third-party payors provide coverage for and establish adequate reimbursement levels for the procedures during which our product candidates are administered and for the drugs themselves. Failure by physicians, hospitals, ambulatory surgery centers, and other users of our products to obtain sufficient coverage and reimbursement from healthcare payors for the procedures administering our product candidates or for the product candidates themselves, or adverse changes in government and private third-party payors' policies would have a material adverse effect on our business, financial condition, results of operations and future growth prospects.

In the U.S., third-party payors continue to implement initiatives that restrict the use of certain technologies to those that meet certain clinical evidentiary requirements. Failure to obtain favorable coverage policies could have a material adverse effect on our business and operations.

In addition to uncertainties surrounding coverage policies, third-party payors from time to time update reimbursement amounts and also revise the methodologies used to determine reimbursement amounts. This includes annual updates to payments to ambulatory surgery centers, hospitals and physicians for the procedure performed administering our products, which could directly impact the demand for any of our product candidates that may be approved. An example of payment updates is the Medicare program updates to physician payments, which is done on an annual basis using a prescribed statutory formula. In the past, when the application of the formula resulted in lower payment, Congress has passed interim legislation to prevent the reductions. Most recently, the Protecting Access to Medicare Act of 2014, signed into law in April 2014, provided for a 0.5% change from 2013 payment rates under the Medicare Physician Fee Schedule through 2014 and a 0% update from January 1 until April 1, 2015. If Congress fails to intervene to prevent the negative update factor in future years, the resulting decrease in payment may adversely affect our revenues and results of operations. Any changes in coverage and reimbursement that further restricts coverage of our products or lowers reimbursement for procedures using our products could materially affect our business.

Healthcare Reform

Current and future legislative proposals to further reform healthcare or reduce healthcare costs may result in lower reimbursement for our products, or for the procedures associated with the use of our products, or limit coverage of our products. The cost containment measures that payors and providers are

instituting and the effect of any healthcare reform initiative implemented in the future could significantly reduce our revenues from the sale of our products. However, we believe that the shift away from fee-for-service agreements to capitated payment models supports the value of our products, as we believe that our products reduce longitudinal resource utilization, which can be cost saving-for both payors and providers.

For example, implementation of the Affordable Care Act has the potential to substantially change healthcare financing and delivery by both governmental and private insurers, and significantly impact the pharmaceutical and medical device industries.

The Affordable Care Act imposed, among other things, a new federal excise tax on the sale of certain medical devices, established an annual nondeductible fee on any entity that manufactures or imports certain specified branded prescription drugs and biologic agents, addressed a new methodology by which rebates owed by manufacturers under the Medicaid Drug Rebate Program are calculated for drugs that are inhaled, infused, instilled, implanted or injected, increased the minimum Medicaid rebates owed by most manufacturers under the Medicaid Drug Rebate Program, extended the Medicaid Drug Rebate program to utilization of prescriptions of individuals enrolled in Medicaid managed care organizations, and provided incentives to programs that increase the federal government's comparative effectiveness research. In addition, the Affordable Care Act implemented payment system reforms including a national pilot program on payment bundling to encourage hospitals, physicians and other providers to improve the coordination, quality and efficiency of certain healthcare services through bundled payment models.

In addition, other legislative changes have been proposed and adopted since the Affordable Care Act was enacted. On August 2, 2011, the President signed into law the Budget Control Act of 2011, which, among other things, created the Joint Select Committee on Deficit Reduction to recommend to Congress proposals in spending reductions. The Joint Select Committee did not achieve a targeted deficit reduction of at least \$1.2 trillion for the years 2013 through 2021, triggering the legislation's automatic reduction to several government programs. This includes reductions to Medicare payments to providers of 2% per fiscal year, which went into effect on April 1, 2013 and will remain in effect through 2024 unless additional Congressional action is taken. On January 2, 2013, President Obama signed into law the American Taxpayer Relief Act of 2012, which, among other things, reduced Medicare payments to several providers, including hospitals, imaging centers and cancer treatment centers and increased the statute of limitations period for the government to recover overpayments to providers from three to five years. We expect that additional state and federal healthcare reform measures will be adopted in the future, any of which could limit the amounts that federal and state governments will pay for healthcare products and services, which could result in reduced demand for our products or additional pricing pressure.

The Foreign Corrupt Practices Act

The Foreign Corrupt Practices Act, or the FCPA, prohibits any U.S. individual or business from paying, offering, or authorizing payment or offering of anything of value, directly or indirectly, to any foreign official, political party or candidate for the purpose of influencing any act or decision of the foreign entity in order to assist the individual or business in obtaining or retaining business. The FCPA also obligates companies whose securities are listed in the United States to comply with accounting provisions requiring the company to maintain books and records that accurately and fairly reflect all transactions of the corporation, including international subsidiaries, and to devise and maintain an adequate system of internal accounting controls for international operations. Activities that violate the FCPA, even if they occur wholly outside the United States, can result in criminal and civil fines, imprisonment, disgorgement, oversight, and debarment from government contracts.

Employees

As of June 30, 2014, we had 18 employees, all of whom are located in the United States. None of our employees is represented by a labor union or covered by a collective bargaining agreement. We consider our relationship with our employees to be good.

Facilities

Our principal offices occupy approximately 8,800 square feet of leased office space in Alpharetta, Georgia, pursuant to a lease agreement that expires in March 2017. We also lease approximately 6,600 square feet of space in Durham, North Carolina for our clinical development and regulatory functions under a lease agreement that expires in May 2015. We believe that our current facilities are suitable and adequate to meet our current needs. We intend to add new facilities or expand existing facilities as we add employees, and we believe that suitable additional or substitute space will be available as needed to accommodate any such expansion of our operations.

Legal Proceedings

We are not currently a party to any material legal proceedings.

MANAGEMENT

Directors and Executive Officers

The following table sets forth information concerning our directors, executive officers and other key employees, including their ages as of August 31, 2014:

<u>Name</u>	<u>Age</u>	<u>Position</u>
<i>Executive Officers:</i>		
Daniel H. White	47	President, Chief Executive Officer and Director
Charles A. Deignan	50	Chief Financial Officer
Glenn Noronha, Ph.D.	49	Executive Vice President, Research and Development
<i>Other Key Employees:</i>		
Rafael V. Andino	49	Vice President, Product Development
Stephen H. Lang	57	Vice President, Commercial Operations
<i>Non-Management Directors:</i>		
Christy L. Shaffer, Ph.D	56	Chairman of the Board of Directors
Clay B. Thorp	46	Director
Evgeny Zaytsev, M.D.	46	Director
Gerald D. Cagle, Ph.D.	70	Director
William D. Humphries	48	Director

Executive Officers

Daniel H. White

Mr. White is the founder of our company and has served as our President and Chief Executive Officer and as a member of our board of directors since our inception in May 2011. From 2008 to 2011, Mr. White served as Executive Director, Global Corporate Development, for Stiefel Laboratories, Inc., a dermatology pharmaceutical company acquired by GlaxoSmithKline in 2009. From 2007 to 2008, he co-founded and served as President and Chief Executive Officer of Percept BioScience, Inc., a biotechnology company. In 2003, Mr. White co-founded, and until 2007 served as Vice President of Finance and Corporate Development of Alimera Sciences, Inc., a biopharmaceutical company focused on ophthalmology. Previously, he was Head of Business Development and Licensing for CIBA Vision, a Novartis company, and Director of Licensing and Business Development for AAI Pharma. Mr. White holds an M.B.A. degree from Wake Forest University and a B.S. degree in Molecular Biology from Auburn University. Our board of directors believes that Mr. White's leadership of our company since its inception, extensive entrepreneurial experience, knowledge of our company as founder and experience with biotechnology companies prior to founding our company provides him with the qualifications and skills to serve as a director of our company.

Charles A. Deignan

Mr. Deignan has served as our Chief Financial Officer since January 2012. From 2009 to December 2011, Mr. Deignan was Vice President of Finance and Administration for Salutria Pharmaceuticals. Previously, from 1999 to 2009, Mr. Deignan served in a number of roles with AtheroGenics, Inc., a publicly held biopharmaceutical company, including as its Vice President of Finance and Administration. Prior to that, he held management positions at AAI Pharma, Inc. and Schering-Plough. Mr. Deignan received his B.S. degree in Business Administration from Boston University.

Glenn Noronha, Ph.D.

Dr. Noronha has served as our Executive Vice President, Research and Development since August 2013. From August 2012 to May 2013, he served as Vice President, Research and Development at Sucampo Pharma Americas, LLC, a pharmaceutical company. From July 2011 to July 2012, Dr. Noronha was Chief Scientific Officer for JW Theriac, Inc., a pharmaceutical company focused on new drug research and development. From 2008 to July 2011, Dr. Noronha was Global Project Head for Retinal Development at Alcon Laboratories, Inc., a Novartis company. From 2002 to 2008, Dr. Noronha held several positions at TargeGen, Inc., a pharmaceutical company, including as co-lead for its ophthalmology programs. Previously, from 2000 to 2002, he was a research scientist and project leader for Medtronic MiniMed, Inc. Dr. Noronha received his Ph.D. degree from Loyola University of Chicago and was a post-doctoral fellow at the University of California at Irvine.

Other Key Employees

Rafael V. Andino

Mr. Andino has served as our Vice President, Product Development since February 2013. Since June 2013, Mr. Andino has also served as an adjunct professor of biomedical engineering at the Georgia Institute of Technology. From January 2009 to February 2013, he served as Director of Research and Development for Bard Medical, a division of C.R. Bard, Inc., a publicly traded company. In 1999, Mr. Andino founded, and until 2008 served as President and Chief Executive Officer of Biofisica, Inc., a biotechnology company that developed technology to expedite wound healing and regeneration of connective tissue in humans. Prior to founding Biofisica, from 1995 to 2005 Mr. Andino was a mechanical engineer, research and development director and senior project manager for CIBA Vision, a Novartis company. He has also worked for the global technology companies E.I. Dupont de Nemours, General Electric and IBM. Mr. Andino received his Bachelor's degree in Mechanical Engineering from the Georgia Institute of Technology, a Master of Science in Biomedical Engineering from the University of Alabama at Birmingham and an M.B.A. degree from Mercer University.

Stephen H. Lang

Mr. Lang has served as our Vice President, Commercial Operations since March 2013. From 2006 to June 2012, Mr. Lang served as Vice President of Sales for ISTA Pharmaceuticals, Inc., an ophthalmic product company acquired by Bausch + Lomb in 2012. Previously, Mr. Lang held leadership roles in sales and marketing, serving with Novartis Ophthalmics, Inc. from 1995 to 2003, including as its Senior Vice President of U.S. Sales from 2002 to 2003. Mr. Lang began his career with Allergan, Inc. where he served for more than 16 years in various positions, including sales, marketing, national accounts and global business development. He earned his B.B.A. degree in Management from Georgia Southern University.

Non-Management Directors

Christy L. Shaffer, Ph.D.

Dr. Shaffer has served as a director of our company and as the chairman of our board of directors since January 2012. Since 2011, Dr. Shaffer has served as a Venture Partner with Hatteras Venture Partners, an investment firm, and as Managing Director of Hatteras Discovery, which invests in early-stage companies in the life sciences industry sector. From 1995 to March 2010, Dr. Shaffer served in increasing leadership positions at Inspire Pharmaceuticals, a publicly held biopharmaceutical company, beginning as the

[Table of Contents](#)

company's first full-time employee and Director of Clinical Operations and eventually being appointed as Chief Executive Officer and a director of the company in 1999, as its President in 2005 and a member of its Development Committee in 2009. Prior to Inspire, she was a clinical research scientist, international project leader and Associate Director of Pulmonary and Critical Care Medicine at Burroughs Wellcome Co. Dr. Shaffer currently serves in leadership roles on several non-profit boards, including as chair of the Morehead Planetarium and Science Center's advisory board, on the Board of Trustees for the Cystic Fibrosis Foundation, and as chair of the board of CFF Therapeutic, Inc. Dr. Shaffer is a receptor pharmacologist by training, earning her Ph.D. in Pharmacology from the University of Tennessee's Health Science Center in Memphis, Tennessee. She completed post-doctoral training at The Chicago Medical School as well as the University of North Carolina at Chapel Hill. In September 2008, the Securities and Exchange Commission approved a non-monetary settlement of its investigation relating to Inspire Pharmaceuticals' disclosures in its periodic reports relating to a clinical trial. The Commission also approved a settlement with Dr. Shaffer, as Inspire's President and Chief Executive Officer and a member of its board of directors, under which she consented to a cease and desist order against future violations of Section 13(a) of the Exchange Act and Rules 12b-20 and 13a-13 thereunder. The cease and desist order followed a finding by the Commission that three Quarterly Reports on Form 10-Q filed by Inspire included misleading disclosure about a clinical trial, specifically that the trial was described as "confirmatory" and "replicating" the efficacy found in an earlier trial. Dr. Shaffer did not admit or deny any findings in the order, and the order did not include any finding of any violation of any statute or regulation that involved any intentional wrongdoing or fraud, any monetary payments or other sanctions or otherwise affect Dr. Shaffer's future employment status, nor did it prohibit Dr. Shaffer from serving in any capacity on public company boards of directors. Our board of directors believes that Dr. Shaffer's clinical background as a scientist and her leadership experience as chief executive of a public company in the biopharmaceutical industry provides her with the qualifications and skills to serve as a director of our company.

Clay B. Thorp

Mr. Thorp has served as a director of our company since January 2012. In 2001, Mr. Thorp co-founded and has since served as General Partner of Hatteras Venture Partners, an investment firm, where he leads investments in a range of life science companies in the biopharmaceutical, medical device, diagnostics and research informatics sectors. Previously, he was instrumental in the founding of several life sciences companies, including serving as co-founder, Chief Executive Officer and Chairman of Synthematix, Inc., a chemistry informatics company that was acquired by Symyx Technologies in 2005, co-founder and former Chairman of PhaseBio Pharmaceuticals, Inc., co-founder and head of corporate development for Novalon Pharmaceutical Corporation, which was sold to Karo Bio in 2000, and co-founder and president of Xanthon, Inc., a bioinformatics company with electro-chemical detection technology for direct analysis of DNA, RNA and proteins. Mr. Thorp holds a Masters of Public Policy degree from Harvard University and a B.A. degree in Mathematics and History from the University of North Carolina at Chapel Hill. Our board of directors believes that Mr. Thorp's experience as an entrepreneur and an investor in life sciences companies provides him with the qualifications and skills to serve as a director of our company.

Evgeny Zaytsev, M.D.

Dr. Zaytsev has served as a director of our company since August 2014. Dr. Zaytsev has served as President, Chief Executive Officer and a managing partner of RMI Partners LLC since May 2013 and as the President and Chief Executive Officer of RMI Partners Inc. since November 2013. Dr. Zaytsev has also served as a general partner at Helix Ventures, which he co-founded to exclusively invest in novel

[Table of Contents](#)

therapeutic opportunities, since July 2009. Previously Dr. Zaytsev was a partner at Asset Management Company, one of the oldest venture firms in Silicon Valley, from 2002 to 2009. Dr. Zaytsev received his M.D. degree and Ph.D. degree from the Altai State Medical University and his M.B.A. degree from the Stanford Graduate School of Business. Our board of directors believes that Dr. Zaytsev's scientific background and experience as an investor in life sciences companies provides him with the qualifications and skills to serve as a director of our company.

Gerald D. Cagle, Ph.D.

Dr. Cagle has served on our board of directors since July 2013. Dr. Cagle has served as Chief Operating Officer at Cognoptix, Inc., a biotechnology company focused on the diagnosis of Alzheimer's disease, since December 2008. He also is Senior Advisor and Head of Business Development for GrayBug, LLC, a platform drug delivery company. Previously, Dr. Cagle served as Senior Vice President of Research & Development at Alcon Laboratories Inc. from 1997 to 2008, assuming the responsibility of Chief Scientific Officer in 2006. He currently serves on the board of directors of Aerie Pharmaceuticals, Inc., a publicly held biopharmaceutical company. Dr. Cagle has also served on the Wilmer Eye Institute Advisory Council and is a member of the ARVO Foundation Board of Governors. Dr. Cagle received his B.S. degree from Wayland College and earned M.S. and Ph.D. degrees from the University of North Texas. Our board of directors believes that Dr. Cagle's scientific background and experience provides him with the qualifications and skills to serve as a director of our company.

William D. Humphries

Mr. Humphries has served as a director of our company since January 2012. Mr. Humphries has served as President and Chief Executive Officer of the North American business of Merz, Inc., an affiliate of Merz Pharma Group, a specialty healthcare company, since March 2012. From 2006 to March 2012, he served in a number of leadership positions with Stiefel Laboratories, Inc., a dermatology pharmaceutical company, including as its Chief Commercial Officer and then as its President beginning in 2008. Stiefel was acquired by GlaxoSmithKline in 2009. After the acquisition, Mr. Humphries served as the President of Dermatology for Stiefel from 2009 until March 2012. Before Stiefel, Mr. Humphries served in executive roles in sales and marketing, business development, and international marketing for Allergan, Inc., concluding as vice president of its U.S. skincare business. Mr. Humphries has served on the board of ZARS Pharma, the GlaxoSmithKline Portfolio Investment Board and the GlaxoSmithKline Ophthalmology Board. Mr. Humphries received his M.B.A. degree from Pepperdine University and a B.A. degree from Bucknell University. Our board of directors believes that Mr. Humphries' experience as pharmaceutical company executive provides him with the qualifications and skills to serve as a director of our company.

Board Composition

Our board of directors currently consists of six members. Each director is currently elected to the board for a one-year term, to serve until the election and qualification of successor directors at the annual meeting of stockholders, or until the director's earlier removal, resignation or death.

Our directors were elected to and currently serve on the board pursuant to a voting agreement among us and several of our largest stockholders. This agreement will terminate upon the completion of this offering, after which there will be no further contractual obligations regarding the election of our directors.

In accordance with our amended and restated certificate of incorporation, which will be in effect upon the closing of this offering, our board of directors will be divided into three classes, each of which will

Table of Contents

consist, as nearly as possible, of one-third of the total number of directors constituting our entire board and which will serve staggered three-year terms. At each annual meeting of stockholders, the successors to directors whose terms then expire will be elected to serve from the time of election and qualification until the third annual meeting following election. Our directors will be divided among the three classes as follows:

- Class I, which will consist of _____ and _____, and their term will expire at our first annual meeting of stockholders to be held after the completion of this offering;
- Class II, which will consist of _____ and _____, and their term will expire at our second annual meeting of stockholders to be held after the completion of this offering; and
- Class III, which will consist of _____ and _____, and their term will expire at our third annual meeting of stockholders to be held after the completion of this offering.

Our amended and restated bylaws, which will become effective upon completion of this offering, will provide that the authorized number of directors may be changed only by resolution approved by a majority of our board of directors. Any additional directorships resulting from an increase in the number of directors will be distributed among the three classes so that, as nearly as possible, each class will consist of one-third of the directors.

The division of our board of directors into three classes with staggered three-year terms may delay or prevent a change of our management or a change in control.

Director Independence

Our board of directors has undertaken a review of the independence of the directors and considered whether any director has a material relationship with us that could compromise his or her ability to exercise independent judgment in carrying out his or her responsibilities. As a result of this review, our board of directors has determined that Drs. Shaffer, Cagle and Zaytsev and Messrs. Humphries and Thorp, representing five of our six directors, are “independent directors” as defined under NASDAQ rules.

Committees of the Board of Directors

Our board of directors has established an audit committee, a compensation committee and a nominating and corporate governance committee, each of which has the composition and responsibilities described below. From time to time, the board may establish other committees to facilitate the management of our business.

Audit Committee

Our audit committee reviews our internal accounting procedures and consults with and reviews the services provided by our independent registered public accountants. Our audit committee consists of three directors, _____, _____ and _____, and our board of directors has determined that each of them is independent within the meaning of the applicable stock exchange listing requirements and the independence requirements contemplated by Rule 10A-3 under the Securities Exchange Act of 1934, as amended, or the Exchange Act. _____ is the chairman of the audit committee and our board of directors has determined that _____ is an “audit committee financial expert” as defined by SEC rules and regulations. Our board of directors has determined that the composition of our audit committee meets the criteria for independence under, and the functioning of our audit committee complies with, the

[Table of Contents](#)

applicable requirements of the Sarbanes-Oxley Act, applicable stock exchange listing requirements and SEC rules and regulations. We intend to continue to evaluate the requirements applicable to us and we intend to comply with the future requirements to the extent that they become applicable to our audit committee. The principal duties and responsibilities of our audit committee include:

- appointing and retaining an independent registered public accounting firm to serve as independent auditor to audit our financial statements, overseeing the independent auditor's work and determining the independent auditor's compensation;
- approving in advance all audit services and non-audit services to be provided to us by our independent auditor;
- establishing procedures for the receipt, retention and treatment of complaints received by us regarding accounting, internal accounting controls, auditing or compliance matters, as well as for the confidential, anonymous submission by our employees of concerns regarding questionable accounting or auditing matters;
- reviewing and discussing with management and our independent auditor the results of the annual audit and the independent auditor's review of our quarterly financial statements; and
- conferring with management and our independent auditor about the scope, adequacy and effectiveness of our internal accounting controls, the objectivity of our financial reporting and our accounting policies and practices.

Compensation Committee

Our compensation committee reviews and determines the compensation of all our executive officers. Our compensation committee consists of directors, _____, each of whom is a non-employee member of our board of directors as defined in Rule 16b-3 under the Exchange Act. _____ is the chairman of the compensation committee. Our board of directors has determined that the composition of our compensation committee satisfies the applicable independence requirements under, and the functioning of our compensation committee complies with the applicable requirements of, stock exchange listing rules and SEC rules and regulations. We intend to continue to evaluate and intend to comply with all future requirements applicable to our compensation committee. The principal duties and responsibilities of our compensation committee include:

- establishing and approving, and making recommendations to the board of directors regarding, performance goals and objectives relevant to the compensation of our chief executive officer, evaluating the performance of our chief executive officer in light of those goals and objectives and setting, or recommending to the full board of directors for approval, the chief executive officer's compensation, including incentive-based and equity-based compensation, based on that evaluation;
- setting the compensation of our other executive officers, based in part on recommendations of the chief executive officer;
- exercising administrative authority under our stock plans and employee benefit plans;
- establishing policies and making recommendations to our board of directors regarding director compensation;
- reviewing and discussing with management the compensation discussion and analysis that we may be required from time to time to include in SEC filings; and

[Table of Contents](#)

- preparing a compensation committee report on executive compensation as may be required from time to time to be included in our annual proxy statements or annual reports on Form 10-K filed with the SEC.

Nominating and Corporate Governance Committee

The nominating and corporate governance committee consists of _____ directors, _____ is the chairman of the nominating and corporate governance committee. Our board of directors has determined that the composition of our nominating and corporate governance committee satisfies the applicable independence requirements under, and the functioning of our nominating and corporate governance committee complies with the applicable requirements of, stock exchange listing standards and SEC rules and regulations. We will continue to evaluate and will comply with all future requirements applicable to our nominating and corporate governance committee. The nominating and corporate governance committee's responsibilities include:

- assessing the need for new directors and identifying individuals qualified to become directors;
- recommending to the board of directors the persons to be nominated for election as directors and to each of the board's committees;
- assessing individual director performance, participation and qualifications;
- developing and recommending to the board corporate governance principles;
- monitoring the effectiveness of the board and the quality of the relationship between management and the board; and
- overseeing an annual evaluation of the board's performance.

Code of Business Conduct and Ethics for Employees, Executive Officers and Directors

Effective upon completion of this offering, we will adopt a Code of Business Conduct and Ethics, or the Code of Conduct, applicable to all of our employees, executive officers and directors. Following the completion of this offering, the Code of Conduct will be available on our website at www.clearsidebio.com. The nominating and corporate governance committee of our board of directors will be responsible for overseeing the Code of Conduct and must approve any waivers of the Code of Conduct for employees, executive officers and directors. We expect that any amendments to the Code of Conduct, or any waivers of its requirements, will be disclosed on our website.

Compensation Committee Interlocks and Insider Participation

None of our directors who currently serve as members of our compensation committee is, or has at any time during the past year been, one of our officers or employees. None of our executive officers currently serves, or in the past year has served, as a member of the board of directors or compensation committee of any other entity that has one or more executive officers serving on our board of directors or compensation committee.

Non-Employee Director Compensation

We have not historically paid cash retainers or other compensation with respect to service on our board of directors, except for reimbursement of direct expenses incurred in connection with attending meetings of

[Table of Contents](#)

the board or committees. In January 2012, we awarded an option to purchase 50,000 shares of our common stock to Mr. Humphries at an exercise price of \$0.07 per share. In October 2013, we awarded an option to purchase 50,000 shares of our common stock to Mr. Humphries at an exercise price of \$0.18 per share. In June 2013, in connection with his appointment to our board of directors, we awarded an option to purchase 100,000 shares of our common stock to Dr. Cagle at an exercise price of \$0.18 per share. Other than Mr. Humphries and Mr. Cagle, none of our non-employee directors serving as of December 31, 2013 held any options to purchase our common stock.

The following table sets forth information regarding compensation earned for service on our board of directors during the year ended December 31, 2013 by our non-employee directors. Mr. White, our President and Chief Executive Officer, is also a director but does not receive any additional compensation for his service as director. Mr. White's compensation as an executive officer is set forth below under "Executive Compensation — Summary Compensation Table."

<u>Name</u>	<u>Stock Awards (\$)</u>	<u>Option Awards (\$)(1)</u>
Gerald D. Cagle, Ph.D.	—	44,900(2)
William D. Humphries	—	29,800(3)
Mark R. Prausnitz, Ph.D.	—(4)	—
Christy L. Shaffer, Ph.D.	—	—
Clay B. Thorp	—	—

- (1) This column reflects the full grant date fair value for options granted during the year as measured pursuant to ASC Topic 718 as stock-based compensation in our consolidated financial statements. Unlike the calculations contained in our financial statements, this calculation does not give effect to any estimate of forfeitures related to service-based vesting but assumes that the director will perform the requisite service for the award to vest in full. The assumptions we used in valuing options are described in Note 11 to our audited consolidated financial statements included in this prospectus.
- (2) As of December 31, 2013, Dr. Cagle held an option to purchase 100,000 shares of common stock, which vests in 16 equal quarterly installments at the end of each calendar quarter beginning September 30, 2013 and is subject to full acceleration of vesting upon a change of control of our company.
- (3) As of December 31, 2013, Mr. Humphries held options to purchase 100,000 shares of common stock, of which 50,000 shares were vested and the remainder vests in eight equal quarterly installments at the end of each calendar quarter beginning March 31, 2014 and are subject to full acceleration of vesting upon a change of control of our company.
- (4) As a founder of our company, Dr. Prausnitz acquired 617,500 shares of common stock from us in August 2011, which shares were subject to a right of repurchase in favor of us through June 30, 2014. Dr. Prausnitz resigned from our board of directors in August 2014. As of December 31, 2013, there were 77,188 shares not yet vested.

We expect that our board of directors will adopt a director compensation policy for non-employee directors to be effective following the completion of this offering.

EXECUTIVE COMPENSATION

Our named executive officers for the year ended December 31, 2013 include our principal executive officer and our two other executive officers:

- Daniel H. White, our President and Chief Executive Officer;
- Charles A. Deignan, our Chief Financial Officer; and
- Glenn Noronha, Ph.D., our Executive Vice President of Research and Development.

No other individuals served as executive officers of our company at any point during 2013.

2013 Summary Compensation Table

The following table presents the compensation awarded to, earned by or paid to each of our named executive officers for the year ended December 31, 2013.

<u>Name and Principal Position</u>	<u>Salary \$(1)</u>	<u>Bonus \$(2)</u>	<u>Option Awards \$(3)</u>	<u>Non-Equity Incentive Plan Compensation (\$)</u>	<u>All Other Compensation (\$)</u>	<u>Total (\$)</u>
Daniel H. White President and Chief Executive Officer	256,771	12,906	102,225	51,625	—	423,527
Charles A. Deignan Chief Financial Officer	179,740	—	68,025	28,910	—	276,675
Glenn Noronha, Ph.D. Executive Vice President, Research and Development(4)	108,333	—	105,250	17,333	48,809(5)	279,725

- (1) Salary amounts represent actual amounts paid during 2013. See “—Narrative to Summary Compensation Table—Annual Base Salary” for a description of adjustments to base salaries made during the year.
- (2) Represents the amount above the specified level of achievement under the annual bonus incentive plan. The compensation committee exercised its discretion to award Mr. White additional compensation in light of his role in the achievement of corporate objectives outside of the scope of the stated objectives described below under “Narrative to Summary Compensation Table—Annual Bonus.”
- (3) The amounts reflect the full grant date fair value for awards granted during 2013. The grant date fair value was computed in accordance with ASC Topic 718, *Compensation — Stock Compensation*. Unlike the calculations contained in our financial statements, this calculation does not give effect to any estimate of forfeitures related to service-based vesting, but assumes that the executive will perform the requisite service for the award to vest in full. The assumptions we used in valuing options are described in note 11 to our financial statements included in this prospectus.
- (4) Dr. Noronha became an executive officer of our company in August 2013, and amounts represent compensation earned since that date.
- (5) Amount represents relocation allowance plus associated tax gross-up.

Narrative to Summary Compensation Table

We review compensation annually for all employees, including our executives. In setting executive base salaries and bonuses and granting equity incentive awards, we consider compensation for comparable positions in the market, the historical compensation levels of our executives, individual performance as compared to our expectations and objectives, our desire to motivate our employees to achieve short- and long-term results that are in the best interests of our stockholders, and a long-term commitment to our company. We do not target a specific competitive position or a specific mix of compensation among base salary, bonus or long-term incentives.

The compensation committee of our board of directors has historically determined our executives' compensation. Our compensation committee typically reviews and discusses management's proposed compensation with the chief executive officer for all executives other than the chief executive officer. Based on those discussions and its discretion, the compensation committee then recommends the compensation for each executive officer. Our compensation committee, without members of management present, discusses and ultimately approves the compensation of our executive officers. To date, our compensation committee has not engaged a compensation consultant or adopted a peer group of companies for purposes of determining executive compensation.

Annual Base Salary

Our named executive officers' base salaries are reviewed periodically by our board of directors, and adjustments may be made upon the recommendations of the compensation committee. In 2012, we entered into an employment agreement with Mr. White under which his annual base salary was established at \$250,000. In February 2013, our compensation committee recommended, and our board of directors approved, an increase in Mr. White's annual base salary to \$258,125. At the same time, our compensation committee recommended, and our board of directors approved, an increase in Mr. Deignan's base salary from \$175,000 to \$180,688. In July 2014, our compensation committee recommended, and the board of directors approved, further increases in our named executive officers' salaries as follows: \$268,450 for Mr. White, \$215,722 for Mr. Deignan and \$266,500 for Dr. Noronha. Dr. Noronha's annual base salary of \$260,000 was approved by our board of directors in connection with the commencement of his employment with us in August 2013.

Annual Bonus

We seek to motivate and reward our executives for achievements relative to our corporate goals and expectations for each fiscal year. Each named executive officer has a target bonus opportunity, defined as a percentage of his annual salary. For 2013 and 2014, the target bonus was and is as follows:

<u>Name</u>	<u>Target Bonus (% of Salary)</u>
Daniel H. White	25
Charles A. Deignan	20
Glenn Noronha, Ph.D.	20

To reinforce the importance of integrated and collaborative leadership, our executives' bonuses have historically been solely based on company performance, and we did not include an individual performance component. For 2013, 40% of each executive officer's target bonus was attributable to the commencement of our Phase 1/2 clinical trial for CLS-1001, 30% was attributable to our further drug and product development efforts and 30% was attributable to our capital raising activities. In 2014, our compensation

[Table of Contents](#)

committee determined that the performance goals had been achieved at an 80% level in the aggregate. The bonuses to be paid to the named executive officers for 2013 performance at the 80% level are reflected in the “Non-Equity Incentive Plan Compensation” column of the Summary Compensation Table above.

Long-Term Incentives

Our 2011 Stock Incentive Plan authorizes us to make grants to eligible recipients of non-qualified stock options, incentive stock options, restricted stock awards, restricted stock units and other stock-based awards. All of our awards under this plan have been in the form of stock options.

We typically grant stock options at the start of employment to each executive and our other employees. Through 2013, we have not maintained a practice of granting additional equity on an annual basis, but we have retained discretion to provide additional targeted grants in appropriate circumstances.

We award stock options on the date the compensation committee approves the grant. We set the option exercise price and grant date fair value based on our per-share valuation on the date of grant.

In February 2013, our compensation committee awarded options to Messrs. White and Deignan to purchase 128,000 shares and 52,000 shares, respectively. Each of these options has an exercise price of \$0.18 per share. In August 2013, in connection with the commencement of Dr. Noronha’s employment with us, our board of directors approved the grant of an option to purchase 125,000 shares of our common stock with an exercise price of \$0.18 per share. In November 2013, our board of directors approved additional options grants to Messrs. White and Deignan and Dr. Noronha to purchase 75,000 shares, 75,000 shares and 50,000 shares, respectively.

Employment Arrangements and Potential Payments upon Termination of Employment

In September 2012, we entered into an employment agreement with Mr. White under which he serves as our President and Chief Executive Officer. The agreement is renewable for one-year terms beginning in September 2014, unless either we or Dr. White give written notice of non-renewal at least 60 days prior to the end of the term. Under this agreement, Mr. White is eligible to receive severance benefits in specified circumstances.

In the event that we terminate Mr. White without cause, he resigns for good reason or we elect not to renew the term of the employment agreement, then, upon execution and effectiveness of a settlement agreement and release of claims in a form acceptable to us, Mr. White will be entitled to receive (a) an amount equal to nine months of his annual base salary, less applicable deductions, payable in accordance with our normal payroll schedule, (b) reimbursement of the cost of health insurance premiums for 12 months or, if shorter, until he obtains reasonably comparable health insurance coverage.

The following definitions have been adopted in Mr. White’s employment agreement:

- “cause” means that our board of directors, without Mr. White, has determined that any of the following occurred: (a) Mr. White’s material breach of the employment agreement, his failure to diligently and properly perform his duties or his failure to achieve the objectives specified by the board of directors, and any such breach or failure has not been cured within 30 days after written notice thereof, (b) his misappropriation or unauthorized use of our property or breach of his agreements with us relating to confidentiality, intellectual property rights, non-competition or non-solicitation, (c) his material failure to comply with our company policies or directives of our board of directors, and any such failure has not been cured within 30 days after written notice thereof,

[Table of Contents](#)

provided that the failure to comply with company policies relating to harassment, unlawful discrimination, retaliation or workplace violence do not require notice or permit a cure period, (d) his use of illegal drugs or any illegal substance, or his use of alcohol in any manner that materially interferes with the performance of his duties to the company, (e) any dishonest or illegal action by him, or any action determined to be materially detrimental to the interest and well-being of our company, including harm to our reputation, (f) his failure to fully disclose any material conflict of interest that he may have in a transaction between us and a third party, which conflict is materially detrimental to our interest and well-being, or (g) any adverse action or omission by him that would be required to be disclosed under securities laws or which would limit our ability to sell securities or would disqualify us from an exemption otherwise available to us; and

- “good reason” means the existence of any of the following without Mr. White’s prior consent: (a) any substantial reduction or diminution of his duties and responsibilities or salary, (b) any material breach of the employment agreement by us, or (c) a relocation of his place of employment by more than 50 miles from the location of our principal office, in each case after notice to us within 90 days following the initial existence of the event and after we have had the opportunity to but have not cured the event for 30 days following such notice, and the executive terminates his employment with us no later than two years after the initial existence of the event.

Outstanding Equity Awards at End of 2013

The following table provides information about outstanding stock options held by each of our executive officers at December 31, 2013. All of these options were granted under our 2011 Stock Incentive Plan.

Name	Option Awards				Stock Awards	
	Number of Securities Underlying Unexercised Options (#) Exercisable	Number of Securities Underlying Unexercised Options (#) Unexercisable	Option Exercise Price (\$)	Option Expiration Date	Number of Shares or Units of Stock That Have Not Vested (#)	Market Value of Shares or Units of Stock That Have Not Vested (\$)(8)
Daniel H. White	—	128,000(1) 75,000(3)	0.18 0.18	02/28/2023 11/24/2023	118,750(4)	78,375
Charles A. Deignan	—	52,000(1) 75,000(3)	0.18 0.18	02/28/2023 11/24/2023	20,833(5) 38,750(6) 17,188(7)	13,750 25,575 11,344
Glenn Noronha, Ph.D.	—	125,000(2) 50,000(3)	0.18 0.18	08/06/2023 11/24/2023		

(1) The unvested shares underlying this option vest as to 25% of the shares on March 1, 2014, with the remainder vesting in 36 equal monthly installments thereafter, subject to the officer’s continued service through each applicable vesting date. The option is subject to full acceleration of vesting following a change of control if the option is assumed or a substantially equivalent award is substituted, but the officer’s employment is terminated without cause within 12 months following the change of control.

(2) The unvested shares underlying this option vest as to 25% of the shares on August 1, 2014, with the remainder vesting in 36 equal monthly installments thereafter, subject to the officer’s continued service through each applicable vesting date. The option is subject to full acceleration of vesting (a) upon a change of control if the option is not assumed or a substantially equivalent award is not substituted by

[Table of Contents](#)

the acquiring or succeeding entity or (b) following a change of control if the option is assumed or a substantially equivalent award is substituted, but the officer's employment is terminated without cause within 12 months following the change of control.

- (3) The invested shares underlying this option vest as to 25% of the shares on December 13, 2014, with the remainder vesting in 36 equal monthly installments thereafter, subject to the officer's continued service through each applicable vesting date. The option is subject to full acceleration of vesting (a) upon a change of control if the option is not assumed or a substantially equivalent award is not substituted by the acquiring or succeeding entity or (b) following a change of control if the option is assumed or a substantially equivalent award is substituted, but the officer's employment is terminated without cause or for good reason within 12 months following the change of control.
- (4) These restricted shares will vest in equal monthly installments through June 30, 2014. These restricted shares are subject to full acceleration of vesting upon a change of control or in the event the officer's employment is terminated without cause.
- (5) These restricted shares will vest in equal monthly installments through January 31, 2016. These restricted shares are subject to full acceleration of vesting (a) upon a change of control if the option is not assumed or a substantially equivalent award is not substituted by the acquiring or succeeding entity or (b) following a change of control if the option is assumed or a substantially equivalent award is substituted, but the officer's employment is terminated without cause or for good reason within 12 months following the change of control.
- (6) These restricted shares will vest in equal monthly installments through July 1, 2016. These restricted shares are subject to full acceleration of vesting (a) upon a change of control if the option is not assumed or a substantially equivalent award is not substituted by the acquiring or succeeding entity or (b) following a change of control if the option is assumed or a substantially equivalent award is substituted, but the officer's employment is terminated without cause or for good reason within 12 months following the change of control.
- (7) These restricted shares will vest in equal monthly installments through September 30, 2016. These restricted shares are subject to full acceleration of vesting (a) upon a change of control if the option is not assumed or a substantially equivalent award is not substituted by the acquiring or succeeding entity or (b) following a change of control if the option is assumed or a substantially equivalent award is substituted, but the officer's employment is terminated without cause or for good reason within 12 months following the change of control.
- (8) Based on the valuation of our common stock of \$0.66 per share as of December 31, 2013.

Stock Option Exercises and Stock Vested During 2013

The following table shows information regarding options exercised and stock awards vested during the year ended December 31, 2013 with respect to our named executive officers.

Name	Option Awards		Stock Awards	
	Number of Shares Acquired on Exercise (#)	Value Realized on Exercise (\$)	Number of Shares Acquired on Vesting (#)	Value Realized on Vesting (\$)
Daniel H. White	—	—	237,500	137,750(2)
Charles A. Deignan	46,875	27,655(1)	1,354(3)	799(4)

- (1) The aggregate dollar amount represents the amount by which the aggregate fair value of the shares of our common stock on the date of exercise, as calculated using an assumed per share value of \$0.66,

[Table of Contents](#)

which is the assumed fair value as of the date of exercise, exceeds the aggregate exercise price of the option, as calculated using the per share exercise price of \$0.07.

- (2) For shares that vested during the first six months of 2013, we have determined the value realized by Mr. White using an assumed value of \$0.50 per share. For shares vested during the last six months of 2013, we have determined the value of such shares using an assumed value of \$0.66 per share.
- (3) Mr. Deignan acquired shares of stock pursuant to the early exercise of stock options, which stock was subject to a right of repurchase in favor of us. The number of shares acquired on vesting represents the number of such shares for which our rights of repurchase lapsed during the year.
- (4) Represents the difference between the estimated value of the shares for which our repurchase right lapsed and the exercise price of \$0.07 per share paid by Mr. Deignan in respect of such shares upon the early exercise of the options. The assumed value of such shares upon vesting was \$0.66 per share.

Health and Welfare Benefits

We maintain a defined contribution employee retirement plan for our employees. Our 401(k) plan is intended to qualify as a tax-qualified plan under Section 401 of the Internal Revenue Code so that contributions to our 401(k) plan, and income earned on such contributions, are not taxable to participants until withdrawn or distributed from the 401(k) plan. Our 401(k) plan provides that each participant may contribute a portion of his or her pre-tax compensation, up to a statutory limit, which is \$17,500 for 2014. Participants who are at least 50 years old can also make “catch-up” contributions, which in 2014 may be up to an additional \$5,500 above the statutory limit. Under our 401(k) plan, each employee is fully vested in his or her deferred salary contributions. Employee contributions are held and invested by the plan’s trustee, subject to participants’ ability to give investment directions by following specified procedures. We do not currently make discretionary contributions or matching contributions to our 401(k) plan.

Except for the benefits described above under “Narrative to Summary Compensation Table — Other Compensation,” we do not provide perquisites or personal benefits to our named executive officers. We do, however, pay the premiums for medical, dental and term life insurance for all of our employees, including our named executive officers.

Equity Incentive Plans

2014 Equity Incentive Plan

We expect that our board of directors will adopt, and our stockholders will approve, prior to the completion of this offering our 2014 Equity Incentive Plan, or our 2014 plan. We do not expect to issue equity awards under our 2014 plan until after the completion of this offering. No awards have been granted and no shares of our common stock have been issued under our 2014 plan. Our 2014 plan will provide for the grant of incentive stock options within the meaning of Section 422 of the Internal Revenue Code, or the Code, to our employees and our parent and subsidiary corporations’ employees, and for the grant of nonstatutory stock options, restricted stock awards, restricted stock unit awards, stock appreciation rights, performance stock awards and other forms of stock compensation to our employees, including officers, consultants and directors. Our 2014 plan will also provide for the grant of performance cash awards to our employees, consultants and directors.

Authorized Shares

The maximum number of shares of our common stock that may be issued under our 2014 plan is _____ shares. The number of shares of our common stock reserved for issuance under our 2014 plan will _____

[Table of Contents](#)

automatically increase on January 1 of each year, for a period of 10 years, from January 1, 2015 continuing through January 1, 2024, by % of the total number of shares of our common stock outstanding on December 31 of the preceding calendar year, or a lesser number of shares as may be determined by our board of directors. The maximum number of shares that may be issued pursuant to the exercise of incentive stock options under the 2014 plan is .

Shares issued under our 2014 plan may be authorized but unissued or reacquired shares of our common stock. Shares subject to stock awards granted under our 2014 plan that expire or terminate without being exercised in full, or that are paid out in cash rather than in shares, will not reduce the number of shares available for issuance under our 2014 plan. Additionally, shares issued pursuant to stock awards under our 2014 plan that we repurchase or that are forfeited, as well as shares reacquired by us as consideration for the exercise or purchase price of a stock award or to satisfy tax withholding obligations related to a stock award, will become available for future grant under our 2014 plan.

Administration

Our board of directors, or a duly authorized committee thereof, has the authority to administer our 2014 plan. Our board of directors has delegated its authority to administer our 2014 plan to our compensation committee under the terms of the compensation committee's charter. Our board of directors may also delegate to one or more of our officers the authority to (i) designate employees other than officers to receive specified stock awards and (ii) determine the number of shares of our common stock to be subject to such stock awards. Subject to the terms of our 2014 plan, the administrator has the authority to determine the terms of awards, including recipients, the exercise price or strike price of stock awards, if any, the number of shares subject to each stock award, the fair market value of a share of our common stock, the vesting schedule applicable to the awards, together with any vesting acceleration, the form of consideration, if any, payable upon exercise or settlement of the stock award and the terms and conditions of the award agreements for use under our 2014 plan.

The administrator has the power to modify outstanding awards under our 2014 plan. Subject to the terms of our 2014 plan, the administrator has the authority to reprice any outstanding option or stock appreciation right, cancel and re-grant any outstanding option or stock appreciation right in exchange for new stock awards, cash or other consideration, or take any other action that is treated as a repricing under generally accepted accounting principles, with the consent of any adversely affected participant.

Section 162(m) Limits

No participant may be granted stock awards covering more than shares of our common stock under our 2014 plan during any calendar year pursuant to stock options, stock appreciation rights and other stock awards whose value is determined by reference to an increase over an exercise price or strike price of at least 100% of the fair market value of our common stock on the date of grant. Additionally, no participant may be granted in a calendar year a performance stock award covering more than shares of our common stock or a performance cash award having a maximum value in excess of \$ under our 2014 plan. These limitations enable us to grant awards that will be exempt from the \$1,000,000 limitation on the income tax deductibility of compensation paid per covered executive officer imposed by Section 162(m) of the Code.

[Table of Contents](#)

Performance Awards

Our 2014 plan permits the grant of performance-based stock and cash awards that may qualify as performance-based compensation that is not subject to the \$1,000,000 limitation on the income tax deductibility of compensation paid per covered executive officer imposed by Section 162(m) of the Code. To enable us to grant performance-based awards that will qualify, our compensation committee can structure such awards so that the stock or cash will be issued or paid pursuant to such award only following the achievement of specified pre-established performance goals during a designated performance period.

Corporate Transactions

Our 2014 plan provides that in the event of a specified corporate transaction, including without limitation a consolidation, merger, or similar transaction involving our company, the sale, lease or other disposition of all or substantially all of the assets of our company or the consolidated assets of our company and our subsidiaries, or a sale or disposition of at least 50% of the outstanding capital stock of our company, the administrator will determine how to treat each outstanding stock award. The administrator may:

- arrange for the assumption, continuation or substitution of a stock award by a successor corporation;
- arrange for the assignment of any reacquisition or repurchase rights held by us to a successor corporation;
- accelerate the vesting of the stock award and provide for its termination prior to the effective time of the corporate transaction;
- arrange for the lapse, in whole or in part, of any reacquisition or repurchase right held by us; or
- cancel the stock award prior to the transaction in exchange for a cash payment, which may be reduced by the exercise price payable in connection with the stock award.

The administrator is not obligated to treat all stock awards or portions of stock awards, even those that are of the same type, in the same manner. The administrator may take different actions with respect to the vested and unvested portions of a stock award.

Change in Control

The administrator may provide, in an individual award agreement or in any other written agreement between us and the participant, that the stock award will be subject to additional acceleration of vesting and exercisability in the event of a change in control. In the absence of such a provision, no such acceleration of the stock award will occur.

Plan Amendment or Termination

Our board has the authority to amend, suspend, or terminate our 2014 plan, provided that such action does not materially impair the existing rights of any participant without such participant's written consent. No incentive stock options may be granted after the tenth anniversary of the date our board of directors adopts our 2014 plan.

2011 Stock Incentive Plan

Our board of directors adopted, and our stockholders approved, the 2011 Stock Incentive Plan, or the 2011 plan, in November 2011. Our 2011 plan was amended by our board of directors and our stockholders

[Table of Contents](#)

in December 2011. Our 2011 plan provides for the grant of incentive stock options within the meaning of Section 422 of the Code to our employees, and for the grant of nonstatutory stock options, restricted stock awards, restricted stock units and other stock-based awards to our officers, directors, employees, consultants and advisers.

Authorized Shares

There are 3,338,776 shares of our common stock reserved for issuance under our 2011 plan. As of June 30, 2014, 420,934 shares of our common stock have been issued upon the exercise of options granted under our 2011 plan and options to purchase 1,619,235 shares of our common stock were outstanding at a weighted average exercise price of \$0.16 per share. Effective upon the completion of this offering, no further options or stock awards may be granted under our 2011 plan, but all outstanding stock awards will continue to be governed by their existing terms.

Administration

Our board of directors, or a committee thereof appointed by our board of directors, administers our 2011 plan and the option and stock awards granted under it. Our board of directors delegated its authority to administer our 2011 plan to our compensation committee.

Corporate Transactions

Our 2011 plan provides that, in the event of a specified change of control transaction, including without limitation a dissolution, merger, consolidation or reorganization of our company with one or more other entities in which our company is not the surviving entity, a sale of substantially all of the assets of our company, or any transaction which results in the disposition of at least a majority of the voting power or value of the securities of our company, the board may take any one or more of the following actions with respect to awards other than restricted stock awards:

- the assumption or substitution of the options by a successor corporation;
- the termination of the options immediately prior to the change of control transaction;
- provide that the options become exercisable, realizable or derivable, or that restrictions applicable to options shall lapse;
- the purchase of outstanding options for an amount of cash that could have been received upon the exercise of the options or the conversion of the options into a right to receive liquidation proceeds; or
- any combination of the foregoing.

With respect to restricted stock awards, upon the occurrence of a change of control transaction involving the liquidation or dissolution of our company, all restrictions and conditions on all restricted stock awards then outstanding shall automatically be deemed terminated or satisfied. With respect to a change of control transaction not involving the liquidation or dissolution of our company, all of our repurchase or other rights under each outstanding restricted stock award shall inure to the benefit of our successor.

2014 Employee Stock Purchase Plan

We expect that our board of directors will adopt, and our stockholders will approve, prior to the completion of this offering, our 2014 Employee Stock Purchase Plan, or our 2014 ESPP. We expect that the

2014 ESPP will become effective upon the completion of this offering, but we have no current plans to grant purchase rights under our 2014 ESPP.

The maximum number of shares of our common stock that may be issued under our 2014 ESPP is _____ shares. Additionally, the number of shares of our common stock reserved for issuance under our 2014 ESPP will automatically increase on January 1 of each year, beginning on January 1 of the year after the completion of this offering and ending on and including January 1, 2024, by the lesser of (i) 1% of the total number of shares of our common stock outstanding on December 31 of the preceding calendar year or (ii) 1,000,000 shares of common stock; provided, however, our board of directors may act prior to the first day of any calendar year to provide that there will be no January 1 increase in the share reserve for such calendar year or that the increase in the share reserve for such calendar year will be a lesser number of shares of common stock. Shares subject to purchase rights granted under our 2014 ESPP that terminate without having been exercised in full will not reduce the number of shares available for issuance under our 2014 ESPP.

Our board of directors, or a duly authorized committee thereof, will administer our 2014 ESPP. Our board of directors has delegated its authority to administer our 2014 ESPP to our compensation committee under the terms of the compensation committee's charter.

Employees, including executive officers, of ours or any of our designated affiliates may have to satisfy one or more of the following service requirements before participating in our 2014 ESPP, as determined by the administrator: (i) customary employment with us or one of our affiliates for more than 20 hours per week and more than five months per calendar year; or (ii) continuous employment with us or one of our affiliates for a minimum period of time, not to exceed two years, prior to the first date of an offering. An employee may not be granted rights to purchase stock under our 2014 ESPP if such employee (i) immediately after the grant would own stock possessing 5% or more of the total combined voting power or value of all classes of our common stock, or (ii) holds rights to purchase stock under our 2014 ESPP that would accrue at a rate that exceeds \$25,000 worth of our stock for each calendar year that the rights remain outstanding.

A component of our 2014 ESPP is intended to qualify as an employee stock purchase plan under Section 423 of the Code and the provisions of this component will be construed in a manner that is consistent with the requirements of Section 423 of the Code. In addition, the 2014 ESPP authorizes the grant of options to purchase shares of our common stock that do not meet the requirements of Section 423 of the Code because of deviations necessary to permit participation in the 2014 ESPP by employees who are foreign nationals or employed outside of the United States while complying with applicable foreign laws. Any such options must be granted pursuant to rules, procedures or subplans adopted by our board designed to achieve these objectives for eligible employees and our company. The administrator may specify offerings with a duration of not more than 27 months, and may specify one or more shorter purchase periods within each offering. Each offering will have one or more purchase dates on which shares of our common stock will be purchased for the employees who are participating in the offering. The administrator, in its discretion, will determine the terms of offerings under our 2014 ESPP.

Our 2014 ESPP permits participants to purchase shares of our common stock through payroll deductions up to 15% of their earnings. Unless otherwise determined by the administrator, the purchase price of the shares will be 85% of the lower of the fair market value of our common stock on the first day of

[Table of Contents](#)

an offering or on the date of purchase. Participants may end their participation at any time during an offering and will be paid their accrued contributions that have not yet been used to purchase shares. Participation ends automatically upon termination of employment with us.

A participant may not transfer purchase rights under our 2014 ESPP other than by will, the laws of descent and distribution or as otherwise provided under our 2014 ESPP.

In the event of a specified corporate transaction, such as a merger or change in control of our company, a successor corporation may assume, continue or substitute each outstanding purchase right. If the successor corporation does not assume, continue or substitute for the outstanding purchase rights, the offering in progress will be shortened and a new exercise date will be set. The participants' purchase rights will be exercised on the new exercise date and such purchase rights will terminate immediately thereafter.

Our board of directors has the authority to amend, suspend or terminate our 2014 ESPP, at any time and for any reason. Our 2014 ESPP will remain in effect until terminated by our board of directors in accordance with the terms of the 2014 ESPP.

Limitations on Liability and Indemnification Matters

Our amended and restated certificate of incorporation, which will become effective upon the completion of this offering, contains provisions that limit the liability of our current and former directors for monetary damages to the fullest extent permitted by Delaware law. Delaware law provides that directors of a corporation will not be personally liable for monetary damages for any breach of fiduciary duties as directors, except liability for:

- any breach of the director's duty of loyalty to the corporation or its stockholders;
- any act or omission not in good faith or that involves intentional misconduct or a knowing violation of law;
- unlawful payments of dividends or unlawful stock repurchases or redemptions as provided in Section 174 of the Delaware General Corporation Law; or
- any transaction from which the director derived an improper personal benefit.

This limitation of liability does not apply to liabilities arising under federal securities laws and does not affect the availability of equitable remedies such as injunctive relief or rescission.

Our amended and restated certificate of incorporation and our amended and restated bylaws will provide that we are required to indemnify our directors to the fullest extent permitted by Delaware law. Our amended and restated bylaws will also provide that, upon satisfaction of certain conditions, we are required to advance expenses incurred by a director in advance of the final disposition of any action or proceeding, and permit us to secure insurance on behalf of any officer, director, employee or other agent for any liability arising out of his or her actions in that capacity regardless of whether we would otherwise be permitted to indemnify him or her under the provisions of Delaware law. Our amended and restated bylaws will also provide our board of directors with discretion to indemnify our officers and employees when determined appropriate by the board. We have entered into and expect to continue to enter into agreements to indemnify our directors, and we also expect to enter into agreements to indemnify our officers, as determined by the board of directors. With certain exceptions, these agreements provide for indemnification for related expenses including, among other things, attorneys' fees, judgments, fines and settlement amounts incurred

[Table of Contents](#)

by any of these individuals in any action or proceeding. We believe that these bylaw provisions and indemnification agreements are necessary to attract and retain qualified persons as directors. We also maintain customary directors' and officers' liability insurance.

The limitation of liability and indemnification provisions in our amended and restated certificate of incorporation and amended and restated bylaws may discourage stockholders from bringing a lawsuit against our directors for breach of their fiduciary duty. They may also reduce the likelihood of derivative litigation against our directors and officers, even though an action, if successful, might benefit us and other stockholders. Further, a stockholder's investment may be adversely affected to the extent that we pay the costs of settlement and damage awards against directors and officers as required by these indemnification provisions. At present, there is no pending litigation or proceeding involving any of our directors, officers or employees for which indemnification is sought and we are not aware of any threatened litigation that may result in claims for indemnification.

Rule 10b5-1 Sales Plans

Our directors and executive officers may adopt written plans, known as Rule 10b5-1 plans, in which they will contract with a broker to buy or sell shares of our common stock on a periodic basis. Under a Rule 10b5-1 plan, a broker executes trades pursuant to parameters established by the director or officer when entering into the plan, without further direction from them. The director or officer may amend a Rule 10b5-1 plan in some circumstances and may terminate a plan at any time. Our directors and executive officers also may buy or sell additional shares outside of a Rule 10b5-1 plan when they are not in possession of material nonpublic information subject to compliance with the terms of our insider trading policy. Prior to 180 days after the date of this offering, subject to early termination, the sale of any shares under such plan would be prohibited by the lock-up agreement that the director or officer has entered into with the underwriters.

CERTAIN RELATIONSHIPS AND RELATED PARTY TRANSACTIONS

The following is a description of transactions since our inception on May 26, 2011 to which we have been a participant in which the amount involved exceeded or will exceed \$120,000, and in which any of our directors, executive officers or holders of more than 5% of our capital stock, or any members of their immediate family, had or will have a direct or indirect material interest, other than compensation arrangements which are described under "Executive Compensation."

Sales of Series A Convertible Preferred Stock

In January 2012, February 2012 and July 2012, we sold an aggregate of 5,198,826 shares of our Series A convertible preferred stock at a price of \$0.78589 per share for an aggregate price of \$4.1 million, 5,071,582 shares of which were sold to holders of more than 5% of our voting securities and members of our board of directors. The table below summarizes these sales.

<u>Purchaser</u>	<u>Shares of Series A Convertible Preferred Stock Purchased</u>	<u>Aggregate Purchase Price</u>
Entities affiliated with Hatteras Venture Partners(1)	4,071,815	\$ 3,199,999
Entities affiliated with GRA Venture Fund(2)	866,533	681,000
Daniel H. White(3)	133,234	104,707
Total	5,071,582	\$ 3,985,706

- (1) Consists of 1,866,418 shares purchased by Hatteras Venture Partners III, LP, 169,489 shares purchased by Hatteras Venture Affiliates III, LP and 2,035,908 shares purchased by Hatteras Venture Partners IV SBIC, LP. Entities affiliated with Hatteras Venture Partners are holders of more than 5% of our voting securities, and Christy L. Shaffer, Ph.D. and Clay B. Thorp are affiliated with Hatteras Venture Partners and are members of our board of directors.
- (2) Consists of 547,492 shares purchased by GRA Venture Fund, LLC and 319,041 shares purchased by GRA Venture Fund (T.E.), LLC. Entities affiliated with GRA Venture Fund are holders of more than 5% of our voting securities.
- (3) Mr. White's investment represented the conversion of \$100,000 in principal amount under convertible promissory notes issued to Mr. White between June 2011 and December 2011, plus accrued interest of \$4,707.

Sales of Series A-1 Convertible Preferred Stock

In January 2013, we sold an aggregate of 4,356,931 shares of our Series A-1 convertible preferred stock at a price of \$1.8132 per share for an aggregate price of \$7.9 million, 3,860,573 shares of which were sold to entities affiliated with members of our board of directors and holders of more than 5% of our voting securities. The table below summarizes these sales.

<u>Purchaser</u>	<u>Shares of Series A-1 Convertible Preferred Stock Purchased</u>	<u>Aggregate Purchase Price</u>
Entities affiliated with Hatteras Venture Partners(1)	1,654,531	\$ 2,999,996
Entities affiliated with GRA Venture Fund(2)	275,754	499,997
Santen Pharmaceutical Co., Ltd.	1,930,288	3,499,998
Total	3,860,573	\$ 6,999,991

[Table of Contents](#)

- (1) Consists of 631,997 shares purchased by Hatteras Venture Partners III, LP, 57,391 shares purchased by Hatteras Venture Affiliates III, LP, 689,388 shares purchased by Hatteras Venture Partners IV SBIC, LP and 275,755 shares purchased by Hatteras NC Fund, LP. Entities affiliated with Hatteras Venture Partners are holders of more than 5% of our voting securities, and Christy Shaffer, Ph.D. and Clay B. Thorp are affiliated with Hatteras Venture Partners and are members of our board of directors.
- (2) Consists of 174,226 shares purchased by GRA Venture Fund, LLC and 101,528 shares purchased by GRA Venture Fund (T.E.), LLC. Entities affiliated with GRA Venture Fund are holders of more than 5% of our voting securities.

Unsecured 7% Convertible Promissory Notes

In April and May 2014, we issued an aggregate principal amount of \$3.0 million of unsecured 7% convertible promissory notes, or the bridge notes, and warrants to purchase an aggregate of 248,175 shares of our common stock at an exercise price of \$0.01 per share. We issued an aggregate principal amount of \$2,196,993 of our bridge notes and warrants to purchase up to 181,750 shares of our common stock to holders of more than 5% of our voting securities, members of our board of directors and our officers. The table below summarizes these issuances.

<u>Name</u>	<u>Principal Amount of Unsecured Convertible Notes</u>	<u>Common Stock Warrants</u>
Entities affiliated with Hatteras Venture Partners(1)	\$ 1,154,496	95,508
Entities affiliated with GRA Venture Fund(2)	499,997	41,363
Santen Pharmaceutical Co., Ltd.	500,000	41,363
Daniel H. White	17,500	1,448
Charles A. Deignan	12,500	1,034
Gerald D. Cagle, Ph.D.	12,500	1,034
Total	\$ 2,196,993	181,750

- (1) Consists of bridge notes and warrants issued to Hatteras Venture Partners III, LP. Entities affiliated with Hatteras Venture Partners are holders of more than 5% of our voting securities, and Christy L. Shaffer, Ph.D. and Clay B. Thorp are affiliated with Hatteras Venture Partners and are members of our board of directors.
- (2) Consists of a principal amount of \$315,911 of our bridge notes and warrants to purchase 31,023 shares of our common stock issued to GRA Venture Fund, LLC, and a principal amount of \$184,086 of our bridge notes and warrants to purchase 10,341 shares of our common stock issued to GRA Venture Fund (T.E.), LLC. Entities affiliated with GRA Venture Fund are holders of more than 5% of our voting securities.

All principal and interest under the bridge notes was converted into shares of our Series B convertible preferred stock in connection with our August 2014 financing described below.

Sales of Series B Convertible Preferred Stock

In August 2014, we issued an aggregate of 6,009,202 shares of our Series B convertible preferred stock at a price of \$2.69783 per share for an aggregate price of \$16.2 million, 4,302,359 shares of which were

[Table of Contents](#)

sold to entities affiliated with members of our board of directors, our officers and holders of more than 5% of our voting securities. In some cases, some or all of the purchase price for these shares took the form of conversion of principal and interest under outstanding bridge notes held by the respective investors. In connection with this financing, we also issued warrants to purchase an aggregate of 1,716,914 shares of common stock at an exercise price of \$0.01 per share, which will become exercisable in February 2015 if we have not completed an initial public offering by then, but otherwise will terminate upon the closing of our initial public offering. Of these warrants issued, warrants to purchase an aggregate of 1,229,248 shares of common stock were issued to entities affiliated with members of our board of directors, our officers and holders of more than 5% of our voting securities. The table below summarizes the issuances of shares of Series B convertible preferred stock and warrants to purchase common stock to entities affiliated with members of our board of directors, our officers and holders of more than 5% of our voting securities.

<u>Purchaser</u>	<u>Shares of Series B Convertible Preferred Stock Purchased</u>	<u>Warrants to Purchase Common Stock Issued</u>	<u>Aggregate Purchase Price</u>	
			<u>Cash</u>	<u>Note Conversion</u>
Entities affiliated with Hatteras Venture Partners(1)	865,881	247,395	\$1,154,492	\$1,181,508
Entities affiliated with GRA Venture Fund(2)	434,937	124,268	499,994	673,394
Santen Pharmaceutical Co., Ltd.	375,004	107,144	499,998	511,699
RMI Investments(3)	2,594,677	741,340	6,999,997	—
Daniel H. White	13,110	3,745	17,499	17,871
Charles A. Deignan	9,375	2,678	12,500	12,792
Gerald D. Cagle	9,375	2,678	12,500	12,792
Total	4,302,359	1,229,248	\$9,196,980	\$2,410,056

- (1) Consists of (i) 396,920 shares of Series B convertible preferred stock and a warrant to purchase 113,406 shares of common stock issued to Hatteras Venture Partners III, LP, (ii) 36,020 shares of Series B convertible preferred stock and a warrant to purchase 10,291 shares of common stock issued to Hatteras Venture Affiliates III, LP and (iii) 432,941 shares of Series B convertible preferred stock and a warrant to purchase 123,698 shares of common stock issued to Hatteras Venture Partners IV SBIC, LP. Entities affiliated with Hatteras Venture Partners are holders of more than 5% of our voting securities, and Christy L. Shaffer, Ph.D. and Clay B. Thorp are affiliated with Hatteras Venture Partners and are members of our board of directors.
- (2) Consists of (i) 236,711 shares of Series B convertible preferred stock and a warrant to purchase 67,632 shares of common stock issued to GRA Venture Fund, LLC, (ii) 137,935 shares of Series B convertible preferred stock and a warrant to purchase 39,410 shares of common stock issued to GRA Venture Fund (T.E.), LLC and (iii) 60,291 shares of Series B convertible preferred stock and a warrant to purchase 17,226 shares of common stock issued to Georgia Research Alliance, Inc. Entities affiliated with GRA Venture Fund are holders of more than 5% of our voting securities.
- (3) Entities affiliated with RMI Investments, or RMI, are holders of more than 5% of our voting securities and Evgeny Zaytsev, M.D. is affiliated with RMI Investments and is a member of our board of directors.

NovaMedica License Agreement

In connection with our Series B financing, in August 2014, we entered into a license agreement with NovaMedica LLC, or NovaMedica. Under this agreement, we granted to NovaMedica the exclusive right in Russia and specified adjacent territories to use our intellectual property to develop and commercialize products involving the use of TA as the sole active pharmaceutical ingredient for administration in the SCS. In connection with this license, NovaMedica will make an upfront payment to us of \$200,000. In addition, NovaMedica has agreed to make royalty payments equal to the amount of any royalties we owe to Emory or GTRC pursuant to the Emory/GTRC License Agreement as a result of NovaMedica's sales of products and services covered by the license granted to NovaMedica, up to a maximum of a low single-digit percentage of net sales of such products or services by NovaMedica. NovaMedica has also agreed to make milestone payments, totaling \$12.7 million in the aggregate, upon the achievement of specified development and commercial milestones. NovaMedica is jointly owned by Rusnano MedInvest LLC, or Rusnano MedInvest, and Domain Russia Investments Limited. RMI, which beneficially owns more than 5% of our voting securities, is a wholly owned subsidiary of Rusnano MedInvest.

The term of the NovaMedica license agreement will expire upon the expiration of the last-to-expire valid claim within the patents licensed to NovaMedica pursuant to the agreement. Either we or NovaMedica may terminate the agreement upon written notice in the event of the other party's material breach that is not cured within 20 business days of receiving written notice requesting a cure of a payment breach, and within 60 business days of receiving written notice requesting a cure in the case of all other breaches. Each party also has the right to terminate in the event of the other party's bankruptcy or insolvency. We may terminate the agreement immediately upon written notice in the event that NovaMedica commences or participates in any action challenging the validity or enforceability of the patents licensed to NovaMedica under the agreement.

Santen Research Collaboration

In January 2013, we entered into a collaboration agreement with Santen, which beneficially owns more than 5% of our common stock. Under this agreement, we and Santen agreed to conduct feasibility studies to identify compounds for further development. Each party to the agreement bears its own costs, except that some of the costs we may incur are limited to a maximum amount. We incurred research and development costs under this agreement of \$162,000 during the year ended December 31, 2013 and \$51,000 during the six months ended June 30, 2014.

Investor Rights Agreement

We have entered into an investor rights agreement, as amended, with our preferred stockholders, including entities affiliated with Hatteras, Santen, GRA and RMI, each of which beneficially own more than 5% of our common stock. The investor rights agreement, among other things:

- grants our preferred stockholders specified registration rights with respect to shares of our common stock, including shares of common stock issued or issuable upon conversion of shares of our convertible preferred stock held by them;
- obligates us to deliver periodic financial statements to some of the stockholders who are parties to the investor rights agreement; and
- grants a right of first refusal with respect to sales of our shares by us, subject to specified exclusions, which exclusions include the sale of the shares pursuant to this prospectus, to the stockholders who are parties to the investor rights agreement.

[Table of Contents](#)

For more information regarding the registration rights provided in this agreement, please refer to the section titled “Description of Capital Stock — Registration Rights.” The provisions of this agreement other than those relating to registration rights will terminate upon the completion of this offering.

Voting Agreement

We have entered into a voting agreement, as amended, with some of our stockholders, including entities affiliated with Hatteras, Santen, GRA and RMI. The voting agreement, among other things, provides for the voting of shares with respect to the constituency of our board of directors. The voting agreement will terminate upon the completion of this offering.

Stock Sale Agreement

We have entered into a stock sale agreement, as amended, with some of our stockholders, including entities affiliated with Hatteras, Santen, GRA and RMI. The stock sale agreement, among other things:

- grants our investors and our founders rights of first refusal and co-sale with respect to proposed transfers of our securities by specified stockholders;
- grants us rights of first refusal with respect to proposed transfers of our securities by specified stockholders; and
- provides for the voting of shares with respect to specified transactions approved by a majority of holders of our outstanding convertible preferred stock.

The stock sale agreement will terminate upon the completion of this offering.

Indemnification Agreements

Our amended and restated certificate of incorporation will contain provisions limiting the liability of directors, and our amended and restated bylaws will provide that we will indemnify each of our directors to the fullest extent permitted under Delaware law. Our amended and restated certificate of incorporation and amended and restated bylaws will also provide our board of directors with discretion to indemnify our officers and employees when determined appropriate by the board.

In addition, we have entered into an indemnification agreement with each of our directors. For more information regarding these agreements, see “Executive Compensation — Limitations on Liability and Indemnification Matters.”

Related Person Transaction Policy

Prior to this offering, we have not had a formal policy regarding approval of transactions with related parties. Prior to the completion of this offering, we expect to adopt a related person transaction policy that sets forth our procedures for the identification, review, consideration and approval or ratification of related person transactions. The policy will become effective immediately upon the execution of the underwriting agreement for this offering. For purposes of our policy only, a related person transaction is a transaction, arrangement or relationship, or any series of similar transactions, arrangements or relationships, in which we and any related person are, were or will be participants in which the amount involved exceeds \$120,000. Transactions involving compensation for services provided to us as an employee or director are not covered by this policy. A related person is any executive officer, director or beneficial owner of more than 5% of any class of our voting securities, including any of their immediate family members and any entity owned or controlled by such persons.

[Table of Contents](#)

Under the policy, if a transaction has been identified as a related person transaction, including any transaction that was not a related person transaction when originally consummated or any transaction that was not initially identified as a related person transaction prior to consummation, our management must present information regarding the related person transaction to our audit committee, or, if audit committee approval would be inappropriate, to another independent body of our board of directors, for review, consideration and approval or ratification. The presentation must include a description of, among other things, the material facts, the interests, direct and indirect, of the related persons, the benefits to us of the transaction and whether the transaction is on terms that are comparable to the terms available to or from, as the case may be, an unrelated third party or to or from employees generally. Under the policy, we will collect information that we deem reasonably necessary from each director, executive officer and, to the extent feasible, significant stockholder to enable us to identify any existing or potential related-person transactions and to effectuate the terms of the policy. In addition, under our Code of Business Conduct and Ethics that we expect to adopt prior to the completion of this offering, our employees and directors will have an affirmative responsibility to disclose any transaction or relationship that reasonably could be expected to give rise to a conflict of interest. In considering related person transactions, our audit committee, or other independent body of our board of directors, will take into account the relevant available facts and circumstances including, but not limited to:

- the risks, costs and benefits to us;
- the impact on a director's independence in the event that the related person is a director, immediate family member of a director or an entity with which a director is affiliated;
- the availability of other sources for comparable services or products; and
- the terms available to or from, as the case may be, unrelated third parties or to or from employees generally.

The policy requires that, in determining whether to approve, ratify or reject a related person transaction, our audit committee, or other independent body of our board of directors, must consider, in light of known circumstances, whether the transaction is in, or is not inconsistent with, our best interests and those of our stockholders, as our audit committee, or other independent body of our board of directors, determines in the good faith exercise of its discretion.

PRINCIPAL STOCKHOLDERS

The following table sets forth the beneficial ownership of our common stock as of August 31, 2014 for:

- each person, or group of affiliated persons, who is known by us to beneficially own more than 5% of our common stock;
- each of our named executive officers;
- each of our directors; and
- all of our current executive officers and directors as a group.

The percentage ownership information shown in the table is based upon 19,580,059 shares of common stock outstanding as of August 31, 2014, after giving effect to the conversion of all of our convertible preferred stock into 15,564,959 shares of common stock, which will occur automatically upon the closing of this offering.

We have determined beneficial ownership in accordance with the rules of the SEC. These rules generally attribute beneficial ownership of securities to persons who possess sole or shared voting power or investment power with respect to those securities. In addition, the rules include shares of common stock issuable pursuant to the exercise of stock options or warrants that are exercisable on or before October 30, 2014, which is 60 days after August 31, 2014. These shares are deemed to be outstanding and beneficially owned by the person holding those options or warrants for the purpose of computing the percentage ownership of that person, but they are not treated as outstanding for the purpose of computing the percentage ownership of any other person. Unless otherwise indicated, the persons or entities identified in this table have sole voting and investment power with respect to all shares shown as beneficially owned by them, subject to applicable community property laws.

[Table of Contents](#)

Except as otherwise noted below, the address for persons listed in the table is c/o Clearside Biomedical, Inc., 1220 Old Alpharetta Road, Suite 300, Alpharetta, Georgia 30005.

Name of Beneficial Owner	Number of Shares Beneficially Owned	Percentage of Shares Beneficially Owned	
		Before Offering	After Offering
<i>Principal Stockholders:</i>			
Entities affiliated with Hatteras Venture Partners(1)	6,687,733	34.1%	
Santen Pharmaceutical Co., Ltd.(2)	2,346,655	12.0	
Entities affiliated with GRA Venture Fund(3)	1,618,586	8.3	
RMI Investments(4)	2,594,677	13.3	
<i>Executive Officers and Directors:</i>			
Daniel H. White(5)	1,309,881	6.7	
Charles A. Deignan(6)	155,991	*	
Glenn Noronha, Ph.D.(7)	36,458	*	
Christy L. Schaffer, Ph.D.(1)	—	—	
Clay B. Thorp(1)	6,687,733	34.1	
William D. Humphries(8)	68,750	*	
Gerald D. Cagle, Ph.D.(9)	41,659	*	
Evgeny Zaytsev, M.D.(4)	2,594,677	13.3	
All current directors and executive officers as a group (8 persons)(10)	10,895,149	54.8	

* Represents beneficial ownership of less than 1%.

- (1) Consists of (a) 2,895,335 shares of common stock issuable upon conversion of shares of preferred stock and 43,780 shares of common stock issuable upon exercise of immediately exercisable warrants held by Hatteras Venture Partners III, LP (“HVP III”), (b) 262,900 shares of common stock issuable upon conversion of shares of preferred stock and 3,973 shares of common stock issuable upon exercise of immediately exercisable warrants held by Hatteras Venture Affiliates III, LP (“HVA III”), (c) 3,158,237 shares of common stock issuable upon conversion of shares of preferred stock and 47,753 shares of common stock issuable upon exercise of immediately exercisable warrants held by Hatteras Venture Partners IV SBIC, LP (“HVP IV SBIC”) and (d) 275,755 shares of common stock issuable upon conversion of shares of preferred stock held by Hatteras NC Fund (“Hatteras NC” and together with HVP III, HVA III, and HVP IV SBIC, the “Hatteras Entities”). The shares directly held by HVA III and HVP III are indirectly held by Hatteras Venture Advisors III, LLC (“HVA III LLC”), their general partner. The individual general partners of HVA III LLC are Clay B. Thorp, Robert A. Ingram, Kenneth B. Lee, Douglas Reed, MD and John Crumpler (the “HVA III LLC Directors”). HVA III LLC and the HVA III LLC Directors may be deemed to have shared voting and dispositive power over, and be deemed indirect beneficial owners of, the shares directly held by Hatteras Venture Affiliates III and Hatteras Venture Partners III. The shares directly held by HVP IV SBIC are indirectly held by Hatteras Venture Advisors IV SBIC, LLC (“HVA IV SBIC LLC”), its general partner. The individual general partners of HVA IV SBIC LLC are Clay B. Thorp, Robert A. Ingram, Kenneth B. Lee, Douglas Reed, MD and John Crumpler (the “HVA IV SBIC LLC Directors”). HVA IV SBIC LLC and the HVA IV SBIC LLC Directors may be deemed to have shared voting and dispositive power over, and be deemed indirect beneficial owners

Table of Contents

of, the shares directly held by HVA IV SBIC. The shares directly held by Hatteras NC are indirectly held by Hatteras Venture Advisors IV, LLC (“HVA IV”), its general partner. The individual general partners of HVA IV are Clay B. Thorp, Robert A. Ingram, Kenneth B. Lee, Douglas Reed, MD and John Crumpler (the “HVA IV Directors”). HVA IV and the HVA IV Directors may be deemed to have shared voting and dispositive power over, and be deemed indirect beneficial owners of, the shares directly held by Hatteras NC. Christy Shaffer, one of our directors, is a Venture Partner with Hatteras Venture Partners, but she does not have beneficial ownership over the shares held by HVP III, HVA III, HVP IV SBIC and Hatteras NC. The principal business address of the Hatteras Entities is 280 S. Mangum St., Suite 350, Durham, NC 27701.

- (2) Consists of (a) 2,305,292 shares of common stock issuable upon conversion of shares of preferred stock and (b) 41,363 shares of common stock issuable upon exercise of immediately exercisable warrants, in each case held by Santen Pharmaceutical Co., Ltd. (“Santen”). The principal business address of Santen is 9-19 Shimoshinjo 3-chome, Higashiyodogawa-ku, Osaka 533-8651, Japan.
- (3) Consists of (a) 958,429 shares of common stock issuable upon conversion of shares of preferred stock and 26,134 shares of common stock issuable upon exercise of immediately exercisable warrants held by GRA Venture Fund, LLC (“GRAVF”), (b) 558,504 shares of common stock issuable upon conversion of shares of preferred stock and 15,228 shares of common stock issuable upon exercise of immediately exercisable warrants held by GRA Venture Fund (T.E.), LLC (“GRAVF TE”) and (c) 60,291 shares of common stock issuable upon conversion of shares of preferred stock held in the name of Georgia Research Alliance, Inc. (“GRA, Inc.” and, together with GRAVF and GRAVF TE, the “GRA Entities”) in an account administered by GRA, Inc. pursuant to a contract with the State of Georgia’s Department of Economic Development. The shares directly held by GRAVF are indirectly held by the members of its board of managers, Duane Ackerman, Jim Balloun, Chris Carr, Russ Chandler, Frederick E. Cooper, Brad Currey, Bill Fickling, Rusty French, William Linginfelter, Diana M. Murphy, Sig Mosley, Tom Parker, David Ratcliffe and Fran Rogers (the “GRAVF Directors”). The GRAVF Directors may be deemed to have shared voting and dispositive power over, and be deemed indirect beneficial owners of, the shares directly held by GRAVF. The shares directly held by GRAVF TE are indirectly held by the members of its board of managers, who are the same individuals as the GRAVF Directors (the “GRAVF TE Directors”). The GRAVF TE Directors may be deemed to have shared voting and dispositive power over, and be deemed indirect beneficial owners of, the shares directly held by GRAVF TE. The shares directly held by GRA, Inc. are indirectly held by the members of its board of trustees, Clyde Tuggle, Douglas Hertz, Kathelen Amos, Paul Amos, Dr. Ricardo Azziz, Dr. Mark Becker, Paul Bowers, Carlton Brown, Chris Carr, Dr. Max Cooper, Kenneth Cornelius, Thomas Cousins, Larry Gellerstedt III, George Deese, James Hannan, Robert Hatcher, Edward Heys, William Linginfelter, Kelly Loeffler, Charles McTier, Jere Morehead, Allen Mosley, Thomas Noonan, Charles Ogburn, Kenneth Ostrowski, G.P. Peterson, Parker Petit, David Ratcliffe, Joseph Rogers, William Rogers, John Somerhalder II, Lizanne Thomas, T. Rogers Wade, James Wagner and Felker Ward (the “GRA, Inc. Trustees”). The GRA, Inc. Trustees may be deemed to have shared voting and dispositive power over, and be deemed indirect beneficial owners of, the shares directly held in the name of GRA, Inc. pursuant to a contact with the State of Georgia’s Department of Economic Development. The principal business address of the GRA Entities is 191 Peachtree Street, NE, Suite 846, Atlanta, Georgia 30303.
- (4) Consists of 2,594,677 shares of common stock issuable upon conversion of shares of preferred stock held by RMI. The shares directly held by RMI are indirectly held by Rusnano MedInvest, the parent company of RMI. RMI Partners LLC is the management company for RusnanoMedInvest. The CEO of RMI Partners

Table of Contents

LLC is Vladimir Gurdus and the Managing Partner of RMI Partners LLC is Evgeny Zaytsev, M.D., a member of our board of directors (the "RMI Partners LLC Directors"). RusnanoMedInvest, RMI Partners LLC and the RMI Partners LLC Directors may be deemed to have shared voting and dispositive power over, and be deemed indirect beneficial owners of, the shares directly held by RMI. The principal business address of RMI is Business-Center 29/22, Capital Tower 1st Brestskaya Street, Moscow, 12 125047.

- (5) Consists of (a) 1,016,290 shares of common stock held by Mr. White directly and vested within 60 days of August 31, 2014, (b) 90,000 shares of common stock held by the White Family Trust, for which Mr. White's wife serves as trustee, (c) 5,136 shares of common stock held for the benefit of Mr. White's children under the Georgia Uniform Transfers to Minors Act, for which Mr. White serves as custodian, (d) 141,617 shares of common stock issuable upon conversion of shares of preferred stock held by Mr. White directly, (e) 4,727 shares of common stock issuable upon conversion of shares of preferred stock held by the Daniel H. White (IRA), for which Mr. White serves as trustee, (f) 50,663 shares of common stock underlying options that are exercisable and vested within 60 days of August 31, 2014 and (g) 1,448 shares of common stock issuable upon exercise of immediately exercisable warrants.
- (6) Consists of (a) 79,686 shares of common stock that are vested within 60 days of August 31, 2014, (b) 45,314 shares of restricted stock that are not vested within 60 days of August 31, 2014, (c) 9,375 shares of common stock issuable upon conversion of shares of preferred stock, (d) 20,582 shares of common stock underlying options that are exercisable and vested within 60 days of August 31, 2014 and (e) 1,034 shares of common stock issuable upon exercise of immediately exercisable warrants. Shares of restricted stock that are not vested are subject to a repurchase right in favor of the company if Mr. Deignan does not satisfy the applicable vesting requirements. In any event, unvested restricted shares may not be disposed of until the vesting period has been satisfied.
- (7) Consists of 36,458 shares of common stock underlying options that are exercisable and vested within 60 days of August 31, 2014.
- (8) Consists of 68,750 shares of common stock underlying options that are exercisable and vested within 60 days of August 31, 2014.
- (9) Consists of (a) 9,375 shares of common stock issuable upon conversion of shares of preferred stock (b) 31,250 shares of common stock underlying options that are exercisable and vested within 60 days of August 31, 2014 and (c) 1,034 shares of common stock issuable upon exercise of immediately exercisable warrants.
- (10) Consists of (a) 1,191,112 shares of common stock that are vested within 60 days of August 31, 2014, (b) 9,351,998 shares of common stock issuable upon conversion of shares of preferred stock, (c) 45,314 shares of restricted stock that are not vested within 60 days of August 31, 2014, (d) 207,703 shares of common stock underlying options that are exercisable and vested within 60 days of August 31, 2014 and (e) 99,022 shares of common stock issuable upon exercise of immediately exercisable warrants. Shares of restricted stock that are not vested are subject to a repurchase right in favor of the company if the officer or director does not satisfy the applicable vesting requirements. In any event, unvested restricted shares may not be disposed of until the vesting period has been satisfied.

DESCRIPTION OF CAPITAL STOCK

The following description of our capital stock and provisions of our amended and restated certificate of incorporation and amended and restated bylaws are summaries. You should also refer to the amended and restated certificate of incorporation and the amended and restated bylaws, which are filed as exhibits to the registration statement of which this prospectus is part.

General

Upon the completion of this offering, our amended and restated certificate of incorporation will authorize us to issue up to _____ shares of common stock, \$0.001 par value per share, and _____ shares of preferred stock, \$0.001 par value per share, all of which shares of preferred stock will be undesignated. Our board of directors may establish the rights and preferences of the preferred stock from time to time. As of August 31, 2014, we had outstanding 4,015,100 shares of common stock, held by 17 stockholders of record. As of August 31, 2014, after giving effect to the conversion of all outstanding preferred stock into shares of common stock, there would have been 19,580,059 shares of common stock issued and outstanding, held of record by 46 stockholders.

Common Stock

Voting Rights

Each holder of our common stock is entitled to one vote for each share on all matters submitted to a vote of the stockholders, including the election of directors. Under our amended and restated certificate of incorporation and amended and restated bylaws, our stockholders will not have cumulative voting rights. Because of this, the holders of a majority of the shares of common stock entitled to vote in any election of directors can elect all of the directors standing for election, if they should so choose.

Dividends

Subject to preferences that may be applicable to any then-outstanding preferred stock, holders of common stock are entitled to receive ratably those dividends, if any, as may be declared from time to time by the board of directors out of legally available funds.

Liquidation

In the event of our liquidation, dissolution or winding up, holders of common stock will be entitled to share ratably in the net assets legally available for distribution to stockholders after the payment of all of our debts and other liabilities and the satisfaction of any liquidation preference granted to the holders of any then-outstanding shares of preferred stock.

Rights and Preferences

Holders of common stock have no preemptive, conversion or subscription rights and there are no redemption or sinking fund provisions applicable to the common stock. The rights, preferences and privileges of the holders of common stock are subject to, and may be adversely affected by, the rights of the holders of shares of any series of preferred stock that we may designate in the future.

Preferred Stock

As of August 31, 2014, there were outstanding 15,564,959 shares of convertible preferred stock. All currently outstanding shares of convertible preferred stock will be converted automatically into an aggregate of 15,564,959 shares of common stock immediately prior to the completion of this offering.

Following the completion of this offering, our board of directors will have the authority, without further action by our stockholders, to issue up to 5,000,000 shares of preferred stock in one or more series, to establish from time to time the number of shares to be included in each such series, to fix the rights, preferences and privileges of the shares of each wholly unissued series and any qualifications, limitations or restrictions thereon, and to increase or decrease the number of shares of any such series, but not below the number of shares of such series then outstanding.

Our board of directors may authorize the issuance of preferred stock with voting or conversion rights that could adversely affect the voting power or other rights of the holders of our common stock. The purpose of authorizing our board of directors to issue preferred stock and determine its rights and preferences is to eliminate delays associated with a stockholder vote on specific issuances. The issuance of preferred stock, while providing flexibility in connection with possible acquisitions and other corporate purposes, could, among other things, have the effect of delaying, deferring or preventing a change in control of us and may adversely affect the market price of our common stock and the voting and other rights of the holders of our common stock. It is not possible to state the actual effect of the issuance of any shares of preferred stock on the rights of holders of common stock until the board of directors determines the specific rights attached to that preferred stock.

We have no present plans to issue any shares of preferred stock following completion of this offering.

Options

As of August 31, 2014, under our 2011 plan, options to purchase an aggregate of 1,685,902 shares of common stock were outstanding. For additional information regarding the terms of this plan, see “Executive Compensation — Equity Incentive Plans.”

Warrants

We have outstanding an immediately exercisable warrant to purchase 16,550 shares of our Series A-1 convertible preferred stock at an exercise price of \$1.81 per share, which expires in February 2023. This warrant has a net exercise provision under which the holder may, in lieu of payment of the exercise price in cash, surrender the warrant and receive a net amount of shares based on the fair market value of our stock at the time of exercise of the warrant after deduction of the aggregate exercise price. The warrant also contain provisions for the adjustment of the exercise price and the number of shares issuable upon the exercise of the warrant in the event of certain stock dividends, stock splits, reorganizations, reclassifications and consolidations. Upon the completion of this offering, unless earlier exercised, this warrant will be converted into a warrant to purchase 16,550 shares of our common stock at \$1.81 per share.

We also have outstanding immediately exercisable warrants to purchase an aggregate of 248,175 shares of our common stock at an exercise price of \$0.01 per share, which expire in April 2024. These warrants have net exercise provisions under which the holder may, in lieu of payment of the exercise price in cash, surrender the warrant and receive a net amount of shares based on the fair market value of our common stock at the time of exercise of the warrant after deduction of the aggregate exercise price. These

warrants also contain provisions for the adjustment of the exercise price and the number of shares issuable upon the exercise of the warrant in the event of certain stock dividends, stock splits, reorganizations, reclassifications and consolidations. If unexercised, these warrants will expire upon the closing of this offering, and therefore we expect that these warrants will be exercised in connection with the completion of this offering and that we will issue approximately 248,175 shares of our common stock upon their exercise.

Registration Rights

We and the holders of our existing convertible preferred stock have entered into an investor rights agreement. The registration rights provisions of this agreement provide those holders with demand and piggyback registration rights with respect to the shares of our common stock currently held by them and issuable to them upon exercise of warrants and upon conversion of our convertible preferred stock in connection with this offering.

Demand Registration Rights

At any time beginning 180 days following the effective date of the registration statement of which this prospectus is a part, the holders of at least 40% of the shares issuable upon conversion of our convertible preferred stock in the aggregate have the right to demand that we file up to two Form S-1 registration statements, as long as the anticipated aggregate offering price, net of underwriting discounts and commissions, would exceed \$10.0 million. These registration rights are subject to specified conditions and limitations, including the right of the underwriters, if any, to limit the number of shares included in any such registration under specified circumstances. Upon such a request, we are required to effect the registration as soon as reasonably possible. An aggregate of _____ shares of common stock and _____ shares issuable upon the exercise of warrants will be entitled to these demand registration rights.

Piggyback Registration Rights

At any time after the completion of this offering, if we propose to register any of our securities under the Securities Act either for our own account or for the account of other stockholders, the holders of shares of common stock that are issued upon conversion of our convertible preferred stock, some holders of shares of our common stock and the holders of our currently outstanding warrants will each be entitled to notice of the registration and will be entitled to include their shares of common stock in the registration statement. These piggyback registration rights are subject to specified conditions and limitations, including the right of the underwriters to limit the number of shares included in any such registration under specified circumstances. An aggregate of _____ shares of common stock and _____ shares issuable upon the exercise of warrants will be entitled to these piggyback registration rights.

Registration on Form S-3

At any time after we become eligible to file a registration statement on Form S-3, holders of shares of our common stock that are issued upon conversion of our convertible preferred stock will be entitled, upon their written request, to have such shares registered by us on a Form S-3 registration statement at our expense, provided that such requested registration has an anticipated aggregate offering size to the public of at least \$1.0 million and subject to other specified conditions and limitations. An aggregate of _____ shares of common stock and _____ shares issuable upon the exercise of warrants will be entitled to these Form S-3 registration rights.

[Table of Contents](#)

Expenses of Registration

We will pay all expenses relating to any demand, piggyback or Form S-3 registration, other than underwriting discounts and commissions, subject to specified conditions and limitations.

Termination of Registration Rights

The registration rights granted under the investor rights agreement will terminate upon the earliest to occur of (i) the fifth anniversary of the completion of this offering or (ii) the closing of a liquidating event.

Anti-Takeover Provisions

Section 203 of the Delaware General Corporation Law

We are subject to Section 203 of the Delaware General Corporation Law, which prohibits a Delaware corporation from engaging in any business combination with any interested stockholder for a period of three years after the date that such stockholder became an interested stockholder, with the following exceptions:

- before such date, the board of directors of the corporation approved either the business combination or the transaction that resulted in the stockholder becoming an interested stockholder;
- upon completion of the transaction that resulted in the stockholder becoming an interested stockholder, the interested stockholder owned at least 85% of the voting stock of the corporation outstanding at the time the transaction began, excluding for purposes of determining the voting stock outstanding, but not the outstanding voting stock owned by the interested stockholder, those shares owned (i) by persons who are directors and also officers and (ii) employee stock plans in which employee participants do not have the right to determine confidentially whether shares held subject to the plan will be tendered in a tender or exchange offer; or
- on or after such date, the business combination is approved by the board of directors and authorized at an annual or special meeting of the stockholders, and not by written consent, by the affirmative vote of at least 66 2/3% of the outstanding voting stock that is not owned by the interested stockholder.

In general, Section 203 defines a “business combination” to include the following:

- any merger or consolidation involving the corporation and the interested stockholder;
- any sale, transfer, pledge or other disposition of 10% or more of the assets of the corporation involving the interested stockholder;
- subject to certain exceptions, any transaction that results in the issuance or transfer by the corporation of any stock of the corporation to the interested stockholder;
- any transaction involving the corporation that has the effect of increasing the proportionate share of the stock or any class or series of the corporation beneficially owned by the interested stockholder; or
- the receipt by the interested stockholder of the benefit of any loans, advances, guarantees, pledges or other financial benefits by or through the corporation.

In general, Section 203 defines an “interested stockholder” as an entity or person who, together with the person’s affiliates and associates, beneficially owns, or within three years prior to the time of determination of interested stockholder status did own, 15% or more of the outstanding voting stock of the corporation.

Certificate of Incorporation and Bylaws to be in Effect Upon the Completion of this Offering

Our amended and restated certificate of incorporation to be in effect upon the completion of this offering, or our restated certificate, will provide for our board of directors to be divided into three classes with staggered three-year terms. Only one class of directors will be elected at each annual meeting of our stockholders, with the other classes continuing for the remainder of their respective three-year terms. Because our stockholders do not have cumulative voting rights, stockholders holding a majority of the shares of common stock outstanding will be able to elect all of our directors. Our restated certificate and our amended and restated bylaws to be effective upon the completion of this offering, or our restated bylaws, will also provide that directors may be removed by the stockholders only for cause upon the vote of 66 2/3% or more of our outstanding common stock. Furthermore, the authorized number of directors may be changed only by resolution of the board of directors, and vacancies and newly created directorships on the board of directors may, except as otherwise required by law or determined by the board, only be filled by a majority vote of the directors then serving on the board, even though less than a quorum.

Our restated certificate and restated bylaws will also provide that all stockholder actions must be effected at a duly called meeting of stockholders and will eliminate the right of stockholders to act by written consent without a meeting. Our restated bylaws will also provide that only our chairman of the board, chief executive officer or the board of directors pursuant to a resolution adopted by a majority of the total number of authorized directors may call a special meeting of stockholders.

Our restated bylaws will also provide that stockholders seeking to present proposals before a meeting of stockholders or to nominate candidates for election as directors at a meeting of stockholders must provide timely advance notice in writing, and will specify requirements as to the form and content of a stockholder's notice.

Our restated certificate and restated bylaws will provide that the stockholders cannot amend many of the provisions described above except by a vote of 66 2/3% or more of our outstanding common stock.

The combination of these provisions will make it more difficult for our existing stockholders to replace our board of directors as well as for another party to obtain control of us by replacing our board of directors. Since our board of directors has the power to retain and discharge our officers, these provisions could also make it more difficult for existing stockholders or another party to effect a change in management. In addition, the authorization of undesignated preferred stock makes it possible for our board of directors to issue preferred stock with voting or other rights or preferences that could impede the success of any attempt to change our control.

These provisions are intended to enhance the likelihood of continued stability in the composition of our board of directors and its policies and to discourage coercive takeover practices and inadequate takeover bids. These provisions are also designed to reduce our vulnerability to hostile takeovers and to discourage certain tactics that may be used in proxy fights. However, such provisions could have the effect of discouraging others from making tender offers for our shares and may have the effect of delaying changes in our control or management. As a consequence, these provisions may also inhibit fluctuations in the market price of our stock that could result from actual or rumored takeover attempts. We believe that the benefits of these provisions, including increased protection of our potential ability to negotiate with the proponent of an unfriendly or unsolicited proposal to acquire or restructure our company, outweigh the disadvantages of discouraging takeover proposals, because negotiation of takeover proposals could result in an improvement of their terms.

Choice of Forum

Our restated certificate will provide that the Court of Chancery of the State of Delaware will be the exclusive forum for:

- any derivative action or proceeding brought on our behalf;
- any action asserting a breach of fiduciary duty;
- any action asserting a claim against us arising pursuant to the Delaware General Corporation Law, our restated certificate, or our amended and restated bylaws; or
- any action asserting a claim against us that is governed by the internal affairs doctrine.

The enforceability of similar choice of forum provisions in other companies' certificates of incorporation has been challenged in legal proceedings, and it is possible that, in connection with any action, a court could find the choice of forum provisions contained in our restated certificate to be inapplicable or unenforceable in such action.

Transfer Agent and Registrar

The transfer agent and registrar for our common stock is American Stock Transfer & Trust Company. The transfer agent's address is 6201 15th Avenue, Brooklyn, NY 11219.

NASDAQ Global Market Listing

We have applied for listing of our common stock on The NASDAQ Global Market under the trading symbol "CLSD."

SHARES ELIGIBLE FOR FUTURE SALE

Prior to this offering, no public market existed for our common stock. Future sales of shares of our common stock in the public market after this offering, or the perception that these sales could occur, could adversely affect prevailing market prices for our common stock and could impair our future ability to raise equity capital.

Based on the number of shares outstanding as of August 31, 2014, upon completion of this offering and assuming no exercise of the underwriters' option to purchase additional shares, _____ shares of common stock will be outstanding, assuming no outstanding options or warrants are exercised. All of the shares of common stock sold in this offering will be freely tradable without restrictions or further registration under the Securities Act, except for any shares sold to our "affiliates," as that term is defined under Rule 144 under the Securities Act. The remaining shares of common stock held by existing stockholders are "restricted securities," as that term is defined in Rule 144 under the Securities Act. Restricted securities may be sold in the public market only if registered or if their resale qualifies for exemption from registration described below under Rule 144 promulgated under the Securities Act.

As a result of contractual restrictions described below and the provisions of Rules 144 and 701, the shares sold in this offering and the restricted securities will be available for sale in the public market as follows:

- the _____ shares sold in this offering and _____ of the existing restricted shares will be eligible for immediate sale upon the completion of this offering;
- approximately _____ restricted shares will be eligible for sale in the public market 90 days after the date of this prospectus, subject to the volume, manner of sale and other limitations under Rule 144 and Rule 701; and
- approximately _____ restricted shares will be eligible for sale in the public market upon expiration of lock-up agreements 180 days after the date of this prospectus, subject in certain circumstances to the volume, manner of sale and other limitations under Rule 144 and Rule 701.

Rule 144

In general, persons who have beneficially owned restricted shares of our common stock for at least six months, and any affiliate of the company who owns either restricted or unrestricted shares of our common stock, are entitled to sell their securities without registration with the SEC under an exemption from registration provided by Rule 144 under the Securities Act.

Non-Affiliates

Any person who is not deemed to have been one of our affiliates at the time of, or at any time during the three months preceding, a sale may sell an unlimited number of restricted securities under Rule 144 if:

- the restricted securities have been held for at least six months, including the holding period of any prior owner other than one of our affiliates;
- we have been subject to the Exchange Act periodic reporting requirements for at least 90 days before the sale; and
- we are current in our Exchange Act reporting at the time of sale.

[Table of Contents](#)

Any person who is not deemed to have been an affiliate of ours at the time of, or at any time during the three months preceding, a sale and has held the restricted securities for at least one year, including the holding period of any prior owner other than one of our affiliates, will be entitled to sell an unlimited number of restricted securities without regard to the length of time we have been subject to Exchange Act periodic reporting or whether we are current in our Exchange Act reporting.

Affiliates

Persons seeking to sell restricted securities who are our affiliates at the time of, or any time during the three months preceding, a sale, would be subject to the restrictions described above. They are also subject to additional restrictions, by which such person would be required to comply with the manner of sale and notice provisions of Rule 144 and would be entitled to sell within any three-month period only that number of securities that does not exceed the greater of either of the following:

- 1% of the number of shares of our common stock then outstanding, which will equal approximately _____ shares immediately after the completion of this offering based on the number of shares outstanding as of August 31, 2014; or
- the average weekly trading volume of our common stock on The NASDAQ Global Market during the four calendar weeks preceding the filing of a notice on Form 144 with respect to the sale.

Additionally, persons who are our affiliates at the time of, or any time during the three months preceding, a sale may sell unrestricted securities under the requirements of Rule 144 described above, without regard to the six month holding period of Rule 144, which does not apply to sales of unrestricted securities.

Rule 701

Rule 701 under the Securities Act, as in effect on the date of this prospectus, permits resales of shares in reliance upon Rule 144 but without compliance with certain restrictions of Rule 144, including the holding period requirement. Most of our employees, executive officers or directors who purchased shares under a written compensatory plan or contract may be entitled to rely on the resale provisions of Rule 701, but all holders of Rule 701 shares are required to wait until 90 days after the date of this prospectus before selling their shares. However, substantially all Rule 701 shares are subject to lock-up agreements as described below and in the section of this prospectus titled "Underwriting" and will become eligible for sale upon the expiration of the restrictions set forth in those agreements.

Form S-8 Registration Statements

As soon as practicable after the completion of this offering, we intend to file with the SEC one or more registration statements on Form S-8 under the Securities Act to register the shares of our common stock that are issuable pursuant to our 2011 plan, 2014 plan and 2014 ESPP. These registration statements will become effective immediately upon filing. Shares covered by these registration statements will then be eligible for sale in the public markets, subject to vesting restrictions, any applicable lock-up agreements described below and Rule 144 limitations applicable to affiliates.

Lock-Up Agreements

We and the holders of substantially all of our common stock outstanding on the date of this prospectus, including each of our executive officers and directors, have entered into lock-up agreements with the

[Table of Contents](#)

underwriters or otherwise agreed, subject to certain exceptions, that we and they will not, directly or indirectly, offer, sell, contract to sell, pledge, grant any option to purchase, make any short sale, or otherwise dispose of or hedge any of our shares of common stock, any options or warrants to purchase shares of our common stock, or any securities convertible into, or exchangeable for or that represent the right to receive shares of our common stock, without the prior written consent of the representatives of the underwriters for a period of 180 days from the date of this prospectus.

MATERIAL U.S. FEDERAL INCOME AND ESTATE TAX CONSEQUENCES TO NON-U.S. HOLDERS

The following is a general discussion of the material U.S. federal income and estate tax considerations applicable to non-U.S. holders with respect to their ownership and disposition of shares of our common stock issued pursuant to this offering. All prospective non-U.S. holders of our common stock should consult their own tax advisors with respect to the U.S. federal, state, local and non-U.S. tax consequences of the purchase, ownership and disposition of our common stock. In general, a non-U.S. holder means a beneficial owner of our common stock (other than a partnership or an entity or arrangement treated as a partnership for U.S. federal income tax purposes) that is not, for U.S. federal income tax purposes:

- an individual who is a citizen or resident of the United States;
- a corporation, or an entity treated as a corporation for U.S. federal income tax purposes, created or organized in the United States or under the laws of the United States or of any state thereof or the District of Columbia;
- an estate, the income of which is subject to U.S. federal income tax regardless of its source; or
- a trust if (1) a U.S. court can exercise primary supervision over the trust's administration and one or more U.S. persons have the authority to control all of the trust's substantial decisions or (2) the trust has a valid election in effect under applicable U.S. Treasury Regulations to be treated as a U.S. person.

This discussion is based on current provisions of the U.S. Internal Revenue Code of 1986, as amended, which we refer to as the Code, existing U.S. Treasury Regulations promulgated thereunder, published administrative rulings and judicial decisions, all as in effect as of the date of this prospectus. These laws are subject to change and to differing interpretation, possibly with retroactive effect. Any change or differing interpretation could alter the tax consequences to non-U.S. holders described in this prospectus.

We assume in this discussion that a non-U.S. holder holds shares of our common stock as a capital asset within the meaning of Section 1221 of the Code (generally, for investment). This discussion does not address all aspects of U.S. federal income and estate taxation that may be relevant to a particular non-U.S. holder in light of that non-U.S. holder's individual circumstances, nor does it address any aspects of U.S. state, local or non-U.S. taxes. This discussion also does not consider any specific facts or circumstances that may apply to a non-U.S. holder and does not address the special tax rules applicable to particular non-U.S. holders, such as holders that own, or are deemed to own, more than 5% of our capital stock (except to the extent specifically set forth below), corporations that accumulate earnings to avoid U.S. federal income tax, tax-exempt organizations, banks, financial institutions, insurance companies, brokers, dealers or traders in securities, commodities or currencies, tax-qualified retirement plans, holders subject to the alternative minimum tax or the Medicare contribution tax, holders who hold or receive our common stock pursuant to the exercise of employee stock options or otherwise as compensation, holders holding our common stock as part of a hedge, straddle or other risk reduction strategy, conversion transaction or other integrated investment, holders deemed to sell our common stock under the constructive sale provisions of the Code, controlled foreign corporations, passive foreign investment companies and certain former U.S. citizens or long-term residents.

In addition, this discussion does not address the tax treatment of partnerships (or entities or arrangements that are treated as partnerships for U.S. federal income tax purposes) or persons that hold their common stock through such partnerships. If a partnership, including any entity or arrangement treated as a

partnership for U.S. federal income tax purposes, holds shares of our common stock, the U.S. federal income tax treatment of a partner in such partnership will generally depend upon the status of the partner and the activities of the partnership. Such partners and partnerships should consult their own tax advisors regarding the tax consequences of the purchase, ownership and disposition of our common stock.

There can be no assurance that the Internal Revenue Service, which we refer to as the IRS, will not challenge one or more of the tax consequences described herein, and we have not obtained, nor do we intend to obtain, a ruling with respect to the U.S. federal income or estate tax consequences to a non-U.S. holder of the purchase, ownership or disposition of our common stock.

Distributions on Our Common Stock

Distributions, if any, on our common stock generally will constitute dividends for U.S. federal income tax purposes to the extent paid from our current or accumulated earnings and profits, as determined under U.S. federal income tax principles. If a distribution exceeds our current and accumulated earnings and profits, the excess will be treated as a tax-free return of the non-U.S. holder's investment, up to such holder's adjusted tax basis in the common stock. Any remaining excess will be treated as capital gain from the sale or exchange of such common stock, subject to the tax treatment described below in "Gain on Sale, Exchange or Other Disposition of Our Common Stock." Any such distribution will also be subject to the discussion below under the heading "Foreign Accounts."

Dividends paid to a non-U.S. holder will generally be subject to withholding of U.S. federal income tax at a 30% rate or such lower rate as may be specified by an applicable income tax treaty between the United States and such holder's country of residence.

Dividends that are treated as effectively connected with a trade or business conducted by a non-U.S. holder within the United States and, if an applicable income tax treaty so provides, that are attributable to a permanent establishment or a fixed base maintained by the non-U.S. holder within the United States, are generally exempt from the 30% withholding tax if the non-U.S. holder satisfies applicable certification and disclosure requirements. However, such U.S. effectively connected income, net of specified deductions and credits, is taxed at the same graduated U.S. federal income tax rates applicable to U.S. persons (as defined in the Code). Any U.S. effectively connected income received by a non-U.S. holder that is a corporation may also, under certain circumstances, be subject to an additional "branch profits tax" at a 30% rate or such lower rate as may be specified by an applicable income tax treaty between the United States and such holder's country of residence.

A non-U.S. holder of our common stock who claims the benefit of an applicable income tax treaty between the United States and such holder's country of residence generally will be required to provide a properly executed IRS Form W-8BEN or W-8BEN-E (or successor form) and satisfy applicable certification and other requirements. Non-U.S. holders are urged to consult their tax advisors regarding their entitlement to benefits under a relevant income tax treaty.

A non-U.S. holder that is eligible for a reduced rate of U.S. withholding tax under an income tax treaty may obtain a refund or credit of any excess amounts withheld by timely filing an appropriate claim for refund with the IRS.

Gain on Sale, Exchange or Other Disposition of Our Common Stock

Subject to the discussion below regarding backup withholding and foreign accounts, in general, a non-U.S. holder will not be subject to any U.S. federal income tax on any gain realized upon such holder's sale, exchange or other disposition of shares of our common stock unless:

- the gain is effectively connected with a U.S. trade or business of the non-U.S. holder and, if an applicable income tax treaty so provides, is attributable to a permanent establishment or a fixed base maintained in the United States by such non-U.S. holder, in which case the non-U.S. holder generally will be taxed at the graduated U.S. federal income tax rates applicable to U.S. persons (as defined in the Code) and, if the non-U.S. holder is a foreign corporation, the branch profits tax described above in "Distributions on Our Common Stock" also may apply;
- the non-U.S. holder is a nonresident alien individual who is present in the United States for 183 days or more in the taxable year of the disposition and certain other conditions are met, in which case the non-U.S. holder will be subject to a 30% tax (or such lower rate as may be specified by an applicable income tax treaty) on the net gain derived from the disposition, which may be offset by U.S. source capital losses of the non-U.S. holder, if any (even though the individual is not considered a resident of the United States); or
- our common stock constitutes a U.S. real property interest because we are, or have been, at any time during the five-year period preceding such disposition (or the non-U.S. holder's holding period, if shorter) a "U.S. real property holding corporation." Even if we are or become a U.S. real property holding corporation, provided that our common stock is regularly traded on an established securities market, our common stock will be treated as a U.S. real property interest only with respect to a non-U.S. holder that holds more than 5% of our outstanding common stock, directly or indirectly, actually or constructively, during the shorter of the 5-year period ending on the date of the disposition or the period that the non-U.S. holder held our common stock. In such case, such non-U.S. holder generally will be taxed on its net gain derived from the disposition at the graduated U.S. federal income tax rates applicable to U.S. persons (as defined in the Code). Generally, a corporation is a U.S. real property holding corporation only if the fair market value of its U.S. real property interests equals or exceeds 50% of the sum of the fair market value of its worldwide real property interests plus its other assets used or held for use in a trade or business. Although there can be no assurance, we do not believe that we are, or have been, a U.S. real property holding corporation, or that we are likely to become one in the future. No assurance can be provided that our common stock will be regularly traded on an established securities market for purposes of the rules described above.

U.S. Federal Estate Tax

Shares of our common stock that are owned or treated as owned at the time of death by an individual who is not a citizen or resident of the United States, as specifically defined for U.S. federal estate tax purposes, are considered U.S. situs assets and will be included in the individual's gross estate for U.S. federal estate tax purposes. Such shares, therefore, may be subject to U.S. federal estate tax, unless an applicable estate tax or other treaty provides otherwise.

Backup Withholding and Information Reporting

We must report annually to the IRS and to each non-U.S. holder the gross amount of the dividends on our common stock paid to such holder and the tax withheld, if any, with respect to such dividends. Non-

[Table of Contents](#)

U.S. holders will have to comply with specific certification procedures to establish that the holder is not a U.S. person (as defined in the Code) in order to avoid backup withholding at the applicable rate with respect to dividends on our common stock. Dividends paid to non-U.S. holders subject to the U.S. withholding tax, as described above in “Distributions on Our Common Stock,” generally will be exempt from U.S. backup withholding.

Information reporting and backup withholding will generally apply to the proceeds of a disposition of our common stock by a non-U.S. holder effected by or through the U.S. office of any broker, U.S. or foreign, unless the holder certifies its status as a non-U.S. holder and satisfies certain other requirements, or otherwise establishes an exemption. Generally, information reporting and backup withholding will not apply to a payment of disposition proceeds to a non-U.S. holder where the transaction is effected outside the United States through a non-U.S. office of a broker. However, for information reporting purposes, dispositions effected through a non-U.S. office of a broker with substantial U.S. ownership or operations generally will be treated in a manner similar to dispositions effected through a U.S. office of a broker. Non-U.S. holders should consult their own tax advisors regarding the application of the information reporting and backup withholding rules to them.

Copies of information returns may be made available to the tax authorities of the country in which the non-U.S. holder resides or is incorporated under the provisions of a specific treaty or agreement.

Backup withholding is not an additional tax. Any amounts withheld under the backup withholding rules from a payment to a non-U.S. holder may be allowed as a credit against the non-U.S. holder’s U.S. federal income tax liability, if any, and may entitle such holder to a refund, provided that the required information is timely furnished to the IRS.

Foreign Accounts

The Code generally imposes a U.S. federal withholding tax of 30% on dividends and the gross proceeds of a disposition of our common stock paid to a “foreign financial institution” (as specifically defined for this purpose), unless such institution enters into an agreement with the U.S. government to, among other things, withhold on certain payments and to collect and provide to the U.S. tax authorities substantial information regarding U.S. account holders of such institution (which includes certain equity and debt holders of such institution, as well as certain account holders that are foreign entities with U.S. owners). A U.S. federal withholding tax of 30% also applies to dividends and the gross proceeds of a disposition of our common stock paid to a non-financial foreign entity, unless such entity provides the withholding agent with either a certification that it does not have any substantial direct or indirect U.S. owners or provides information regarding substantial direct and indirect U.S. owners of the entity. The withholding provisions described above will generally apply to dividends on our common stock paid on or after July 1, 2014 and with respect to gross proceeds of a sale or other disposition of our common stock on or after January 1, 2017. Under certain circumstances, a non-U.S. holder might be eligible for refunds or credits of such taxes.

UNDERWRITING

Subject to the terms and conditions set forth in the underwriting agreement, to be dated the date of this prospectus, between us and RBC Capital Markets, LLC and Wells Fargo Securities, LLC, as the representatives of the underwriters named below and the joint book-running managers of this offering, we have agreed to sell to the underwriters, and each of the underwriters has agreed, severally and not jointly, to purchase from us, the respective number of shares of common stock shown opposite its name below:

<u>Underwriter</u>	<u>Number of Shares</u>
RBC Capital Markets, LLC	
Wells Fargo Securities, LLC	
Needham & Company, LLC	
Oppenheimer & Co. Inc.	
Total	

The underwriting agreement provides that the obligations of the several underwriters are subject to certain conditions precedent such as the receipt by the underwriters of officers' certificates and legal opinions and approval of certain legal matters by their counsel. The underwriting agreement provides that the underwriters will purchase all of the shares of common stock if any of them are purchased. If an underwriter defaults, the underwriting agreement provides that the purchase commitments of the non-defaulting underwriters may be increased or the underwriting agreement may be terminated. We have agreed to indemnify the underwriters, their affiliates and certain of their controlling persons against certain liabilities, including liabilities under the Securities Act, and to contribute to payments that the underwriters may be required to make in respect of those liabilities.

The underwriters have advised us that, following the completion of this offering, they currently intend to make a market in the common stock as permitted by applicable laws and regulations. However, the underwriters are not obligated to do so, and the underwriters may discontinue any market-making activities at any time without notice in their sole discretion. Accordingly, no assurance can be given as to the liquidity of the trading market for the common stock, that you will be able to sell any of the common stock held by you at a particular time or that the prices that you receive when you sell will be favorable.

The underwriters are offering the shares of common stock subject to their acceptance of the shares of common stock from us and subject to prior sale. The underwriters reserve the right to withdraw, cancel or modify offers to the public and to reject orders in whole or in part. In addition, the underwriters have advised us that they do not expect sales to accounts over which they have discretionary authority to exceed 5% of the common stock being offered.

Commissions and Expenses

The underwriters have advised us that they propose to offer the shares of common stock to the public at the initial public offering price set forth on the cover page of this prospectus and to certain dealers, which may include the underwriters, at that price less a concession not in excess of \$ per share of common stock. After the offering, the initial public offering price and the concession to dealers may be reduced by the representatives. No such reduction will change the amount of proceeds to be received by us as set forth on the cover page of this prospectus.

[Table of Contents](#)

The following table shows the public offering price, the underwriting discounts and commissions that we are to pay the underwriters and the proceeds, before expenses, to us in connection with this offering. Such amounts are shown assuming both no exercise and full exercise of the underwriters' option to purchase additional shares.

	Per Share		Total	
	Without Option To Purchase Additional Shares	With Option To Purchase Additional Shares	Without Option To Purchase Additional Shares	With Option To Purchase Additional Shares
Public offering price	\$	\$	\$	\$
Underwriting discounts and commissions paid by us	\$	\$	\$	\$
Proceeds to us, before expenses	\$	\$	\$	\$

We estimate expenses payable by us in connection with this offering, other than the underwriting discounts and commissions referred to above, will be approximately \$ million. We have agreed to reimburse the underwriters for certain expenses in an amount up to \$.

Determination of Offering Price

Prior to this offering, there has not been a public market for our common stock. Consequently, the initial public offering price for our common stock will be determined by negotiations between us and the representatives. Among the factors to be considered in these negotiations will be prevailing market conditions, our financial information, market valuations of other companies that we and the underwriters believe to be comparable to us, estimates of our business potential, the present state of our development and other factors deemed relevant.

We offer no assurances that the initial public offering price will correspond to the price at which the common stock will trade in the public market subsequent to the offering or that an active trading market for the common stock will develop and continue after the offering.

Listing

We have applied to list our common stock on The NASDAQ Global Market under the symbol "CLSD."

Option to Purchase Additional Shares

We have granted to the underwriters an option, exercisable for 30 days from the date of this prospectus, to purchase, from time to time, in whole or in part, up to an aggregate of shares from us at the public offering price set forth on the cover page of this prospectus, less underwriting discounts and commissions. If the underwriters exercise this option, each underwriter will be obligated, subject to specified conditions, to purchase a number of additional shares proportionate to that underwriter's initial purchase commitment as indicated in the table above. This option may be exercised only if the underwriters sell more shares than the total number set forth on the cover page of this prospectus.

No Sales of Similar Securities

We, our officers, directors and holders of substantially all of our outstanding capital stock have agreed, subject to specified exceptions, not to directly or indirectly:

- sell, offer to sell, contract to sell, effect any short sale, pledge, transfer, establish a “put equivalent position” within the meaning of Rule 16a-1(h) under the Exchange Act;
- otherwise dispose of, or enter into any swap, hedge or similar arrangement that transfers, in whole or in part, the economic risk of ownership of, any shares of common stock, options or warrants to acquire shares of common stock, or securities exchangeable or exercisable for or convertible into shares of common stock currently or hereafter owned either of record or beneficially;
- make any demand for, or exercise any right with respect to, the registration under the Securities Act of the offer and sale of any shares of common stock, options or warrants to acquire shares of common stock, or securities exchangeable or exercisable for or convertible into shares of common stock; or
- publicly announce any intention to do any of the foregoing;

for a period of 180 days after the date of this prospectus without the prior written consent of the representatives.

The lock-up restrictions terminate after the close of trading of the common stock on and including the 180th day after the date of this prospectus. The representatives may, in their sole discretion and at any time or from time to time before the termination of the 180-day period release all or any portion of the securities subject to lock-up agreements.

The restrictions described above do not apply to:

- transfers of shares of our common stock or any security convertible into or exercisable or exchangeable for shares of our common stock (i) as a bona fide gift, or gifts, (ii) to an immediate family member or a trust for the direct or indirect benefit of the lock-up signatory or such immediate family member of the lock-up signatory or (iii) by will or intestacy;
- transactions relating to shares of our common stock or other securities acquired in the open market after the completion of this offering;
- if the lock-up signatory is a corporation, limited partnership, trust or other business entity, transfers of shares of our common stock to (i) another corporation, member, partner, trust or other business entity that is a direct or indirect affiliate (as defined in Rule 405 promulgated under the Securities Act) of the lock-up signatory or (ii) as part of a pro rata distribution or transfer by the lock-up signatory to its stockholders, members, partners, beneficiaries or other equity holders provided however, that any such transfer or distribution shall not involve a disposition for value;
- transfers to us in connection with the “cashless” exercise of options to purchase shares of our common stock pursuant to employee benefit plans disclosed in this prospectus;
- transfers in connection with the “net exercise” of warrants held by the lock-up signatory;
- transfers to us to satisfy tax withholding obligations in connection with the vesting or exercise of equity incentive awards under our employee benefit plans after the completion of this offering;

[Table of Contents](#)

- transfers to us in connection with the repurchase of shares of our common stock issued pursuant to employee benefit plans disclosed in this prospectus upon the termination of service pursuant to an existing company right;
- transfers, sales, tenders or other dispositions of shares of our common stock, or any securities convertible into or exercisable or exchangeable for our common stock, occurring after the consummation of this offering, pursuant to a bona fide third party tender offer, merger, consolidation or other similar transaction made to all holders of our capital stock that would result in the disposition of not less than a majority of the outstanding shares of our voting securities, provided that in the event that the tender offer, merger, consolidation or other such transaction is not completed, our common stock and any security convertible into or exchangeable for our common stock shall remain subject to the same restrictions;
- transfers pursuant to operation of law, including pursuant to a domestic order or negotiated divorce settlement;
- any issuance by us of shares of our common stock or securities convertible or exercisable or exchangeable for shares of our common stock pursuant to the exercise or conversion of warrants, options, or other convertible or exchangeable securities, in each case outstanding as of the date of this prospectus; and
- the establishment of a trading plan that complies with Rule 10b5-1 under the Exchange Act; provided, however, that (i) the restrictions shall apply in full force to sales or other dispositions pursuant to such Rule 10b5-1 plan during the 180-day lock-up period described above and (ii) no public announcement or disclosure of entry into such Rule 10b5-1 plan is made or required to be made;

provided, however, that in the case of any transfer or distribution pursuant to the first, third, ninth and tenth clauses above, each donee, distributee recipient or transferee shall sign and deliver a lock-up agreement substantially in the form of the lock-up agreements described above; and in the case of any transfer or distribution pursuant to the first (except for sub-clause (iii)), second, third, fourth, fifth, sixth and eleventh clauses above, no filing under Section 16(a) of the Exchange Act, reporting a reduction in beneficial ownership of shares of our common stock, shall be required or shall be voluntarily made during the restricted period.

There are no existing agreements between the underwriters and any of our stockholders who will execute a lock-up agreement, providing consent to the sale of shares prior to the expiration of the lock-up period.

Stabilization

The underwriters have advised us that they, pursuant to Regulation M under the Exchange Act, certain persons participating in the offering may engage in short sale transactions, stabilizing transactions, syndicate covering transactions or the imposition of penalty bids in connection with this offering. These activities may have the effect of stabilizing or maintaining the market price of the common stock at a level above that which might otherwise prevail in the open market. Establishing short sales positions may involve either “covered” short sales or “naked” short sales.

“Covered” short sales are sales made in an amount not greater than the underwriters’ option to purchase additional shares of our common stock in this offering. The underwriters may close out any

[Table of Contents](#)

covered short position by either exercising their option to purchase additional shares of our common stock or purchasing shares of our common stock in the open market. In determining the source of shares to close out the covered short position, the underwriters will consider, among other things, the price of shares available for purchase in the open market as compared to the price at which they may purchase shares through the option to purchase additional shares.

“Naked” short sales are sales in excess of the option to purchase additional shares of our common stock. The underwriters must close out any naked short position by purchasing shares in the open market. A naked short position is more likely to be created if the underwriters are concerned that there may be downward pressure on the price of the shares of our common stock in the open market after pricing that could adversely affect investors who purchase in this offering.

A stabilizing bid is a bid for the purchase of shares of common stock on behalf of the underwriters for the purpose of fixing or maintaining the price of the common stock. A syndicate covering transaction is the bid for or the purchase of shares of common stock on behalf of the underwriters to reduce a short position incurred by the underwriters in connection with the offering. Similar to other purchase transactions, the underwriter’s purchases to cover the syndicate short sales may have the effect of raising or maintaining the market price of our common stock or preventing or retarding a decline in the market price of our common stock. As a result, the price of our common stock may be higher than the price that might otherwise exist in the open market. A penalty bid is an arrangement permitting the underwriters to reclaim the selling concession otherwise accruing to a syndicate member in connection with the offering if the common stock originally sold by such syndicate member are purchased in a syndicate covering transaction and therefore have not been effectively placed by such syndicate member.

Neither we nor any of the underwriters make any representation or prediction as to the direction or magnitude of any effect that the transactions described above may have on the price of our common stock. The underwriters are not obligated to engage in these activities and, if commenced, any of the activities may be discontinued at any time.

Electronic Distribution

A prospectus in electronic format may be made available by e-mail or through online services maintained by one or more of the underwriters or their affiliates. In those cases, prospective investors may view offering terms online and may be allowed to place orders online. The underwriters may agree with us to allocate a specific number of shares of common stock for sale to online brokerage account holders. Any such allocation for online distributions will be made by the underwriters on the same basis as other allocations. Other than the prospectus in electronic format, the information on the underwriters’ websites and any information contained in any other website maintained by any of the underwriters is not part of this prospectus, has not been approved or endorsed by us or the underwriters and should not be relied upon by investors.

Other Activities and Relationships

The underwriters and certain of their respective affiliates are full service financial institutions engaged in various activities, which may include securities trading, commercial and investment banking, financial advisory, investment management, investment research, principal investment, hedging, financing and brokerage activities. The underwriters and certain of their respective affiliates have, from time to time, performed, and may in the future perform, various commercial and investment banking and financial advisory services for us and our affiliates, for which they received or will receive customary fees and expenses.

[Table of Contents](#)

In the ordinary course of their various business activities, the underwriters and certain of their respective affiliates may make or hold a broad array of investments and actively trade debt and equity securities (or related derivative securities) and financial instruments (including bank loans) for their own account and for the accounts of their customers, and such investment and securities activities may involve securities or instruments issued by us and our affiliates. If the underwriters or their respective affiliates have a lending relationship with us, they routinely hedge their credit exposure to us consistent with their customary risk management policies. The underwriters and their respective affiliates may hedge such exposure by entering into transactions which consist of either the purchase of credit default swaps or the creation of short positions in our securities or the securities of our affiliates, including potentially the common stock offered hereby. Any such short positions could adversely affect future trading prices of the common stock offered hereby. The underwriters and certain of their respective affiliates may also communicate independent investment recommendations, market color or trading ideas or publish or express independent research views in respect of such securities or instruments and may at any time hold, or recommend to clients that they acquire, long or short positions in such securities and instruments.

Notice to Residents of Canada

The securities may be sold only to purchasers purchasing as principal that are both “accredited investors” as defined in National Instrument 45-106 Prospectus and Registration Exemptions and “permitted clients” as defined in National Instrument 31-103 Registration Requirements, Exemptions and Ongoing Registrant Obligations. Any resale of the securities must be made in accordance with an exemption from the prospectus requirements and in compliance with the registration requirements of applicable securities laws.

Notice to Prospective Investors in the EEA

In relation to each Member State of the European Economic Area which has implemented the Prospectus Directive (each, a “Relevant Member State”), including each Relevant Member State that has implemented amendments to Article 3(2) of the Prospectus Directive introduced by the 2010 PD Amending Directive (each, an “Early Implementing Member State”), an offer of shares of common stock to the public may not be made in that Relevant Member State and each initial purchaser represents and agrees that with effect from and including the date on which the Prospectus Directive is implemented in that Relevant Member State (the “Relevant Implementation Date”) it has not made and will not make an offer of the shares of common stock to the public in that Relevant Member State prior to the publication of a prospectus in relation to the shares of common stock which has been approved by the competent authority in that Relevant Member State or, where appropriate, approved in another Relevant Member State and notified to the competent authority in that Relevant Member State, all in accordance with the Prospectus Directive, except that an offer of the shares of common stock to the public in that Relevant Member State may be made at any time under the following exemptions under the Prospectus Directive, if they have been implemented in that Relevant Member State:

- (a) to any legal entity which is a “qualified investor” as defined in the Prospectus Directive;
- (b) to fewer than 100 (or, in the case of Early Implementing Member States, 150) natural or legal persons (other than “qualified investors” as defined in the Prospectus Directive) subject to obtaining the prior consent of the Subscribers; or
- (c) in any other circumstances falling within Article 3(2) of the Prospectus Directive,

provided that no such offer of the shares of common stock referred to in (a) to (c) above shall require the issuer or any Subscriber to publish a prospectus pursuant to Article 3 of the Prospectus Directive or

supplement a prospectus pursuant to Article 16 of the Prospectus Directive and each person who initially acquires any shares of common stock or to whom any offer is made will be deemed to have represented, warranted and agreed to and with the issuer or any Subscriber that it is a qualified investor within the meaning of the law in that Relevant Member State implementing Article 2(1)(e) of the Prospectus Directive.

For the purposes of this provision, the expression an “offer to the public” in relation to any shares of common stock in any Relevant Member State means the communication in any form and by any means of sufficient information on the terms of the offer and the shares of common stock to be offered so as to enable an investor to decide to purchase or subscribe to the shares of common stock, as the same may be varied in that Relevant Member State by any measure implementing the Prospectus Directive in that Relevant Member State and the expression “Prospectus Directive” means Directive 2003/71/EC of the European Parliament and of the Council of 4 November 2003 (and amendments thereto, including the 2010 PD Amending Directive, to the extent implemented in the Relevant Member State), and includes any relevant implementing measure in the Relevant Member State. The expression “2010 PD Amending Directive” means Directive 2010/73/EU.

Notice to Prospective Investors in Hong Kong

The shares may not be offered or sold by means of any document other than (i) in circumstances which do not constitute an offer to the public within the meaning of the Companies Ordinance (Cap.32, Laws of Hong Kong), or (ii) to “professional investors” within the meaning of the Securities and Futures Ordinance (Cap.571, Laws of Hong Kong) and any rules made thereunder, or (iii) in other circumstances which do not result in the document being a “prospectus” within the meaning of the Companies Ordinance (Cap.32, Laws of Hong Kong), and no advertisement, invitation or document relating to the shares may be issued or may be in the possession of any person for the purpose of issue (in each case whether in Hong Kong or elsewhere), which is directed at, or the contents of which are likely to be accessed or read by, the public in Hong Kong (except if permitted to do so under the laws of Hong Kong) other than with respect to shares which are or are intended to be disposed of only to persons outside Hong Kong or only to “professional investors” within the meaning of the Securities and Futures Ordinance (Cap.571 Laws of Hong Kong) and any rules made thereunder.

Notice to Prospective Investors in Israel

The securities offered by this prospectus have not been approved or disapproved by the Israeli Securities Authority, or the ISA, nor have such securities been registered for sale in Israel. The shares of common stock may not be offered or sold, directly or indirectly, to the public in Israel, absent the publication of a prospectus. The ISA has not issued permits, approvals or licenses in connection with the Offering or publishing the prospectus; nor has it authenticated the details included herein, confirmed their reliability or completeness, or rendered an opinion as to the quality of the securities being offered. Any resale in Israel, directly or indirectly, to the public of the securities offered by this prospectus is subject to restrictions on transferability and must be effected only in compliance with the Israeli securities laws and regulations.

Notice to Prospective Investors in Japan

The securities have not been and will not be registered under the Financial Instruments and Exchange Law of Japan (the Financial Instruments and Exchange Law) and may not be offered or sold, directly or indirectly, in Japan or to, or for the benefit of, any resident of Japan (which term as used herein means any person resident in Japan, including any corporation or other entity organized under the laws of Japan), or to

[Table of Contents](#)

others for re-offering or resale, directly or indirectly, in Japan or to a resident of Japan, except pursuant to an exemption from the registration requirements of, and otherwise in compliance with, the Financial Instruments and Exchange Law and any other applicable laws, regulations and ministerial guidelines of Japan.

Notice to Prospective Investors in Singapore

This prospectus has not been registered as a prospectus with the Monetary Authority of Singapore. Accordingly, this prospectus and any other document or material in connection with the offer or sale, or invitation for subscription or purchase, of the shares may not be circulated or distributed, nor may the shares be offered or sold, or be made the subject of an invitation for subscription or purchase, whether directly or indirectly, to persons in Singapore other than (i) to an institutional investor under Section 274 of the Securities and Futures Act, Chapter 289 of Singapore (SFA), (ii) to a relevant person, or any person pursuant to Section 275(1A), and in accordance with the conditions, specified in Section 275 of the SFA or (iii) otherwise pursuant to, and in accordance with the conditions of, any other applicable provision of the SFA.

Where the shares are subscribed or purchased under Section 275 by a relevant person which is: (a) a corporation (which is not an accredited investor) the sole business of which is to hold investments and the entire share capital of which is owned by one or more individuals, each of whom is an accredited investor; or (b) a trust (where the trustee is not an accredited investor) whose sole purpose is to hold investments and each beneficiary is an accredited investor, shares, debentures and units of shares and debentures of that corporation or the beneficiaries' rights and interest in that trust shall not be transferable for six months after that corporation or that trust has acquired the shares under Section 275 except: (1) to an institutional investor under Section 274 of the SFA or to a relevant person, or any person pursuant to Section 275(1A), and in accordance with the conditions, specified in Section 275 of the SFA; (2) where no consideration is given for the transfer; or (3) by operation of law.

Notice to Prospective Investors in the United Kingdom

In addition, in the United Kingdom, this document is being distributed only to, and is directed only at, and any offer subsequently made may only be directed at persons who are "qualified investors" (as defined in the Prospectus Directive) (i) who have professional experience in matters relating to investments falling within Article 19(5) of the Financial Services and Markets Act 2000 (Financial Promotion) Order 2005, as amended, which we refer to as the Order, or (ii) who are high net worth companies (or persons to whom it may otherwise be lawfully communicated) falling within Article 49(2)(a) to (d) of the Order (all such persons together being referred to as "relevant persons"). This document must not be acted on or relied on in the United Kingdom by persons who are not relevant persons. In the United Kingdom, any investment or investment activity to which this document relates is only available to, and will be engaged in with, relevant persons.

Notice to Prospective Investors in Switzerland

The Prospectus does not constitute an issue prospectus pursuant to Article 652a or Article 1156 of the Swiss Code of Obligations ("CO") and the shares will not be listed on the SIX Swiss Exchange. Therefore, the Prospectus may not comply with the disclosure standards of the CO or the listing rules (including any prospectus schemes) of the SIX Swiss Exchange. Accordingly, the shares may not be offered to the public in or from Switzerland, but only to a selected and limited circle of investors, which do not subscribe to the shares with a view to distribution.

LEGAL MATTERS

The validity of the shares of common stock being offered by this prospectus will be passed upon for us by Cooley LLP, Reston, Virginia. Certain legal matters related to this offering will be passed upon for the underwriters by Latham & Watkins LLP, Boston, Massachusetts.

EXPERTS

The financial statements of Clearside Biomedical, Inc. at December 31, 2012 and 2013, and for each of the two years in the period ended December 31, 2013, and for the period from May 26, 2011 (date of inception) to December 31, 2013, appearing in this prospectus and registration statement have been audited by Ernst & Young LLP, independent registered public accounting firm, as set forth in their report thereon appearing elsewhere herein, and are included in reliance upon such report given on the authority of such firm as experts in accounting and auditing.

WHERE YOU CAN FIND ADDITIONAL INFORMATION

We have filed with the SEC a registration statement on Form S-1 under the Securities Act, with respect to the shares of common stock being offered by this prospectus. This prospectus, which constitutes part of the registration statement, does not contain all of the information in the registration statement and its exhibits. For further information with respect to our company and the common stock offered by this prospectus, we refer you to the registration statement and its exhibits. Statements contained in this prospectus as to the contents of any contract or any other document referred to are not necessarily complete, and in each instance, we refer you to the copy of the contract or other document filed as an exhibit to the registration statement. Each of these statements is qualified in all respects by this reference.

You can read our SEC filings, including the registration statement, over the internet at the SEC's website at www.sec.gov. You may also read and copy any document we file with the SEC at its public reference room at 100 F Street, N.E., Room 1580, Washington, D.C. 20549. You may also obtain copies of these documents at prescribed rates by writing to the Public Reference Section of the SEC at 100 F Street, N.E., Washington, D.C. 20549. Please call the SEC at 1-800-SEC-0330 for further information on the operation of the public reference facilities.

Upon completion of this offering, we will be subject to the information reporting requirements of the Exchange Act, and we will file reports, proxy statements and other information with the SEC. These reports, proxy statements and other information will be available for inspection and copying at the public reference room and website of the SEC referred to above. We also maintain a website at www.clearsidebio.com, at which you may access these materials free of charge as soon as reasonably practicable after they are electronically filed with, or furnished to, the SEC. The information contained in, or that can be accessed through, our website is not part of, and is not incorporated into, this prospectus.

Index to Financial Statements

Report of Independent Registered Public Accounting Firm	F-2
Balance sheets as of December 31, 2012 and 2013 and June 30, 2014 (unaudited)	F-3
Statements of operations for the years ended December 31, 2012 and 2013, for the period from May 26, 2011 (date of inception) to December 31, 2013, for the six months ended June 30, 2013 and 2014 (unaudited) and for the period from May 26, 2011 (date of inception) to June 30, 2014 (unaudited)	F-4
Statements of stockholders' deficit for the period from May 26, 2011 (date of inception) to December 31, 2013 and June 30, 2014 (unaudited)	F-5
Statements of cash flows for the years ended December 31, 2012 and 2013, for the period from May 26, 2011 (date of inception) to December 31, 2013, for the six months ended June 30, 2013 and 2014 (unaudited) and for the period from May 26, 2011 (date of inception) to June 30, 2014 (unaudited)	F-6
Notes to financial statements	F-7

Report of Independent Registered Public Accounting Firm

The Board of Directors and Stockholders of
Clearside Biomedical, Inc.

We have audited the accompanying balance sheets of Clearside Biomedical, Inc. (a development stage enterprise) (the Company) as of December 31, 2012 and 2013, and the related statements of operations, stockholders' deficit, and cash flows for each of the two years in the period ended December 31, 2013 and for the period from May 26, 2011 (date of inception) to December 31, 2013. These financial statements are the responsibility of the Company's management. Our responsibility is to express an opinion on these financial statements based on our audits.

We conducted our audits in accordance with the standards of the Public Company Accounting Oversight Board (United States). Those standards require that we plan and perform the audit to obtain reasonable assurance about whether the financial statements are free of material misstatement. We were not engaged to perform an audit of the Company's internal control over financial reporting. Our audits included consideration of internal control over financial reporting as a basis for designing audit procedures that are appropriate in the circumstances, but not for the purpose of expressing an opinion on the effectiveness of the Company's internal control over financial reporting. Accordingly, we express no such opinion. An audit also includes examining, on a test basis, evidence supporting the amounts and disclosures in the financial statements, assessing the accounting principles used and significant estimates made by management, and evaluating the overall financial statement presentation. We believe that our audits provide a reasonable basis for our opinion.

In our opinion, the financial statements referred to above present fairly, in all material respects, the financial position of Clearside Biomedical, Inc. at December 31, 2012 and 2013, and the results of its operations and its cash flows for each of the two years in the period ended December 31, 2013 and for the period from May 26, 2011 (date of inception) to December 31, 2013, in conformity with U.S. generally accepted accounting principles.

/s/ Ernst & Young LLP

Atlanta, Georgia
June 23, 2014

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)
Balance Sheets
(in thousands, except share and per share data)

	<u>December 31,</u>		<u>June 30, 2014</u> <u>(unaudited)</u>	Pro forma
	<u>2012</u>	<u>2013</u>		Liabilities, Convertible Preferred Stock and Stockholders' Deficit
				June 30, 2014 (unaudited)
Assets				
Current assets:				
Cash and cash equivalents	\$ 856	\$ 1,909	\$ 1,453	
Prepaid expenses	27	63	55	
Other current assets	—	45	20	
Total current assets	<u>883</u>	<u>2,017</u>	<u>1,528</u>	
Property and equipment, net	50	98	100	
Deferred offering costs	—	—	1,213	
Other assets	15	22	21	
Total assets	<u>\$ 948</u>	<u>\$ 2,137</u>	<u>\$ 2,862</u>	
Liabilities, convertible preferred stock and stockholders' deficit				
Current liabilities:				
Accounts payable	\$ 129	\$ 248	\$ 1,530	\$ 1,530
Accrued liabilities	400	427	1,107	1,107
Current portion of deferred rent	2	8	11	11
Current portion of long-term debt	—	—	2,778	2,928
Total current liabilities	<u>531</u>	<u>683</u>	<u>5,426</u>	<u>5,576</u>
Deferred rent	26	22	16	16
Long-term debt	150	268	150	—
Other non-current liabilities	2	31	30	30
Total liabilities	<u>709</u>	<u>1,004</u>	<u>5,622</u>	<u>5,622</u>
Convertible preferred stock:				
Series A preferred stock, \$0.001 par value, 5,200,000 shares authorized; 5,198,826 shares issued and outstanding at December 31, 2012 and 2013 and June 30, 2014 (unaudited); no shares authorized, issued or outstanding, pro forma; liquidation preference of \$4,085,705 at December 31, 2013 and June 30, 2014 (unaudited)	4,029	4,040	4,046	—
Series A-1 preferred stock, \$0.001 par value; no shares authorized, issued or outstanding at December 31, 2012; 4,800,000 shares authorized, 4,356,931 shares issued and outstanding at December 31, 2013 and June 30, 2014 (unaudited); no shares authorized, issued or outstanding, pro forma; liquidation preference of \$7,899,987 at December 31, 2013 and June 30, 2014 (unaudited)	—	7,831	7,837	—
Total convertible preferred stock	<u>4,029</u>	<u>11,871</u>	<u>11,883</u>	<u>—</u>
Stockholders' deficit:				
Common stock, \$0.001 par value; 17,000,000 shares authorized; 2,547,243 and 3,482,916 shares issued and outstanding at December 31, 2012 and 2013, respectively; and 3,950,620 shares issued and outstanding at June 30, 2014 (unaudited)	2	3	4	14
Additional paid-in capital	489	794	1,279	13,152
Deficit accumulated during the development stage	<u>(4,281)</u>	<u>(11,535)</u>	<u>(15,926)</u>	<u>(15,926)</u>
Total stockholders' deficit	<u>(3,790)</u>	<u>(10,738)</u>	<u>(14,643)</u>	<u>(2,760)</u>
Total liabilities, convertible preferred stock and stockholders' deficit	<u>\$ 948</u>	<u>\$ 2,137</u>	<u>\$ 2,862</u>	<u>\$ 2,862</u>

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)
Statements of Operations
(in thousands, except share and per share data)

	<u>Year Ended December 31,</u>		<u>Six Months Ended</u>		<u>Period From</u>	<u>Period From</u>
	<u>2012</u>	<u>2013</u>	<u>June 30,</u>		<u>May 26, 2011</u>	<u>May 26, 2011</u>
			<u>2013</u>	<u>2014</u>	<u>(Date of</u>	<u>(Date of</u>
			<u>(unaudited)</u>		<u>Inception) to</u>	<u>Inception) to</u>
					<u>December 31,</u>	<u>June 30,</u>
					<u>2013</u>	<u>2014</u>
						<u>(unaudited)</u>
Operating expenses:						
Research and development	\$ 2,354	\$ 5,045	\$ 2,204	\$ 2,562	\$ 7,606	\$ 10,168
General and administrative	1,575	2,193	1,101	1,731	3,909	5,640
Total operating expenses	<u>3,929</u>	<u>7,238</u>	<u>3,305</u>	<u>4,293</u>	<u>11,515</u>	<u>15,808</u>
Loss from operations	(3,929)	(7,238)	(3,305)	(4,293)	(11,515)	(15,808)
Other income (expense):						
Interest expense	(3)	(23)	(6)	(99)	(28)	(127)
Interest income	1	7	5	1	8	9
Total other expense	<u>(2)</u>	<u>(16)</u>	<u>(1)</u>	<u>(98)</u>	<u>(20)</u>	<u>(118)</u>
Net loss	<u>\$ (3,931)</u>	<u>\$ (7,254)</u>	<u>\$ (3,306)</u>	<u>\$ (4,391)</u>	<u>\$ (11,535)</u>	<u>\$ (15,926)</u>
Net loss per share of common stock — basic and diluted	\$ (2.12)	\$ (2.45)	\$ (1.21)	\$ (1.20)	\$ (5.50)	\$ (6.78)
Weighted average shares outstanding, basic and diluted	1,853,423	2,956,285	2,728,321	3,673,629	2,097,885	2,349,838
Pro forma net loss per share — basic and diluted (unaudited)		\$ (0.58)		\$ (0.33)		
Pro forma weighted average shares outstanding — basic and diluted (unaudited)		12,512,042		13,229,386		

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)
Statement of Stockholders' Deficit
(in thousands, except share and per share data)

	Common Stock		Additional Paid-In Capital	Deficit Accumulated During Development Stage	Total Stockholders' Deficit
	Shares	Amount			
Balance at May 26, 2011 (inception)	—	\$ —	\$ —	\$ —	\$ —
Issuance of Common Stock at par value on August 4, 2011	50,000	—	—	—	—
Vesting of restricted stock	1,315,833	1	—	—	1
Stock-based compensation expense	—	—	217	—	217
Net loss	—	—	—	(350)	(350)
Balance at December 31, 2011	1,365,833	1	217	(350)	(132)
Exercise of stock options at \$0.07 per share on August 24, 2012	77,484	—	6	—	6
Exercise of stock options at \$0.07 per share on December 18, 2012	161,426	—	11	—	11
Vesting of restricted stock	942,500	1	—	—	1
Accretion of stock issuance costs	—	—	(11)	—	(11)
Stock-based compensation expense	—	—	266	—	266
Net loss	—	—	—	(3,931)	(3,931)
Balance at December 31, 2012	2,547,243	2	489	(4,281)	(3,790)
Exercise of stock options at \$0.07 per share on October 1, 2013	41,664	—	3	—	3
Exercise of stock options at \$0.07 per share on November 26, 2013	13,853	—	1	—	1
Exercise of stock options at \$0.07 per share on December 16, 2013	18,591	—	1	—	1
Vesting of restricted stock	861,565	1	—	—	1
Accretion of stock issuance costs	—	—	(22)	—	(22)
Stock compensation expense	—	—	322	—	322
Net loss	—	—	—	(7,254)	(7,254)
Balance at December 31, 2013	3,482,916	3	794	(11,535)	(10,738)
Vesting of restricted stock (unaudited)	443,124	1	2	—	3
Exercise of stock options at \$0.07 per share on June 9, 2014	24,580	—	—	—	—
Issuance of warrants to purchase common stock (unaudited)	—	—	263	—	263
Accretion of stock issuance costs (unaudited)	—	—	(12)	—	(12)
Stock compensation expense (unaudited)	—	—	232	—	232
Net loss (unaudited)	—	—	—	(4,391)	(4,391)
Balance at June 30, 2014 (unaudited)	<u>3,950,620</u>	<u>\$ 4</u>	<u>\$ 1,279</u>	<u>\$ (15,926)</u>	<u>\$ (14,643)</u>

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)
Statement of Cash Flows
(in thousands)

	<u>Year Ended</u> <u>December 31,</u>		<u>Six Months</u> <u>Ended June 30,</u>		<u>Period From</u> <u>May 26, 2011</u> <u>(Date of</u> <u>Inception) to</u> <u>December 31,</u>	<u>Period From</u> <u>May 26, 2011</u> <u>(Date of</u> <u>Inception) to</u> <u>June 30,</u>
	<u>2012</u>	<u>2013</u>	<u>2013</u>	<u>2014</u>	<u>2013</u>	<u>2014</u>
			<u>(unaudited)</u>			<u>(unaudited)</u>
Operating activities						
Net loss	\$(3,931)	\$(7,254)	\$ (3,306)	\$ (4,391)	\$ (11,535)	\$ (15,926)
Adjustments to reconcile net loss to net cash used by operating activities:						
Depreciation	7	15	4	13	22	35
Stock-based compensation expense	266	322	136	232	805	1,037
Accretion of debt discount	—	—	—	55	—	55
Change in fair value of warrant liability	—	6	—	2	6	8
Changes in operating assets and liabilities:						
Prepaid expenses and other current assets	19	(78)	(18)	30	(105)	(75)
Other assets	(15)	2	2	(3)	(13)	(16)
Accounts payable and accrued liabilities	450	146	86	749	680	1,429
Deferred rent	28	2	(10)	(3)	30	27
Net cash used in operating activities	<u>(3,176)</u>	<u>(6,839)</u>	<u>(3,106)</u>	<u>(3,316)</u>	<u>(10,110)</u>	<u>(13,426)</u>
Investing activities						
Acquisition of property and equipment	(57)	(63)	(42)	(15)	(120)	(135)
Net cash used in investing activities	<u>(57)</u>	<u>(63)</u>	<u>(42)</u>	<u>(15)</u>	<u>(120)</u>	<u>(135)</u>
Financing activities						
Proceeds from issuance of convertible shareholder note	—	—	—	—	100	100
Proceeds from issuance of long-term debt	150	125	125	3,000	275	3,275
Principal payments made on long-term debt	—	—	—	(125)	—	(125)
Proceeds from issuance of common stock and restricted stock	—	—	—	—	8	8
Purchase of restricted stock	(4)	—	—	—	(4)	(4)
Proceeds from exercise of stock options	17	10	—	—	27	27
Proceeds from issuance of Series A Preferred Stock, net of issuance cost	3,913	—	—	—	3,913	3,913
Proceeds from issuance of Series A-1 Preferred Stock, net of issuance cost	—	7,820	7,820	—	7,820	7,820
Net cash provided by financing activities	<u>4,076</u>	<u>7,955</u>	<u>7,945</u>	<u>2,875</u>	<u>12,139</u>	<u>15,014</u>
Net increase (decrease) in cash and cash equivalents	843	1,053	4,797	(456)	1,909	1,453
Cash and cash equivalents, beginning of period	13	856	856	1,909	—	—
Cash and cash equivalents, end of period	<u>\$ 856</u>	<u>\$ 1,909</u>	<u>\$ 5,653</u>	<u>\$ 1,453</u>	<u>\$ 1,909</u>	<u>\$ 1,453</u>
Supplemental schedule of noncash investing and financing activities						
Conversion of notes payable and accrued interest into Series A preferred stock	\$ 105	\$ —	\$ —	\$ —	\$ 105	\$ 105
Issuance of warrant to purchase Series A-1 preferred stock	—	19	19	—	19	19
Issuance of warrant to purchase common stock	—	—	—	263	—	263
Accretion of redeemable convertible preferred stock to redemption value	11	22	12	12	33	45
Vesting of restricted stock	1	1	—	3	3	6
Amortization of debt discount	—	2	—	7	2	9
Deferred initial public offering costs in accounts payable and accrued expenses	—	—	—	1,213	—	1,213

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements

1. The Company

Clearside Biomedical, Inc. (the “Company”) is a clinical-stage biopharmaceutical company developing first-in-class drug therapies to treat chronic, blinding diseases of the eye. The Company’s current product candidates focus on diseases affecting the retina and choroid, especially diseases associated with macular edema, and are injected non-surgically into the suprachoroidal space using its proprietary microinjector. Incorporated in the State of Delaware on May 26, 2011, the Company has its corporate headquarters in Alpharetta, Georgia.

To date, the Company is in the development stage since it has not yet commenced primary operations or generated significant revenue as defined by Financial Accounting Standards Board (“FASB”) Accounting Standards Codification (“ASC”) Topic 915, *Development Stage Entities*. The Company’s activities since inception have primarily consisted of developing product and technology rights, raising capital and performing research and development activities. The Company has no current source of revenue to sustain present activities, and does not expect to generate revenue until and unless the Company receives regulatory approval of and successfully commercialize its product candidates. The Company is subject to a number of risks and uncertainties similar to those of other development stage life science companies, including, among others, the need to obtain adequate additional financing, successful development efforts including regulatory approval of products, compliance with government regulations, successful commercialization of potential products, protection of proprietary technology and dependence on key individuals.

Liquidity

The Company has funded its operations to date primarily through the sale of convertible preferred stock. The Company will need to obtain additional financing to fund future operations, including completing the development and commercialization of its primary product candidates. The Company will need to expend substantial resources for research and development, including costs associated with the clinical testing of its product candidates. The Company will need to obtain additional financing to conduct additional trials for the regulatory approval of its drug candidates if requested by regulatory bodies, and completing the development of any additional product candidates that might be acquired. If such products were to receive regulatory approval, the Company would need to prepare for the potential commercialization of its product candidates and fund the commercial launch of the products, if the Company decides to commercialize the products on its own. Moreover, the Company’s fixed expenses such as rent and other contractual commitments are substantial and are expected to increase in the future.

The Company has incurred losses and experienced negative operating cash flows since inception, and has cumulative net cash flows used by operating activities of \$10.1 million and \$13.4 million and cumulative net losses of \$11.5 million and \$15.9 million for the period from May 26, 2011 (inception) to December 31, 2013 and for the period from May 26, 2011 (inception) to June 30, 2014 (unaudited), respectively. The total future need for operating capital and research and development funding significantly exceeds the cash and cash equivalents that the Company has on its balance sheet. As a result, the Company will require additional funding in the future and may not be able to raise such additional funds. In the absence of product or other revenues, the amount, timing, nature or source of which cannot be predicted, the Company’s losses will continue as it conducts its research and development activities. Until the Company

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

can generate a sufficient amount of revenue, the Company may finance future cash needs through public or private equity offerings, license agreements, debt financings, collaborations, strategic alliances and marketing or distribution arrangements. The incurrence of indebtedness would result in increased fixed payment obligations and could also result in restrictive covenants, such as limitations on the Company's ability to incur additional debt, limitations on its ability to acquire, sell or license intellectual property rights and other operating restrictions that could adversely impact the ability of the Company to conduct its business. If adequate funds are not available, the Company plans to delay, reduce or eliminate research and development programs or reduce administrative expenses. The Company may seek to access the public or private capital markets whenever conditions are favorable, even if the Company does not have an immediate need for additional capital at that time. In addition, if the Company raises additional funds through collaborations, strategic alliances or marketing, distribution or licensing arrangements with third parties, the Company may have to relinquish valuable rights to its technologies, future revenue streams or product candidates or to grant licenses on terms that may not be favorable to the Company. There is no assurance that the Company will be successful in obtaining sufficient financing on acceptable terms and conditions to fund continuing operations, if at all. The failure of the Company to obtain sufficient funds on acceptable terms when needed could have a material adverse effect on the Company's business, results of operations and financial condition.

2. Significant Accounting Policies

Basis of Presentation

The Company's financial statements have been prepared in conformity with accounting principles generally accepted in the United States of America ("U.S. GAAP").

Unaudited Interim Financial Information

The accompanying balance sheet as of June 30, 2014, statements of operations and statements of cash flows for the six months ended June 30, 2013 and 2014 and the period from May 26, 2011 (inception) to June 30, 2014 and statement of stockholders' deficit as of June 30, 2014 are unaudited. The interim unaudited financial statements have been prepared on the same basis as the annual financial statements and, in the opinion of management, reflect all adjustments, which include only normal recurring adjustments, necessary for the fair presentation of the Company's financial position as of June 30, 2014 and the results of its operations and its cash flows for the six months ended June 30, 2013 and 2014 and the period from May 26, 2011 (inception) to June 30, 2014. The financial data and other information disclosed in these notes related to the six months ended June 30, 2013 and 2014 and the period from May 26, 2011 (inception) to June 30, 2014 are unaudited. The results for the six months ended June 30, 2014 are not indicative of results to be expected for the year ending December 31, 2014, any other interim periods or any future year or period.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

Unaudited Pro Forma Information

The unaudited pro forma balance sheet data as of June 30, 2014 gives effect to the automatic conversion of all outstanding shares of the Company's convertible preferred stock into an aggregate of 9,555,757 shares of common stock upon completion of the Company's planned initial public offering. The unaudited pro forma basic and diluted net loss per share for the year ended December 31, 2013 and six months ended June 30, 2014 gives effect to such automatic conversion as if it had occurred as of the beginning of the periods presented.

Use of Estimates

The preparation of financial statements in conformity with U.S. GAAP requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities and disclosure of contingent assets and liabilities at the date of the financial statements and reported amounts of income and expenses during the reporting periods. Significant items subject to such estimates and assumptions include the accounting for useful lives to calculate depreciation and amortization, accrued liabilities, share-based compensation expense and income tax valuation allowance. Actual results could differ from these estimates.

Segment Information

Operating segments are defined as components of an enterprise about which separate discrete information is available for evaluation by the chief operating decision maker, or decision-making group, in deciding how to allocate resources and in assessing performance. The Company views its operations and manages its business in one operating segment. All operations are located in the United States.

Property and Equipment, Net

Property and equipment is recorded at historical cost. Depreciation is calculated using the straight-line method over the estimated useful lives of the related assets. Repairs and maintenance are expensed when incurred. Upon retirement or sale, the cost of the assets disposed of and the related accumulated depreciation are removed from the accounts, and any resulting gain or loss is included in the determination of net income.

Cash Equivalents

Cash equivalents consist of short-term, highly liquid investments with an original term of three months or less at the date of purchase.

Concentration of Credit Risk Arising From Cash Deposits in Excess of Insured Limits

The Company maintains its cash in bank deposits that at times may exceed federally insured limits. The Company has not experienced any loss in such accounts. The Company believes it is not exposed to any significant risks with respect to its cash balances.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

Deferred Offering Costs

The Company capitalizes certain legal, accounting and other third-party fees that are directly associated with in-process equity financings as other assets until such financings are consummated. After consummation of the equity financing, these costs are recorded in stockholders' equity (deficit) as a reduction of additional paid-in capital generated as a result of the offering or as a reduction to the carrying value of preferred stock issued, if such stock is classified outside of stockholders' equity (deficit). Should the equity financing no longer be considered probable of being consummated, the deferred offering costs would be expensed immediately as a charge to operating expenses in the statement of operations. The Company did not record any deferred offering costs as of December 31, 2012 or 2013. As of June 30, 2014 (unaudited), the Company had recorded \$1.2 million of deferred offering costs.

Research and Development Costs

Research and development costs are charged to expense as incurred and include, but are not limited to:

- employee-related expenses, including salaries, benefits, travel and stock-based compensation expense for research and development personnel;
- expenses incurred under agreements with contract research organizations, contract manufacturing organizations and consultants that conduct clinical and preclinical studies;
- costs associated with preclinical and development activities;
- costs associated with technology and intellectual property licenses;
- costs for the Company's research and development facility; and
- depreciation expense for assets used in research and development activities.

Costs for certain development activities, such as clinical studies, are recognized based on an evaluation of the progress to completion of specific tasks using data such as patient enrollment, clinical site activations, or information provided to the Company by its vendors on their actual costs incurred. Payments for these activities are based on the terms of the individual arrangements, which may differ from the patterns of costs incurred, and are reflected in the financial statements as prepaid or accrued expense, which are reported in accounts payable. No material adjustments to these estimates have been recorded in these financial statements.

Stock-Based Compensation

Compensation cost related to stock-based awards granted to employees is measured at grant date, based on the estimated fair value of the award. The Company estimates the fair value of stock options using a Black-Scholes option pricing model. Compensation expense for options granted to non-employees is determined as the fair value of consideration received or the fair value of the equity instruments issued, whichever is more reliably measured. The fair value of restricted stock awards is determined based on the fair value of the Company's common stock on the date of grant. Stock-based compensation costs are expensed on a straight-line basis (net of estimated forfeitures) over the relevant vesting period. The fair value of awards granted to non-employees is re-measured each period until the related service is complete.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

All stock-based compensation costs are recorded in general and administrative or research and development costs in the statements of operations based upon the underlying employees' roles within the Company.

Debt Discount

The Company follows the authoritative guidance in ASC 470-20-25-2, *Debt with Conversion of Other Options*, for accounting for debt discount related to the detachable stock purchase warrants issued in connection with a debt obligation. The fair value of the warrants is recorded as a discount against the related debt obligation, and is amortized using the effective interest rate method over the term of the underlying debt obligation and reflected in interest expense.

Fair Value Measurements

The Company records certain financial assets and liabilities at fair value in accordance with the provisions of ASC Topic 820, *Fair Value Measurements and Disclosures*, on fair value measurements. As defined in the guidance, fair value, defined as an exit price, represents the amount that would be received to sell an asset or pay to transfer a liability in an orderly transaction between market participants. As a result, fair value is a market-based approach that should be determined based on assumptions that market participants would use in pricing an asset or a liability. As a basis for considering these assumptions, the guidance defines a three-tier value hierarchy that prioritizes the inputs used in the valuation methodologies in measuring fair value.

- Level 1—Unadjusted quoted prices in active, accessible markets for identical assets or liabilities.
- Level 2—Other inputs that are directly or indirectly observable in the marketplace.
- Level 3—Unobservable inputs that are supported by little or no market activity.

The fair value hierarchy also requires an entity to maximize the use of observable inputs and minimize the use of unobservable inputs when measuring fair value.

The Company's material financial instruments at December 31, 2012 and 2013 and June 30, 2014 (unaudited) consisted primarily of cash and cash equivalents, long-term debt and stock purchase warrant liabilities. The fair value of cash and cash equivalents, other current assets, accounts payable and accrued expenses approximate their respective carrying values due to the short term nature of these instruments. The Company has determined its stock purchase warrants liability to be a Level 3 fair value measurement (Note 10).

Stock Purchase Warrants

The Company accounts for stock purchase warrants as liabilities based upon the characteristics and provisions of the underlying instruments. These liabilities are recorded at their fair value on the date of issuance within other non-current liabilities on the balance sheet and are remeasured on each subsequent reporting date, with fair value changes recognized as income (decreases in fair value) or expenses (increases in fair value) in other income (expense), net in the statements of operations. The fair value of these liabilities is estimated using the Black-Scholes method.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

Income Taxes

Deferred tax assets or liabilities are recorded for temporary differences between financial statement and tax basis of assets and liabilities, using enacted rates in effect for the year in which the differences are expected to reverse. A valuation allowance is recorded if it is more likely than not that a deferred tax asset will not be realized. The Company has provided a full valuation allowance on its deferred tax assets, which primarily consist of cumulative net operating losses of \$10.4 million for the period from May 26, 2011 (inception) to December 31, 2013. Due to its history of operating losses since inception and losses expected to be incurred in the foreseeable future, a full valuation allowance was considered necessary.

Recent Accounting Pronouncements

Management has considered all recently issued accounting pronouncements and does not believe the adoption of such pronouncements will have a material impact on the Company's financial statements.

Net Loss per Share of Common Stock

Basic net loss per share is calculated by dividing the net loss by the weighted average number of shares of common stock outstanding for the period, without consideration of the dilutive effect of potential common stock equivalents. Diluted net loss per share gives effect to all dilutive potential shares of common stock outstanding during this period.

For all periods presented, the Company's potential common stock equivalents, which include convertible preferred stock, stock options, unvested restricted stock and stock purchase warrants, have been excluded from the computation of diluted net loss per share as their inclusion would have the effect of reducing the net loss per share. Therefore, the denominator used to calculate both basic and diluted net loss per share is the same in all periods presented.

The Company's potential common stock equivalents that have been excluded from the computation of diluted net loss per share for all periods presented because of their antidilutive effect consisted of the following:

	Year Ended December 31,		Six Months Ended June 30,		Period From May 26, 2011 (Inception) to December 31, 2013	Period From May 26, 2011 (Inception) to June 30, 2014
	2012	2013	2013	2014		
			(unaudited)			(unaudited)
Convertible preferred stock	5,198,826	9,555,757	9,555,757	9,555,757	9,555,757	9,555,757
Outstanding stock options	617,250	1,694,198	1,207,750	1,619,235	1,694,198	1,619,235
Unvested restricted stock	1,282,500	504,271	855,000	61,147	504,271	61,147
Stock purchase warrants	—	16,550	—	264,725	16,550	264,725
	<u>7,098,576</u>	<u>11,770,776</u>	<u>11,618,507</u>	<u>11,500,864</u>	<u>11,770,776</u>	<u>11,500,864</u>

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

3. Property and Equipment, Net

Property and equipment, net consisted of the following (dollar amounts in thousands):

	Estimated Useful Lives (Years)	December 31,		June 30,
		2012	2013	2014
				(unaudited)
Furniture and fixtures	5	\$ 51	\$ 77	\$ 79
Machinery and equipment	5	—	26	26
Computer equipment	3	—	7	20
	Lesser of useful life or remaining lease term			
Leasehold improvements		6	10	10
		57	120	135
Less: Accumulated depreciation		(7)	(22)	(35)
		<u>\$ 50</u>	<u>\$ 98</u>	<u>\$ 100</u>

Depreciation expense was \$7,000 and \$20,000 for the years ended December 31, 2012 and 2013, respectively; \$4,000 and \$13,000 for the six months ended June 30, 2013 (unaudited) and 2014 (unaudited), respectively; and \$22,000 and \$35,000 for the period from May 26, 2011 (inception) to December 31, 2013 and the period from May 26, 2011 (inception) to June 30, 2014 (unaudited), respectively.

4. Other Current Liabilities

Other current liabilities consisted of the following (in thousands):

	December 31,		June 30,
	2012	2013	2014
			(unaudited)
Accrued expenses	\$164	\$ 93	\$ 608
Accrued bonuses	215	289	378
Accrued vacation	21	33	74
Accrued interest payable	—	12	47
	<u>\$400</u>	<u>\$427</u>	<u>\$ 1,107</u>

5. Long-Term Debt

Note Purchase Agreement

On December 20, 2012, the Company entered into a \$150,000 unsecured note purchase agreement with a lender and borrowed \$150,000 on that date. Amounts borrowed under the agreement bore interest at 5% per annum. All unpaid principal, together with the balance of unpaid and accrued interest, were due and payable on demand at any time after the earlier of (i) the maturity date of December 2017, (ii) the date

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

on which the Company has achieved sustainable profitability for a period of at least two consecutive fiscal years in accordance with generally accepted accounting principles, (iii) without the prior written consent of the lender, the date on which an equity financing of the Company in which the Company issues shares of common stock, preferred stock or other equity interests in the Company in a transaction or series of related transactions and receives an investment of cash in consideration of such issuance in the amount of not less than \$7,000,000 or consolidation of the Company or the sale or transfer by the Company's stockholders of capital stock of the Company representing more than 50% of the voting power occurs or (iv) upon or after the occurrence of an event of default. The repayment acceleration provision specifically excluded the Series A-1 Preferred Stock financing that occurred in January 2013. The unsecured promissory note converted into an aggregate of 60,291 shares of Series B convertible preferred stock in connection with the Company's August 2014 Series B convertible preferred stock financing.

Interest expense on the borrowings under the note purchase agreement was \$0 and \$8,000 for the years ended December 31, 2012 and 2013, respectively; \$4,000 for each of the six months ended June 30, 2013 (unaudited) and 2014 (unaudited); and \$8,000 and \$12,000 for the period from May 26, 2011 (inception) to December 31, 2013 and the period from May 26, 2011 (inception) to June 30, 2014 (unaudited), respectively. As of December 31, 2013 and June 30, 2014 (unaudited), the total amount of borrowings due under this note purchase agreement was \$150,000.

Loan Agreement

In April 2013, the Company borrowed \$125,000 under a loan agreement, which borrowings bore interest at a compounded annual rate of 4.25%. As of December 31, 2013, the total amount of borrowings due under the loan agreement was \$125,000. All unpaid principal, together with the balance of unpaid and accrued interest, was due and payable on demand at any time after the earlier of (i) the maturity date of February 2016, (ii) the date on which the Company sells, leases, transfers or otherwise disposes of all or substantially all of its assets now owned or hereafter acquired, (iii) the date on which the Company makes a public offering of the Company's capital stock or equity interests, (iv) the date on which the Company takes any action which would result in a change in the direct or indirect control of 50% or more of the capital stock or equity interest ownership of the Company, (v) the date after December 31, 2013 on which the Company had received additional equity investments or milestone payments or license fees totaling \$2,500,000 in the aggregate over any 12-month period or (vi) upon or after the occurrence of an event of default. Subsequent to December 31, 2013, the Company repaid this loan in full.

Interest expense on the borrowings under the loan agreement was \$0 and \$4,000 for the years ended December 31, 2012 and 2013, respectively; \$0 and \$2,000 for the six months ended June 30, 2013 (unaudited) and 2014 (unaudited), respectively; and \$4,000 and \$6,000 for the period from May 26, 2011 (inception) to December 31, 2013 and the period from May 26, 2011 (inception) to June 30, 2014 (unaudited), respectively.

As of December 31, 2013, the Company had recorded unamortized debt discount of \$7,000, relating to the detachable warrants issued in conjunction with the loan agreement (Note 10). Debt discounts are amortized using the effective interest method through the earlier of the date of maturity or the conversion of the debt. As of December 31, 2013, cumulative amortization of debt discount amounted to \$3,000.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

6. Convertible Shareholder Notes Payable

During 2011, the Company issued a convertible promissory note (the "Note") to Daniel H. White, the Company's chief executive officer, under which \$100,000 was advanced to the Company in a series of advances between June 2011 and December 2011. The Note, which totaled \$104,707, including accrued interest of \$4,707, was converted, in two tranches, into 133,234 shares of Series A preferred stock of the Company during the year ended December 31, 2012.

In April 2014, the Company authorized the sale of convertible promissory notes (the "Bridge Notes") to its existing stockholders, including two of its executive officers and one of its directors in their individual capacities, in the aggregate principal amount of \$6.0 million. In April 2014, the Company issued \$3.0 million in aggregate principal amount of Bridge Notes. As of June 30, 2014 (unaudited), the total amount of borrowings due under the loan agreement was \$3.0 million. The outstanding notes accrued interest at a rate of 7%, with principal plus interest due upon maturity in April 2015, unless earlier converted. The Bridge Notes were convertible upon the occurrence of a qualified financing. The Company's August 2014 Series B convertible preferred stock financing was a qualified financing as contemplated by the Bridge Notes, and accordingly the principal and interest under all of the Bridge Notes was converted automatically into an aggregate of 1,137,644 shares of Series B convertible preferred stock in connection with this financing. In connection with the issuance of the Bridge Notes, the Company also issued warrants to the lenders to purchase an aggregate of 248,175 shares of common stock at an exercise price of \$0.01 per share. Unless earlier exercised, these warrants will expire upon the closing of an initial public offering.

Interest expense on the borrowings under the Bridge Notes was \$36,000 for the six months ended June 30, 2014 (unaudited) and for the period from May 26, 2011 (inception) to June 30, 2014 (unaudited).

As of June 30, 2014 (unaudited), the Company had recorded unamortized debt discount of \$0.2 million, relating to the detachable warrants issued in conjunction with the Bridge Notes. Debt discounts are amortized using the effective interest method through the earlier of the date of maturity or the conversion of the debt. As of June 30, 2014 (unaudited), cumulative amortization of debt discount amounted to \$41,000.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

7. Income Taxes

No provision for U.S. federal or state income taxes has been recorded as the Company has incurred net operating losses since inception. Significant components of the Company's net deferred income tax assets consist of the following (in thousands):

	December 31,	
	2012	2013
Current		
Deferred tax asset (liability)		
Non-deductible accrued expenses	\$ 8	\$ 125
Deferred rent	1	3
Valuation allowance	(9)	(128)
Net current deferred tax asset	<u>\$ —</u>	<u>\$ —</u>
Non-current		
Deferred tax asset (liability)		
Stock compensation expense	\$ 27	\$ 39
Net operating loss carryforwards	1,447	4,005
Depreciation differences	—	(17)
Tax credits	1	181
Deferred rent	10	8
Charitable contributions	—	2
Valuation allowance	(1,485)	(4,218)
Net non-current deferred tax asset	<u>\$ —</u>	<u>\$ —</u>

A reconciliation of the statutory tax rates and the effective tax rates is as follows:

	Year Ended December 31,	
	2012	2013
U.S. federal tax rate	34.00%	34.00%
State tax rate	4.62	4.43
Permanent Difference and Other	(0.62)	(1.60)
Tax Credit	0.03	2.47
Valuation allowance	<u>(38.03)</u>	<u>(39.30)</u>
	0.00%	0.00%

Realization of the future tax benefits is dependent on the Company's ability to generate sufficient taxable income within the carryforward period. Due to the Company's history of operating losses, the deferred tax assets arising from the aforementioned future tax benefits are currently not likely to be realized and, accordingly, are offset by a full valuation allowance. The income tax provision varies from the expected provision determined by applying the federal statutory income tax rate to net income (loss). The reasons for the difference in the expected provision, as determined by applying the federal statutory income tax rate to net income (loss) is primarily due to the increase in the deferred income tax valuation allowance of \$1.5 million and \$2.8 million for the years ended December 31, 2012 and 2013, respectively.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

As of December 31, 2013, the Company has net deferred tax assets primarily related to net operating loss carryforwards of \$4.0 million, which expire through 2033. Utilization of the net operating loss carryforwards and credits may be subject to a substantial annual limitation due to the ownership change limitations provided by the Internal Revenue Code of 1986, as amended, and similar state provisions. The effect of an ownership change could be an imposition of an annual limitation on the use of net operating loss carryforwards attributable to periods before the change.

The U.S. federal statute of limitations remains open for the periods from inception and forward. The Company has not been the subject of examination by the taxing authorities.

The Company has no uncertain tax positions.

8. Convertible Preferred Stock

The Company has authorized 10,000,000 shares of preferred stock. Of the authorized shares of preferred stock, 5,200,000 shares have been designated as Series A Convertible Preferred Stock ("Series A") and 4,800,000 shares have been designated as Series A-1 Preferred Stock ("Series A-1"). The Series A and Series A-1 shares were issued at a price of \$0.78589 and \$1.81320 per share, respectively.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

The following table summarizes the activity of convertible preferred stock (dollar amounts in thousands, except per share amounts):

	Series A Preferred Stock		Series A-1 Preferred Stock		Total Convertible Preferred Stock
	Shares	Amount	Shares	Amount	Amount
Balance at December 31, 2011	—	\$ —	—	\$ —	\$ —
Issuance of Series A at \$0.78589 per share on January 2, 2012, net of issuance cost of \$26	1,479,213	1,137	—	—	1,137
Issuance of Series A at \$0.78589 per share on January 15, 2012, net of issuance cost of \$31	420,382	300	—	—	300
Issuance of Series A at \$0.78589 per share on July 23, 2012, net of issuance cost of \$12	3,165,997	2,476	—	—	2,476
Conversion of related party note and interest payable at \$0.78589 per share on January 3, 2012	48,598	38	—	—	38
Conversion of related party note and interest payable at \$0.78589 per share on July 23, 2012	84,636	67	—	—	67
Accretion of preferred stock issuance costs	—	11	—	—	11
Balance at December 31, 2012	5,198,826	4,029	—	—	4,029
Issuance of Series A-1 at \$1.81320 per share, on January 31, 2013, net of issuance cost of \$75	—	—	4,081,177	7,325	7,325
Issuance of Series A-1 at \$1.81320 per share, on February 12, 2013, net of issuance cost of \$5	—	—	275,754	495	495
Accretion of preferred stock issuance costs	—	11	—	11	22
Balance at December 31, 2013	5,198,826	4,040	4,356,931	7,831	11,871
Accretion of preferred stock issuance costs (unaudited)	—	6	—	6	12
Balance at June 30, 2014 (unaudited)	<u>5,198,826</u>	<u>\$ 4,046</u>	<u>4,356,931</u>	<u>\$ 7,837</u>	<u>\$ 11,883</u>

Dividends

Holders of Series A and Series A-1 shares, in preference of common stockholders, shall be entitled to receive (a) when, as and if declared by the board of directors (the "Board"), but only out of funds that are legally available therefore, or (b) upon the liquidation, dissolution or winding up of the Company, whether voluntary or involuntary, cumulative cash dividends at the rate of 8% per annum of the applicable original issuance price of such series of preferred stock.

The total cumulative preferred dividends in arrears for the preferred stock as of December 31, 2013 and June 30, 2014 (unaudited) were \$1,115,000 and \$1,594,000, respectively.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

Liquidation

Upon a liquidation event (as defined in the amended and restated certificate of incorporation) the Series A and Series A-1 holders will be paid their liquidation preference of \$0.78589 and \$1.81320 per share, respectively, which is the original issue price plus any accrued and declared but unpaid dividends on such class of capital stock. If the net assets of the Company are insufficient to cover the liquidation preference, the Company will distribute the available funds among the holders of Series A and Series A-1 shares at the time outstanding, ratably in proportion to the full amounts to which they would otherwise be entitled if such amounts had been paid in full.

Conversion

Each share of Series A and Series A-1 is convertible into a number of fully paid shares of common stock at any time at the option of the stockholder. The Series A and Series A-1 shares may be converted into common stock at a conversion price of \$0.78589 and \$1.81320, respectively. As described in the amended and restated certificate of incorporation, a reduction in the conversion price will occur if the Company sells common stock for less than the conversion price of the Series A and Series A-1 shares. Based on the conversion terms, there were no beneficial conversion features associated with Series A and Series A-1 shares.

In addition, the potential reduction in the conversion price did not result in the conversion price feature meeting the definition of a derivative, which would require separate accounting.

Voting

Each holder of the Series A and Series A-1 shares shall be entitled to the number of votes equal to the number of shares of common stock into which such shares of Series A and Series A-1 may be converted, and shall have voting rights and powers equal to the voting rights and powers of the common stock, with certain limitations.

Redemption

Series A and Series A-1 preferred stock will be subject to redemption at the option of the investors holding a majority of the Series A and Series A-1 shares at any time after the sixth anniversary of the issuance in an amount equal to the liquidation preference. On such redemption date, the Company shall redeem, on a pro rata basis in accordance with the number of shares of Series A and Series A-1 owned by each holder, that number of outstanding shares of Series A and Series A-1 determined by dividing (i) the total number of shares of Series A and Series A-1 outstanding immediately prior to such redemption date by (ii) the number of remaining redemption dates including the redemption date to which such calculation applies. If the Company does not have sufficient funds legally available to redeem on any redemption date all Series A and Series A-1 shares to be redeemed on such redemption date, the Company shall redeem a pro rata portion of each holder's Series A and Series A-1 shares out of funds legally available therefor, based on the respective amounts which would otherwise be payable in respect of the shares to be redeemed if the legally available funds were sufficient to redeem all such shares, and shall redeem the remaining shares to have been redeemed as soon as practicable after the Company has funds legally available therefor.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

9. Common Stock

The Company's amended and restated certificate of incorporation authorizes the Company to issue 17,000,000 shares of \$0.001 par value common stock. As of December 31, 2013 and June 30, 2014 (unaudited), there were 3,482,916 and 3,950,620 shares of common stock outstanding, respectively, which excludes 504,271 and 61,147 shares, respectively, of unvested restricted stock. The terms, rights, preferences and privileges of the Company's common stock are as follows:

Voting

Each holder of common stock is entitled to one vote for each share of common stock held on all matters submitted to a vote of the stockholders, including the election of directors. The Company's amended and restated certificate of incorporation and bylaws do not provide for cumulative voting rights.

Dividends

Subject to preferences that may be attributable to any then outstanding convertible preferred stock, the holders of the Company's outstanding shares of common stock are entitled to receive dividends, if any, as may be declared by the Company's board of directors out of legally available funds.

Liquidation

In the event of the Company's liquidation, dissolution or winding up, holders of common stock will be entitled to share ratably in the net assets legally available for distribution to stockholders after the payment of all the Company's debts and other liabilities, subject to satisfaction of the liquidation preferences granted to the holders of any outstanding preferred stock.

Rights and Preference

Holders of the Company's common stock have no preemptive, conversion or subscription rights, and there are no redemption or other related provisions attributable to the common stock. The rights, preferences and privileges of the holders of common stock are subject to, and may be adversely affected by, the rights of the holders of shares of any series of preferred stock of the Company that may be issued.

10. Stock Purchase Warrants

During 2013, in connection with the loan agreement (Note 5), the Company issued a warrant to the lender to purchase up to 16,550 shares of Series A-1 preferred stock at a price per share of \$1.8132. The term of the warrant extends until 10 years from the grant date and the warrant is exercisable at any time during that 10-year period. This warrant was outstanding at December 31, 2013 and June 30, 2014 (unaudited) and had a weighted average remaining life of 9.12 and 8.63 years, respectively.

During 2014, in connection with the issuance of the Bridge Notes (Note 6), the Company issued warrants to the lenders to purchase up to 248,175 shares of common stock at a price per share of \$0.01. The terms of these warrants extend until one year from the grant date and the warrants are exercisable at any time during that one-year period. These warrants were outstanding at June 30, 2014 (unaudited) and had a weighted average remaining life of 0.8 years. If unexercised, these warrants will expire upon the closing of an initial public offering.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

The Company recognizes the warrants on its balance sheet as an adjustment to its long-term debt, which had no effect on the Company's cash flows for any period presented.

The Company estimated the fair value of the warrant at issuance using the Black-Scholes option-pricing model utilizing the fair value of the underlying preferred stock. The estimates in the Black-Scholes option pricing model are based, in part, on assumptions, including but not limited to the volatility of comparable public companies, the expected life of the warrant, the risk-free interest rate and the fair value of the equity underlying the warrant.

Key assumptions utilized in the fair value calculation for the warrant at issuance appear in the table below:

Expected term (years)	5.00
Volatility	100.28%
Risk-free interest rate	1.38%
Dividend yield	0.00%

11. Stock-based Compensation

In November 2011, the Company's Board adopted and approved the Clearside Biomedical, Inc. 2011 Stock Incentive Plan (the "Plan") which provides for the grant of stock-based awards to employees, directors and consultants of the Company. The Company has reserved 2,240,508 shares of common stock for issuance under the Plan. The Board shall determine price, term and vesting conditions of all stock-based awards at their grant date. Absent a public market price for the Company's common stock, the board of directors will determine the estimated fair value for the underlying common stock. Stock-based awards vest over variable periods, generally from one to five years, and expire not more than ten years after the date of grant.

The total stock-based compensation expense recognized is reflected in the statements of operations as follows (in thousands):

	<u>Year Ended</u> <u>December 31,</u>		<u>Six Months</u> <u>Ended June 30,</u>		<u>Period From</u> <u>May 26, 2011</u> <u>(Inception) to</u> <u>December 31, 2013</u>	<u>Period From</u> <u>May 26, 2011</u> <u>(Inception) to</u> <u>June 30, 2014</u>
	<u>2012</u>	<u>2013</u>	<u>2013</u>	<u>2014</u>	<u>December 31, 2013</u>	<u>June 30, 2014</u>
			(unaudited)			(unaudited)
Research and development	\$ 158	\$ 213	\$ 92	\$ 136	\$ 531	\$ 667
General and administrative	108	109	44	96	274	370
Total	<u>\$ 266</u>	<u>\$ 322</u>	<u>\$ 136</u>	<u>\$ 232</u>	<u>\$ 805</u>	<u>\$ 1,037</u>

Stock-based payments are accounted for in accordance with the provisions of ASC 718, *Compensation-Stock Compensation*. The estimated fair value of options granted is determined as of the date of grant using the Black-Scholes option pricing model. The resulting fair value is recognized ratably over the requisite service period, which is generally the vesting period of the awards. Options granted to non-employees are re-measured at each financial reporting period until required services are performed.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

The following table sets forth the weighted average assumptions utilized in the fair value calculation for the underlying common stock for the years ended December 31, 2012 and 2013 and for the six months ended June 30, 2013 (unaudited). The Company did not grant any stock options during the six months ended June 30, 2014.

	<u>Year Ended December 31,</u>		<u>Six Months</u>
	<u>2012</u>	<u>2013</u>	<u>Ended June 30,</u>
			<u>2013</u>
Expected term (years)	7.00	7.00	7.00
Expected stock price volatility	156.84%	97.02%	100.06%
Risk-free interest rate	1.07%	1.69%	1.28%
Expected dividend yield	0.00%	0.00%	0.00%

Expected term (in years): The Company utilized the guidance set forth in ASC 718 to determine the expected term of options. The Company utilized the simplified method as prescribed by ASC 718, as the Company does not have sufficient historical exercise and post-vesting termination data to provide a reasonable basis upon which to estimate the expected term of stock options granted to employees.

Risk-free interest rate: The risk-free interest rate is based on the yields of U.S. Treasury securities with maturities similar to the expected time to a possible liquidity event.

Expected dividend yield: The Company has not paid and does not anticipate paying any dividends in the foreseeable future.

Expected stock price volatility: The volatility assumption is based on the historical volatilities of the stock of several public entities that are similar to the Company, as the Company does not have sufficient historical transactions in its own shares on which to base expected volatility. The same peer group of companies was utilized both for 2012 and for 2013.

Stock Options

The Company has granted stock option awards to employees, directors and consultants. Stock-based compensation expense for options granted is reflected in the statements of operations as follows (in thousands):

	<u>Year Ended</u>		<u>Six</u>		<u>Period From</u>	<u>Period From</u>
	<u>December 31,</u>	<u>December 31,</u>	<u>Months Ended</u>		<u>(Inception) to</u>	<u>(Inception) to</u>
	<u>2012</u>	<u>2013</u>	<u>2013</u>	<u>2014</u>	<u>December 31,</u>	<u>June 30,</u>
			<u>(unaudited)</u>		<u>2013</u>	<u>2014</u>
						<u>(unaudited)</u>
Research and development	\$ 17	\$ 50	\$ 16	\$ 51	\$ 67	\$ 118
General and administrative	70	71	25	76	141	217
Total	<u>\$ 87</u>	<u>\$ 121</u>	<u>\$ 41</u>	<u>\$ 127</u>	<u>\$ 208</u>	<u>\$ 335</u>

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

The following table summarizes the activity related to stock options:

	<u>Number of Shares</u>	<u>Weighted Average Exercise Price</u>
Options outstanding at May 26, 2011, inception	—	\$ —
Granted	50,000	0.04
Options outstanding at December 31, 2011	50,000	0.04
Granted	823,660	0.07
Exercised	(238,910)	0.07
Cancelled/Forfeited	(17,500)	0.07
Options outstanding at December 31, 2012	617,250	0.07
Granted	1,435,500	0.18
Exercised	(157,444)	0.07
Cancelled/Forfeited	(201,108)	0.13
Options outstanding at December 31, 2013	1,694,198	0.16
Exercised (unaudited)	(24,580)	0.07
Cancelled/Forfeited (unaudited)	(50,383)	0.07
Options outstanding at June 30, 2014 (unaudited)	1,619,235	0.16
Options exercisable at December 31, 2013	205,136	0.07
Options exercisable at June 30, 2014 (unaudited)	371,150	0.12

The following table provides additional information about the Company's stock options that were outstanding and exercisable at December 31, 2013 (aggregate intrinsic values in thousands):

<u>Exercise Price</u>	<u>Options Outstanding</u>	<u>Weighted Average Exercise Price</u>	<u>Aggregate Intrinsic Value</u>	<u>Weighted Average Remaining Contractual Life (Years)</u>	<u>Options Exercisable</u>	<u>Weighted Average Exercise Price</u>	<u>Aggregate Intrinsic Value</u>
\$0.01	25,000			7.9	16,667		
0.07	349,698			8.2	182,226		
0.18	1,319,500			9.5	6,243		
	<u>1,694,198</u>	\$ 0.16	\$ 886		<u>205,136</u>	\$ 0.07	\$ 121

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

The following table provides additional information about the Company's stock options that were outstanding and exercisable at June 30, 2014 (unaudited) (aggregate intrinsic values in thousands):

<u>Exercise Price</u>	<u>Options Outstanding</u>	<u>Weighted Average Exercise Price</u>	<u>Aggregate Intrinsic Value</u>	<u>Weighted Average Remaining Contractual Life (Years)</u>	<u>Options Exercisable</u>	<u>Weighted Average Exercise Price</u>	<u>Aggregate Intrinsic Value</u>
\$0.01	25,000			7.4	20,825		
0.07	274,735			8.0	161,380		
0.18	1,319,500			9.1	188,900		
	<u>1,619,235</u>	\$ 0.16	\$ 2,010		<u>371,105</u>	\$ 0.12	\$ 474

As of December 31, 2013 and June 30, 2014 (unaudited), the Company had \$710,000 and \$607,000 of unrecognized compensation expense, respectively, related to unvested stock options granted under the Plan. This cost is expected to be recognized over a weighted average period of 3.3 years as of June 30, 2014 (unaudited). The weighted average remaining contractual life of all outstanding options as of December 31, 2013 and June 30, 2014 (unaudited) was 9.3 and 8.9 years, respectively.

The intrinsic value is calculated as the difference between the estimated fair market value and the exercise price per share of the stock options. The estimated fair market value per share of common stock as of December 31, 2013 and June 30, 2014 (unaudited) was \$0.66 and \$1.40, respectively.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

Restricted Stock

In 2011, the founders of the Company purchased restricted common stock. These shares are subject to repurchase rights whereby the Company has the right to repurchase the unvested shares at price per share equal to the lesser of (i) the fair market value of the shares at the time the right of repurchase is exercised and (ii) the original issuance price of the shares. A summary of the status of unvested restricted stock is presented below:

	<u>Shares</u>	<u>Weighted Average Grant Date Fair Value</u>
Unvested at May 26, 2011 (inception)	—	\$ —
Granted	3,970,000	0.16
Vested	(1,315,833)	0.16
Cancelled	(50,000)	0.16
Unvested at December 31, 2011	2,604,167	0.16
Vested	(942,500)	0.16
Forfeited	(379,167)	0.16
Unvested at December 31, 2012	1,282,500	0.16
Granted	83,336	0.61
Vested	(861,565)	0.16
Unvested at December 31, 2013	504,271	0.23
Vested (unaudited)	(443,124)	0.18
Unvested at June 30, 2014 (unaudited)	61,147	0.61

Stock-based compensation expense for restricted stock granted is reflected in the statements of operations as follows (in thousands):

	<u>Year Ended December 31,</u>		<u>Six Months Ended June 30,</u>		<u>Period From May 26, 2011 (Inception) to December 31, 2013</u>	<u>Period From May 26, 2011 (Inception) to June 30, 2014 (unaudited)</u>
	<u>2012</u>	<u>2013</u>	<u>2013</u>	<u>2014</u>		
Research and development	\$ 141	\$ 163	\$ 76	\$ 86	\$ 464	\$ 550
General and administrative	38	38	19	19	133	152
Total	<u>\$ 179</u>	<u>\$ 201</u>	<u>\$ 95</u>	<u>\$ 105</u>	<u>\$ 597</u>	<u>\$ 702</u>

As of December 31, 2013 and June 30, 2014 (unaudited), the Company had \$106,000 and \$0 of unrecognized compensation expense, respectively, related to unvested restricted stock.

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

12. Commitments and Contingencies

Lease Commitment Summary

The Company leases office space under non-cancelable operating leases which expire in March 2017. The operating leases have renewal options and rent escalation clauses. The following table presents future minimum commitments of the Company due under non-cancelable operating leases with original or remaining terms in excess of one year at December 31, 2013.

Minimum lease payments were as follows at December 31, 2013 (in thousands):

2014	\$254
2015	180
2016	90
2017	23
Total minimum lease payments	<u>\$547</u>

Rent expense, net of sublease income, was \$102,000 and \$182,000 for years ended December 31, 2012 and 2013, respectively; \$72,000 and \$113,000 for the six months ended June 30, 2013 (unaudited) and 2014 (unaudited), respectively; and \$284,000 and \$397,000 for the period from May 26, 2011 (inception) to December 31, 2013 and the period from May 26, 2011 (inception) to June 30, 2014 (unaudited), respectively. Total future rent income from the sub-lease agreement on an operating lease due through May 2015 was \$59,000 as of December 31, 2013.

Litigation

The Company is not party to any litigation and does not have contingency reserves established for any litigation liabilities.

Contract Service Providers

In the course of the Company's normal business operations, it has agreements with contract service providers to assist in the performance of its research and development, clinical research and manufacturing. Substantially all of these contracts are on an as needed basis.

Employment Contracts

The Company has at will employment contracts with substantially all employees providing for salary, benefits and bonuses.

13. License Agreement

On July 4, 2012, the Company entered into an Exclusive License Agreement with Emory University and Georgia Tech Research Corporation ("Emory/GTRC"), whereby the Company purchased a license for Methods and Devices For Drug Delivery Using Microneedles. The Company paid \$30,000 for the license and made a milestone payment of \$35,000 during the year ended December 31, 2012. No payments were made to Emory/GTRC during the year ended December 31, 2013 or the six months ended June 30, 2014. The Exclusive License Agreement requires the Company to make a milestone payment upon the occurrence

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

of a specified event relating to the commercialization of a drug developed using the licensed patents. Beginning on the third anniversary of this license agreement, the Company will be obligated to pay an annual license maintenance fee of \$25,000, which will continue until the first commercial sale of a product covered by the licensed patents. Additionally, the Company will owe a low single-digit royalty on any worldwide net product sales related to the licensed patents, with minimum annual royalties of \$15,000 after commercialization. The minimum annual royalty increases thereafter to \$100,000.

14. Collaborative Agreement

On January 31, 2013, the Company entered into a collaborative research agreement with one of the holders of the Series A-1 preferred stock, whereby the two parties agreed to conduct feasibility studies for certain compounds. Each party to the collaborative research agreement will bear its own costs, except that certain costs incurred by the Company are limited to a defined maximum amount. The Company incurred research and development costs in relation to the collaborative research agreement of \$162,000 in 2013 and \$51,000 for the six months ended June 30, 2014 (unaudited).

15. Unaudited Pro Forma Net Loss Per Share

The unaudited pro forma basic and diluted net loss per share for the year ended December 31, 2013 and six months ended June 30, 2014 gives effect to the conversion of outstanding convertible preferred stock, which will occur automatically upon the completion of the planned initial public offering, as if such conversion occurred as of the beginning of the periods presented. The following table sets forth the computation of unaudited pro forma basic and diluted net loss per share of common stock for the periods indicated (in thousands except share and per share amounts):

	Year Ended December 31, 2013	Six Months Ended June 30, 2014
Numerator for pro forma calculation:		
Net loss	\$ (7,254)	\$ (4,391)
Denominator for pro forma calculation:		
Weighted-average number of shares outstanding—basic and diluted	2,956,285	3,673,629
Pro forma adjustment to reflect automatic conversion of outstanding convertible preferred stock	<u>9,555,757</u>	<u>9,555,757</u>
Weighted-average number of pro forma shares outstanding—basic Diluted	12,512,042	13,229,386
Pro forma net loss per share—basic and diluted	\$ (0.58)	\$ (0.33)

CLEARSIDE BIOMEDICAL, INC.
(A Development Stage Enterprise)

Notes to the Financial Statements (Continued)

16. Subsequent Events (Unaudited)

The Company evaluated subsequent events through September 3, 2014, the date on which these financial statements were issued. In August 2014, the Company issued an aggregate of 6,009,202 shares of Series B convertible preferred stock to 31 accredited investors at a per share price of \$2.69783, for aggregate consideration of approximately \$16.2 million. In some cases, some or all of the purchase price for these shares took the form of conversion of principal and interest under outstanding Bridge Notes held by the respective investors (Note 6). The shares of Series B convertible preferred stock are convertible into common stock immediately upon: (i) the closing of an initial public offering generating net proceeds of not less than \$50.0 million to the Company, at a price per share of at least \$4.04675 if the initial public offering closes on or before February 20, 2015 or \$8.09349 if the initial public offering closes after February 20, 2015; or (ii) the written consent of the holders of at least a majority of the outstanding shares of preferred stock and the holders of at least 70% of the outstanding shares of Series B convertible preferred stock. In connection with the Series B convertible preferred stock financing, the Company also issued warrants to purchase an aggregate of 1,716,914 shares of common stock at an exercise price of \$0.01 per share, which will become exercisable in February 2015 if the Company has not completed an initial public offering by then, but otherwise will terminate upon the closing of the Company's initial public offering.

In connection with the Company's Series B financing, in August 2014, the Company entered into a license agreement with NovaMedica LLC ("NovaMedica"). Under this agreement, the Company granted to NovaMedica the exclusive right in Russia and specified adjacent territories to use the Company's intellectual property to develop and commercialize products involving the use of TA as the sole active pharmaceutical ingredient for administration in the SCS. In connection with this license, NovaMedica will make an upfront payment to the Company of \$200,000. In addition, NovaMedica has agreed to make royalty payments equal to the amount of any royalties the Company owes to Emory or GTRC pursuant to the Emory/GTRC License Agreement as a result of NovaMedica's sales of products and services covered by the license granted to NovaMedica, up to a maximum of a low single-digit percentage of net sales of such products or services by NovaMedica. NovaMedica has also agreed to make milestone payments, totaling \$12.7 million in the aggregate, upon the achievement of specified development and commercial milestones. The term of the NovaMedica license agreement will expire upon the expiration of the last-to-expire valid claim within the patents licensed to NovaMedica pursuant to the agreement. Either the Company or NovaMedica may terminate the agreement upon written notice in the event of the other party's material breach that is not cured within 20 business days of receiving written notice requesting a cure of a payment breach, and within 60 business days of receiving written notice requesting a cure in the case of all other breaches. Each party also has the right to terminate in the event of the other party's bankruptcy or insolvency. The Company may terminate the agreement immediately upon written notice in the event that NovaMedica commences or participates in any action challenging the validity or enforceability of the patents licensed to NovaMedica under the agreement.

Shares



COMMON STOCK

RBC CAPITAL MARKETS

WELLS FARGO SECURITIES

NEEDHAM & COMPANY

OPPENHEIMER & CO.

, 2014

PART II
INFORMATION NOT REQUIRED IN PROSPECTUS

Item 13. Other Expenses of Issuance and Distribution.

The following table sets forth all costs and expenses, other than underwriting discounts and commissions, payable by us in connection with the sale of the common stock being registered. All amounts shown are estimates except for the SEC registration fee, the Financial Industry Regulatory Authority, or FINRA, filing fee and The NASDAQ Global Market initial listing fee.

	<u>Amount to be Paid</u>
SEC registration fee	\$ *
FINRA filing fee	*
NASDAQ Global Market initial listing fee	*
Blue sky fees and expenses	*
Printing and engraving expenses	*
Legal fees and expenses	*
Accounting fees and expenses	*
Transfer agent and registrar fees and expenses	*
Miscellaneous fees and expenses	*
Total	<u>\$ *</u>

* To be filed by amendment.

Item 14. Indemnification of Directors and Officers.

We are incorporated under the laws of the State of Delaware. Section 102 of the Delaware General Corporation Law permits a corporation to eliminate the personal liability of directors of a corporation to the corporation or its stockholders for monetary damages for a breach of fiduciary duty as a director, except where the director breached his duty of loyalty, failed to act in good faith, engaged in intentional misconduct or knowingly violated a law, authorized the payment of a dividend or approved a stock repurchase in violation of Delaware corporate law or obtained an improper personal benefit.

Section 145 of the Delaware General Corporation Law provides that a corporation has the power to indemnify a director, officer, employee or agent of the corporation and certain other persons serving at the request of the corporation in related capacities against expenses (including attorneys' fees), judgments, fines and amounts paid in settlements actually and reasonably incurred by the person in connection with an action, suit or proceeding to which he is or is threatened to be made a party by reason of such position, if such person acted in good faith and in a manner he reasonably believed to be in or not opposed to the best interests of the corporation, and, in any criminal action or proceeding, had no reasonable cause to believe his conduct was unlawful, except that, in the case of actions brought by or in the right of the corporation, no indemnification shall be made with respect to any claim, issue or matter as to which such person shall have been adjudged to be liable to the corporation unless and only to the extent that the Court of Chancery or other adjudicating court determines that, despite the adjudication of liability but in view of all of the circumstances of the case, such person is fairly and reasonably entitled to indemnity for such expenses which the Court of Chancery or such other court shall deem proper.

[Table of Contents](#)

As permitted by the Delaware General Corporation Law, our amended and restated certificate of incorporation and bylaws to be in effect upon the closing of this offering will provide that: (i) we are required to indemnify our directors to the fullest extent permitted by the Delaware General Corporation Law; (ii) we may, in our discretion, indemnify our officers, employees and agents as set forth in the Delaware General Corporation Law; (iii) we are required, upon satisfaction of certain conditions, to advance all expenses incurred by our directors in connection with certain legal proceedings; (iv) the rights conferred in the bylaws are not exclusive; and (v) we are authorized to enter into indemnification agreements with our directors, officers, employees and agents.

We have entered into agreements with our directors that require us to indemnify them against expenses, judgments, fines, settlements and other amounts that any such person becomes legally obligated to pay (including with respect to a derivative action) in connection with any proceeding, whether actual or threatened, to which such person may be made a party by reason of the fact that such person is or was a director or officer of us or any of our affiliates, provided such person acted in good faith and in a manner such person reasonably believed to be in, or not opposed to, our best interests. The indemnification agreements also set forth certain procedures that will apply in the event of a claim for indemnification thereunder. We intend to enter into similar indemnification agreements with our executive officers prior to the completion of this offering. At present, no litigation or proceeding is pending that involves any of our directors or officers regarding which indemnification is sought, nor are we aware of any threatened litigation that may result in claims for indemnification.

We maintain a directors' and officers' liability insurance policy. The policy insures directors and officers against unindemnified losses arising from certain wrongful acts in their capacities as directors and officers and reimburses us for those losses for which we have lawfully indemnified the directors and officers. The policy contains various exclusions.

In addition, the underwriting agreement filed as Exhibit 1.1 to this Registration Statement provides for indemnification by the underwriters of us and our officers and directors for certain liabilities arising under the Securities Act, or otherwise.

Item 15. *Recent Sales of Unregistered Securities.*

Issuances of Capital Stock, Promissory Notes and Warrants

The following list sets forth information regarding all unregistered securities sold by us since May 26, 2011, the date of our inception, through September 8, 2014.

- 1) Between June 2011 and December 2011, we borrowed an aggregate of \$100,000 from Daniel H. White pursuant to a series of convertible promissory notes.
- 2) In January 2012, February 2012 and July 2012, we issued an aggregate of 5,198,826 shares of our Series A convertible preferred stock to seven accredited investors at a per share price of \$0.78589, for aggregate consideration of approximately \$4.1 million, including the conversion of the promissory notes described above.
- 3) In December 2012, we borrowed \$150,000 from a lender pursuant to an unsecured promissory note.
- 4) In January 2013, we issued an aggregate of 4,356,931 shares of our Series A-1 convertible preferred stock to 13 accredited investors at a per share price of \$1.8132, for aggregate consideration of approximately \$7.9 million.

Table of Contents

- 5) In February 2013, in connection with a loan agreement, we borrowed \$125,000 from a lender pursuant to a promissory note and issued a warrant to purchase 16,550 shares of our Series A-1 convertible preferred stock, which will become a warrant to purchase 16,550 shares of our common stock following the completion of this offering.
- 6) In April and May 2014, we issued an aggregate principal amount of \$3.0 million of unsecured 7% convertible promissory notes and warrants to purchase 248,175 shares of our common stock at an exercise price of \$0.01 per share to 10 accredited investors.
- 7) In August 2014, we issued an aggregate of 6,009,202 shares of our Series B convertible preferred stock at a per share price of \$2.69783 and warrants to purchase 1,716,914 shares of our common stock at an exercise price of \$0.01 per share to 31 accredited investors, for aggregate consideration of approximately \$16.2 million. In some cases, some or all of the purchase price for these shares took the form of conversion of principal and interest under outstanding convertible promissory notes held by the respective investors.

The offers, sales and issuances of the securities described in the paragraphs above were exempt from registration under Section 4(a)(2) of the Securities Act and Regulation D promulgated under the Securities Act. The recipients represented to us that they acquired the securities for investment only and not with a view to or for sale in connection with any distribution thereof and appropriate legends were affixed to the securities issued in these transactions. The recipients also represented to us that they were accredited investors as defined in Rule 501 promulgated under the Securities Act.

Stock Option Grants

From May 26, 2011, the date of our inception, through September 8, 2014, we have granted options under our 2011 stock incentive plan to purchase an aggregate of 2,689,160 shares of our common stock to employees, consultants and directors, having exercise prices ranging from \$0.01 to \$1.40 per share. Of these, options to purchase an aggregate of 578,991 shares have been cancelled without being exercised and 424,267 shares were issued upon the exercise of stock options, at an exercise price of \$0.07 per share, for aggregate proceeds of approximately \$30,000.

The offers, sales and issuances of the securities described in the foregoing paragraph were exempt from registration under Rule 701 promulgated under the Securities Act in that the transactions were under compensatory benefit plans and contracts relating to compensation as provided under Rule 701. The recipients of such securities were our employees, directors or consultants and received the securities under our 2011 stock incentive plan. Appropriate legends were affixed to the securities issued in these transactions.

Item 16. Exhibits and Financial Statement Schedules.

The exhibits to the registration statement are listed in the Exhibit Index attached hereto and are incorporated by reference herein.

Item 17. Undertakings.

The undersigned Registrant hereby undertakes to provide to the underwriters at the closing specified in the underwriting agreement, certificates in such denominations and registered in such names as required by the underwriters to permit prompt delivery to each purchaser.

[Table of Contents](#)

Insofar as indemnification for liabilities arising under the Securities Act may be permitted to directors, officers and controlling persons of the Registrant pursuant to the foregoing provisions, or otherwise, the Registrant has been advised that in the opinion of the Securities and Exchange Commission such indemnification is against public policy as expressed in the Securities Act and is, therefore, unenforceable. In the event that a claim for indemnification against such liabilities (other than the payment by the Registrant of expenses incurred or paid by a director, officer or controlling person of the Registrant in the successful defense of any action, suit or proceeding) is asserted by such director, officer or controlling person in connection with the securities being registered, the Registrant will, unless in the opinion of its counsel the matter has been settled by controlling precedent, submit to a court of appropriate jurisdiction the question whether such indemnification by it is against public policy as expressed in the Securities Act and will be governed by the final adjudication of such issue.

The undersigned Registrant hereby undertakes that:

(1) For purposes of determining any liability under the Securities Act, the information omitted from the form of prospectus filed as part of this Registration Statement in reliance upon Rule 430A and contained in a form of prospectus filed by the Registrant pursuant to Rule 424(b)(1) or (4) or 497(h) under the Securities Act shall be deemed to be part of this Registration Statement as of the time it was declared effective.

(2) For the purpose of determining any liability under the Securities Act, each post-effective amendment that contains a form of prospectus shall be deemed to be a new registration statement relating to the securities offered therein, and the offering of such securities at that time shall be deemed to be the initial bona fide offering thereof.

SIGNATURES

Pursuant to the requirements of the Securities Act, the Registrant has duly caused this Registration Statement to be signed on its behalf by the undersigned, thereunto duly authorized, in the City of Alpharetta, State of Georgia, on the _____ day of _____, 2014.

CLEARSIDE BIOMEDICAL, INC.

By: _____
Daniel H. White
President and Chief Executive Officer

KNOW ALL BY THESE PRESENTS, that each person whose signature appears below hereby constitutes and appoints Daniel H. White, Charles A. Deignan and Brent B. Siler, and each of them, his or her true and lawful agent, proxy and attorney-in-fact, with full power of substitution and resubstitution, for him or her and in his or her name, place and stead, in any and all capacities, to (i) act on, sign and file with the Securities and Exchange Commission any and all amendments (including post-effective amendments) to this registration statement together with all schedules and exhibits thereto and any subsequent registration statement filed pursuant to Rule 462(b) under the Securities Act of 1933, as amended, together with all schedules and exhibits thereto, (ii) act on, sign and file such certificates, instruments, agreements and other documents as may be necessary or appropriate in connection therewith, (iii) act on and file any supplement to any prospectus included in this registration statement or any such amendment or any subsequent registration statement filed pursuant to Rule 462(b) under the Securities Act of 1933, as amended, and (iv) take any and all actions which may be necessary or appropriate to be done, as fully for all intents and purposes as he or she might or could do in person, hereby approving, ratifying and confirming all that such agent, proxy and attorney-in-fact or any of his substitutes may lawfully do or cause to be done by virtue thereof.

Pursuant to the requirements of the Securities Act, this Registration Statement has been signed by the following persons in the capacities and on the dates indicated.

<u>Signature</u>	<u>Title</u>	<u>Date</u>
_____ Daniel H. White	President, Chief Executive Officer and Director (<i>Principal Executive Officer</i>)	, 2014
_____ Charles A. Deignan	Chief Financial Officer (<i>Principal Financial Officer and Principal Accounting Officer</i>)	, 2014
_____ Christy L. Shaffer, Ph.D.	Director	, 2014
_____ Clay B. Thorp	Director	, 2014

[Table of Contents](#)

<u>Signature</u>	<u>Title</u>	<u>Date</u>
_____ William D. Humphries	Director	, 2014
_____ Evgeny Zaytsev, M.D.	Director	, 2014
_____ Gerald D. Cagle, Ph.D.	Director	, 2014

EXHIBIT INDEX

Exhibit Number	Description of Document
1.1 †	Form of Underwriting Agreement.
3.1	Fourth Amended and Restated Certificate of Incorporation, as currently in effect.
3.2 †	Form of Certificate of Amendment of Fourth Amended and Restated Certificate of Incorporation to be filed prior to the completion of this offering.
3.3 †	Form of Fifth Amended and Restated Certificate of Incorporation to be effective upon completion of this offering.
3.4 **	Bylaws, as currently in effect.
3.5 †	Form of Amended and Restated Bylaws to be effective upon completion of this offering.
4.1 †	Specimen stock certificate evidencing shares of Common Stock.
4.2	Second Amended and Restated Investor Rights Agreement, dated as of August 29, 2014, by and among the Registrant and certain of its stockholders.
4.3 **	Form of Common Stock Purchase Warrant issued in bridge financing.
4.4 **	Stock Warrant issued to North Carolina Biotechnology Center, dated as of February 12, 2013.
5.1 †	Opinion of Cooley LLP as to legality.
10.1 #**	License Agreement, by and among the Registrant, Emory University and The Georgia Tech Research Corporation, dated as of July 4, 2012, as amended by the First Amendment to License Agreement, dated April 2, 2014.
10.2	Lease Agreement, dated as of March 14, 2012, by and between the Registrant and McDonald Ventures XI, LLC, as amended by the renewal letter from McDonald Ventures XI, LLC to the Registrant, dated March 18, 2014, and by the First Amendment to the Lease Agreement, dated August 22, 2014.
10.3 +	2011 Stock Incentive Plan, as amended to date.
10.4 +**	Form of Incentive Stock Option Grant Notice and Incentive Stock Option Agreement under 2011 Stock Incentive Plan.
10.5 +**	Form of Nonqualified Stock Option Grant Notice and Nonqualified Stock Option Agreement under 2011 Stock Incentive Plan.
10.6 +†	Form of 2014 Equity Incentive Plan
10.7 +†	Form of Stock Option Grant Notice and Stock Option Agreement under 2014 Equity Incentive Plan.
10.8 +†	Form of Restricted Stock Unit Grant Notice and Restricted Stock Unit Award Agreement under 2014 Equity Incentive Plan.
10.9 +**	Form of Indemnification Agreement with non-employee directors.

Table of Contents

<u>Exhibit Number</u>	<u>Description of Document</u>
10.10**	Office Lease, dated as of June 17, 2013, by and between the Registrant and Highwoods Realty Limited Partnership.
10.11#**	Collaboration Agreement, dated as of January 31, 2013, by and among the Registrant and Santen Pharmaceutical Co., Ltd., as amended by Amendment No. 1 to Collaboration Agreement, dated as of April 29, 2014.
10.12+†	Form of 2014 Employee Stock Purchase Plan.
10.13+†	Form of Employment Agreement with executive officers to be in effect upon completion of this offering.
10.14+†	Non-Employee Director Compensation Policy to be in effect upon completion of this offering.
10.15#†	License Agreement, by and between the Registrant and NovaMedica LLC, dated as of August 29, 2014.
23.1†	Consent of Ernst & Young LLP, independent registered public accounting firm.
23.2†	Consent of Cooley LLP (included in Exhibit 5.1).

† To be filed by amendment.

+ Indicates management contract or compensatory plan.

** Previously submitted.

Portions of this exhibit (indicated by asterisks) have been omitted pursuant to a request for confidential treatment and have been separately filed with the Securities and Exchange Commission.

FOURTH AMENDED AND RESTATED CERTIFICATE OF INCORPORATION
OF
CLEARSIDE BIOMEDICAL, INC.

Pursuant to Sections 228, 242 and 245 of the General Corporation Law of the State of Delaware, the undersigned corporation hereby submits the following for the purpose of amending and restating its Third Amended and Restated Certificate of Incorporation, and does hereby certify as follows.

1. The name of the corporation is Clearside Biomedical, Inc. The corporation's original Certificate of Incorporation was filed on May 26, 2011.
2. The corporation's Third Amended and Restated Certificate of Incorporation was filed on April 28, 2014.
3. The corporation's Third Amended and Restated Certificate of Incorporation is hereby amended and restated in its entirety, as set forth in the text of the Fourth Amended and Restated Certificate of Incorporation attached hereto as Exhibit A.
4. This Fourth Amended and Restated Certificate of Incorporation will be effective upon filing.

[Signature page to follow.]

IN WITNESS WHEREOF, said Clearside Biomedical, Inc. has caused this Fourth Amended and Restated Certificate of Incorporation to be signed by its Chief Executive Officer this 27th day of August, 2014.

CLEARSIDE BIOMEDICAL, INC.

By: /s/ Daniel White
Name: Daniel White
Title: Chief Executive Officer

EXHIBIT A

FOURTH AMENDED AND RESTATED CERTIFICATE OF INCORPORATION
OF
CLEARSIDE BIOMEDICAL, INC.

ARTICLE I

The name of the corporation shall be "Clearside Biomedical, Inc." (the "**Corporation**").

ARTICLE II

The address of the registered office of the Corporation in the State of Delaware is 2711 Centerville Road, Suite 400, Wilmington, New Castle County, Delaware 19808, and the name of the registered agent is Corporation Service Company.

ARTICLE III

The purpose for which the Corporation is organized is to engage in any lawful act or activity for which corporations may be organized under the General Corporation Law of Delaware.

ARTICLE IV

The Corporation shall have the authority to issue 46,985,672 shares of capital stock, \$0.001 par value per share, of which 30,000,000 shares shall be designated Common Stock (the "**Common Stock**") and 16,985,672 shares shall be designated Preferred Stock (the "**Preferred Stock**"). Of the authorized shares of Preferred Stock, 5,198,826 shares shall be designated Series A Preferred Stock (the "**Series A Preferred Stock**"), 4,373,481 shares shall be designated Series A-1 Preferred Stock (the "**Series A-1 Preferred Stock**") and together with the Series A Preferred Stock, the "**Series A/A-1 Preferred Stock**") and 7,413,365 shares shall be designated Series B Preferred Stock (the "**Series B Preferred Stock**"). The Preferred Stock shall have the rights, preferences, privileges and restrictions set forth below in Article V.

ARTICLE V

The rights, preferences, privileges, restrictions and other matters relating to the Preferred Stock are as follows.

A. Dividends.

1. Preferred Stock. From and after the date of the issuance of any shares of Preferred Stock, dividends at the rate per annum of: \$0.06287 per share shall accrue on such shares of Series A Preferred Stock; \$0.14506 per share shall accrue on such shares of Series A-1 Preferred Stock; and \$0.215826 per share shall accrue on such shares of Series B Preferred Stock (in each case, subject to appropriate adjustment in the event of any stock dividend, stock split, combination or other similar recapitalization with respect to the Preferred Stock) (the "**Accruing Dividends**"). Accruing Dividends shall accrue from day to day, whether or not declared; provided however, that except as set forth in the following sentence of this Subsection A.1 or in Subsections B.1 or B.2, such Accruing Dividends shall be payable only when, as, and if declared by the Board of Directors and the Corporation shall be under no

obligation to pay such Accruing Dividends. The Corporation shall, upon the written request of any holder of Preferred Stock, furnish or cause to be furnished to such holder a certificate setting forth the accrued dividends with respect to that holder's Preferred Stock and the basis for calculating the accrued dividends.

2. Series B Preferred Stock Dividend. The Corporation shall not declare, pay or set aside any dividends on shares of any other class or series of capital stock of the Corporation (other than dividends on shares of Common Stock payable in shares of Common Stock) unless (in addition to the obtaining of any consents required elsewhere in this Fourth Amended and Restated Certificate of Incorporation (this "**Restated Certificate**")) the holders of the Series B Preferred Stock then outstanding shall first receive, or simultaneously receive, a dividend on each outstanding share of Series B Preferred Stock in an amount equal to the amount of the aggregate Accruing Dividends then accrued on such share of Series B Preferred Stock and not previously paid.

3. Series A/A-1 Preferred Stock Dividend. The Corporation shall not declare, pay or set aside any dividends on shares of any other class or series of capital stock of the Corporation (other than dividends permitted by Section A(2) above or dividends on shares of Common Stock payable in shares of Common Stock) unless (in addition to the obtaining of any consents required elsewhere in this Restated Certificate) the holders of Series A Preferred Stock and Series A-1 Preferred Stock then outstanding shall first receive, or simultaneously receive, a dividend on each outstanding share of Series A Preferred Stock or Series A-1 Preferred Stock in an amount equal to the amount of the aggregate Accruing Dividends then accrued on such share of Series A Preferred Stock or Series A-1 Preferred Stock, as applicable, and not previously paid. Payment of any dividends to the holders of Series A Preferred Stock and Series A-1 Preferred Stock shall be on a *pro rata, pari passu* basis in proportion to the Accrued Dividends for each series of Preferred Stock.

4. Additional Dividends. After the payment or setting aside for payment of the dividends described in Sections A(1) and A(2), any additional dividends (other than dividends on Common Stock payable solely in Common Stock) set aside or paid in any fiscal year shall be set aside or paid among the holders of the Preferred Stock and Common Stock then outstanding in proportion to the greatest whole number of shares of Common Stock which would be held by each such holder if all shares of Preferred Stock were converted at the then-effective Conversion Rate (as defined in Section D).

B. Preference on Liquidation.

1. Upon the occurrence of any Liquidating Event (as defined below), before any payment shall be made in respect of the Corporation's Common Stock, the Series A/A-1 Preferred Stock or any other class or series of the Corporation's capital stock, each holder of Series B Preferred Stock then outstanding shall be entitled to receive, out of the assets of the Corporation available for distribution to its stockholders an amount per share of Series B Preferred Stock equal to the product of (i) \$2.69783, subject to equitable adjustment for any stock splits, combinations, consolidations, recapitalizations, reorganizations, reclassifications, stock distributions, stock dividends or other similar events (collectively, "**Recapitalizations**") with respect to such share (as so adjusted from time to time, the "**Series B Original Price**"), multiplied by (ii) 1.50, plus all declared but unpaid Accruing Dividends plus (without duplication) any other declared but unpaid dividends on such share (such amounts, the "**Series B Preference Amount**").

If, upon the occurrence of a Liquidating Event, the assets and funds distributed among the holders of Series B Preferred Stock shall be insufficient to permit the payment to such holders of the full aforesaid Series B Preference Amount then the entire assets and funds of the Corporation legally available for distribution shall be distributed ratably among the holders of Series B Preferred Stock in proportion to the aggregate of the Series B Preference Amount each such holder is entitled to receive.

2. Upon the occurrence of any Liquidating Event, after payment of the Series B Preference Amount and before any payment shall be made in respect of the Corporation's Common Stock, each holder of Series A/A-1 Preferred Stock then outstanding shall be entitled to receive, out of the assets of the Corporation available for distribution to its stockholders:

(a) an amount per share of Series A Preferred Stock equal to \$0.78589, subject to equitable adjustment for any Recapitalizations with respect to such share (as so adjusted from time to time, the "**Series A Original Price**"), plus all declared but unpaid Accruing Dividends plus (without duplication) any other declared but unpaid dividends on such share;

(b) an amount per share of Series A-1 Preferred Stock equal to \$1.8132, subject to equitable adjustment for any Recapitalizations with respect to such share (as so adjusted from time to time, the "**Series A-1 Original Price**"), plus all declared but unpaid Accruing Dividends plus (without duplication) any other declared but unpaid dividends on such share (together with the amounts set forth in subsection (a), the "**Series A/A-1 Preference Amount**" and, together with the Series B Preference Amount, the "**Preference Amount**");

If, upon the occurrence of a Liquidating Event, and after the payment in full of the Series B Preference Amount, the remaining assets and funds available for distribution among the holders of Series A/A-1 Preferred Stock shall be insufficient to permit the payment to such holders of the full aforesaid Series A/A-1 Preference Amount then such remaining funds shall be distributed ratably among the holders of Series A/A-1 Preferred Stock in proportion to the aggregate of the Series A/A-1 Preference Amount each such holder is entitled to receive.

3. After payment has been made to the holders of Preferred Stock of the full Preference Amount to which they shall be entitled as aforesaid, the remaining assets of the Corporation legally available for distribution, if any, shall be distributed ratably among the holders of the Corporation's Common Stock and Preferred Stock as if such shares of Preferred Stock had been converted voluntarily into Common Stock immediately prior to such Liquidating Event at the then-applicable conversion rate.

4. A "**Liquidating Event**" shall mean (a) any liquidation, dissolution or winding up of the Corporation, either voluntary or involuntary, or (b) a transaction or series of related transactions resulting in any of the following: (1) a sale, lease, transfer, exclusive license, exchange or other disposition of all or substantially all the assets of the Corporation, (2) a merger, consolidation, sale or reorganization as a result of which stockholders of the Corporation immediately prior to such merger, consolidation, sale or reorganization possess less than 50% of the voting power of the acquiring, surviving or successor entity immediately following such merger, consolidation, sale or reorganization, or (3) the transfer by one or more stockholders of the Corporation of securities of the Corporation representing 50% or more of the combined voting power of the then outstanding securities of the Corporation; provided, however, if the holders of at least a majority of the shares of Preferred Stock then outstanding, voting on an as-if converted basis (a "**Preferred Majority**"), so elect by giving written notice to the Corporation before the effective date of a merger, consolidation, sale or reorganization that would otherwise be a Liquidating Event as defined herein, such merger, consolidation, sale or reorganization shall not be deemed a Liquidating Event and the provisions of Subsection D.7 shall apply, and provided further that a "Liquidating Event" shall not include any transaction or series of related transactions

principally undertaken for bona fide equity financing purposes in which cash is received by the Corporation or any successor or indebtedness of the Corporation is cancelled or converted, or a combination thereof. Upon the occurrence of any Liquidating Event that would involve the distribution of assets other than cash with respect to the outstanding shares of Preferred Stock, the amount of such distribution shall be the fair market value thereof at the time of such distribution as determined in good faith by the Board of Directors of the Corporation, and any securities to be distributed in such event shall be valued as follows:

(i) Securities not subject to investment letter or other similar restrictions on free marketability covered by subsection (ii) hereof:

(A) if traded on a securities exchange, the value shall be deemed to be the average of the closing prices of the securities on such exchange over the 30-day period ending three (3) business days prior to the closing;

(B) if actively traded over-the-counter, the value shall be deemed to be the average of the closing bid or sale prices (whichever is applicable) over the 30-day period ending three (3) business days prior to the closing; and

(C) if there is no active public market, the value shall be the fair market value thereof, as reasonably determined by the Board of Directors of the Corporation in good faith.

(ii) The method of valuation of securities subject to investment letter or other restrictions on free marketability (other than restrictions arising solely by virtue of a stockholder's status as an affiliate or former affiliate) shall be to make an appropriate discount from the market value determined as provided in clauses (A), (B) or (C) of subsection (i) hereof, to reflect the adjusted fair market value thereof, as reasonably determined by the Board of Directors of the Corporation in good faith.

5. Allocation of Escrow and Contingent Consideration. Upon the occurrence of any Liquidating Event, if any portion of the consideration payable to the stockholders of the Corporation is payable only upon satisfaction of contingencies (the "**Additional Consideration**"), the transaction documents for such Liquidating Event shall provide that (a) the portion of such consideration that is not Additional Consideration (such portion, the "**Initial Consideration**") shall be allocated among the holders of capital stock of the Corporation in accordance with this Section B as if the Initial Consideration were the only consideration payable in connection with such Liquidating Event and (b) any Additional Consideration which becomes payable to the stockholders of the Corporation upon satisfaction of such contingencies shall be allocated among the holders of capital stock of the Corporation in accordance with this Section B after taking into account the previous payment of the Initial Consideration as part of the same transaction. For the purposes of this Section 5, consideration placed into escrow or retained as holdback to be available for satisfaction of indemnification or similar obligations in connection with such Liquidating Event shall be deemed to be Additional Consideration.

C. Voting.

1. General Rights. Except as otherwise expressly provided herein or as required by law, the holder of each share of Preferred Stock shall be entitled to the number of votes equal to the number of shares of Common Stock into which such share of Preferred Stock could then be converted

and shall have voting rights and powers equal to the voting rights and powers of the Common Stock (except as otherwise expressly provided herein or as required by law, voting together with the Common Stock as a single class) and shall be entitled to notice of any stockholders' meeting in accordance with the Bylaws of the Corporation. Fractional votes shall not, however, be permitted and any fractional voting rights resulting from the above formula (after aggregating all shares of Common Stock into which shares of Preferred Stock held by each holder could be converted) shall be reduced to the nearest whole number.

2. **Protective Provisions.** In addition to any other rights provided by law or as set forth in this Restated Certificate, for so long as any shares of Preferred Stock are outstanding the Corporation shall not, without first obtaining the affirmative vote or written consent of the holders of at least a majority of the then outstanding shares of Preferred Stock, including holders of at least 70% of the then outstanding shares of Series B Preferred Stock (such required holders, the "**Requisite Preferred Holders**"), consenting or voting together as a separate class on an as-converted to Common Stock basis, take any of the following actions (whether by merger, consolidation, recapitalization or otherwise):

(a) authorize or effect any Liquidating Event;

(b) authorize or effect a merger, consolidation or share exchange between the Corporation and another entity or a sale, lease, license or other disposition of all or substantially all of the Corporation's assets, or effect a sale or other disposition which results in the holders of the Corporation's capital stock prior to the transaction owning less than fifty percent (50%) of the voting power of the Corporation's capital stock after the transaction, or some other reorganization or acquisition of the Corporation, whether or not any of the foregoing would constitute a Liquidating Event;

(c) redeem, purchase, pay any dividend on or otherwise acquire for value any shares of Common Stock or any Preferred Stock (other than employee, director or consultant shares repurchased at the lower of cost or fair market value pursuant to equity incentive agreements or other similar arrangements providing for the right to repurchase shares upon the termination of services) prior to the Preferred Stock;

(d) authorize or issue any shares of capital stock having rights, preferences or privileges superior to or on parity with the Preferred Stock, or authorize or issue any securities exchangeable, convertible or exercisable for shares of such capital stock;

(e) reclassify any shares of Common Stock or any other class or series of capital stock of the Corporation;

(f) increase the authorized number of shares of any class or series of capital stock of the Corporation;

(g) alter or change any of the powers, preferences, privileges or rights of the Preferred Stock;

(h) amend, repeal or add to any provision of this Restated Certificate or the Corporation's Bylaws as in effect on the date this Restated Certificate is filed with the Secretary of State of the State of Delaware (the "**Effective Date**");

(i) convert the Corporation into another form of business entity or into a corporation organized in a jurisdiction other than Delaware;

(j) change the fundamental business of the Corporation;

(k) increase or decrease the size of the Corporation's Board of Directors;

(l) create or authorize the creation of any debt security, other than equipment leases or bank lines of credit unless such debt security has received the prior approval of the Board of Directors of the Corporation, including the approval of the Series B Director then serving;

(m) create or hold capital stock in any subsidiary that is not a wholly-owned subsidiary or dispose of any subsidiary stock or all or substantially all of the assets of any subsidiary; or

(n) incur any indebtedness in excess of \$500,000.

3. Election of Directors. At each meeting of the stockholders held for the election of directors, or upon the taking of a written consent of stockholders for such purpose, (i) the holders of Common Stock shall be entitled, voting as a separate class, to elect one (1) member of the Board of Directors of the Corporation, (ii) for so long as at least 1,000,000 shares of the Series A Preferred Stock remain outstanding (subject to equitable adjustment for any Recapitalizations), the holders of Series A/A-1 Preferred Stock shall be entitled, voting together as a separate class on an as-converted to Common Stock basis, to elect two (2) members of the Board of Directors of the Corporation (the "**Series A Directors**"), and (iii) for so long as at least 1,000,000 shares of the Series B Preferred Stock remain outstanding, the holders of Series B Preferred Stock shall be entitled, voting together as a separate class on an as-converted to Common Stock basis, to elect one (1) member of the Board of Directors of the Corporation (the "**Series B Director**"). All remaining directors of the Corporation shall be elected by the holders of Common Stock and Preferred Stock, voting together as a single class on an as-converted to Common Stock basis. In the event the holders of Series B Preferred Stock, Series A/A-1 Preferred Stock or Common Stock, as the case may be, fail to elect a member of the Board of Directors of the Corporation as set forth above, such directorship shall remain vacant until a member of the Board of Directors is elected by the holders of Series B Preferred Stock, Series A/A-1 Preferred Stock or Common Stock, as the case may be, and no such directorship shall be filled by stockholders of the Corporation other than by the stockholders of the Corporation that are entitled to elect a person to fill such directorship, voting as a separate class. Any director who shall have been elected by the holders of Series B Preferred Stock, Series A/A-1 Preferred Stock or Common Stock, as the case may be may be removed during the aforesaid term of office, either with or without cause, by, and only by, the affirmative vote of the holders of a majority of the shares of Series B Preferred Stock, Series A/A-1 Preferred Stock or Common Stock, as the case may be, in accordance with the Bylaws of the Corporation, given either at a special meeting of such stockholders duly called for that purpose or pursuant to a written consent of stockholders, and any vacancy thereby created may be filled by the holders of a majority of the shares of the Series B Preferred Stock, Series A/A-1 Preferred Stock or Common Stock, as the case may be, represented at a meeting or pursuant to written consent.

D. Conversion Rights.

Each share of Preferred Stock shall be convertible at the option of the holder thereof, at any time after the issuance of such share, into fully paid and nonassessable shares of Common Stock of the Corporation. The number of shares of Common Stock into which each share of the Series A Preferred Stock, Series A-1 Preferred Stock or Series B Preferred Stock may be converted shall be determined by dividing the Series A Original Price by the Series A Conversion Price, the Series A-1 Original Price by the Series A-1 Conversion Price or the Series B Original Price by the Series B Conversion Price, as applicable, (each as determined as hereinafter provided) in effect at the time of the conversion.

1. Conversion Price. Before any adjustment pursuant to Section E hereof, the Series A conversion price (the “**Series A Conversion Price**”) shall be equal to the Series A Original Price, the Series A-1 conversion price (the “**Series A-1 Conversion Price**”) shall be equal to the Series A-1 Original Price and the Series B conversion price (the “**Series B Conversion Price**”) shall be equal to the Series B Original Price (the Series A Conversion Price, the Series A-1 Conversion Price and the Series B Conversion Price shall each be a “**Conversion Price**”).

2. Mechanics of Conversion. The holder of any shares of Preferred Stock may exercise the conversion rights as to such shares or any part thereof by delivering to the Corporation during regular business hours, at the office of any transfer agent of the Corporation for the Preferred Stock, or at the principal office of the Corporation or at such other place as may be designated by the Corporation, the certificate or certificates for the shares to be converted, duly endorsed for transfer to the Corporation or accompanied by a written instrument or instruments of transfer (if required by it), accompanied by written notice stating that the holder elects to convert all or a number of such shares represented by the certificate or certificates. Such notice shall also state such holder’s name or the names of the nominees in which such holder wishes the certificate or certificates for shares of Common Stock to be issued. Conversion shall be deemed to have been effected on the date when such delivery is made, and such date is referred to herein as the “**Conversion Date.**” As promptly as practicable thereafter the Corporation shall issue and deliver to such holder, at such office or other place designated by the Corporation, a certificate or certificates for the number of full shares of Common Stock to which such holder is entitled and a check for cash with respect to any fractional interest in a share of Common Stock as provided in Subsection D.3 below. All shares of Preferred Stock which shall have been surrendered for conversion as herein provided shall no longer be deemed to be outstanding and all rights with respect to such shares, including the rights, if any, to receive notices and to vote, shall immediately cease and terminate upon the Conversion Date, except only the right of the holder thereof to receive shares of Common Stock in exchange therefor. The holder shall be deemed to have become a stockholder of record with respect to the shares of Common Stock on the applicable Conversion Date. Upon conversion of only a portion of the number of shares of Preferred Stock represented by a certificate surrendered for conversion, the Corporation shall issue and deliver to the holder of the certificate so surrendered for conversion, at the expense of the Corporation, a new certificate covering the number of shares of Preferred Stock representing the unconverted portion of the certificate so surrendered.

3. Fractional Shares. No fractional shares of Common Stock or scrip shall be issued upon conversion of shares of Preferred Stock. If more than one share of Preferred Stock shall be surrendered for conversion at any one time by the same holder, the number of full shares of Common Stock issuable upon conversion thereof shall be computed on the basis of the aggregate number of shares of Preferred Stock so surrendered. Instead of any fractional shares of Common Stock that would otherwise be issuable upon conversion of any shares of Preferred Stock, the Corporation shall pay a cash adjustment in respect of such fractional interest equal to the fair market value of such fractional interest as determined in good faith by the Corporation’s Board of Directors.

4. Payment of Taxes. The Corporation shall pay any and all issue and transfer taxes that may be payable in respect of any issue or delivery of shares of Common Stock on conversion of the Preferred Stock pursuant hereto. The Corporation shall not, however, be required to pay any tax that may be payable in respect of any transfer involved in the issue and delivery of shares of Common Stock in a name other than that in which the Preferred Stock so converted was registered, and no such issue or delivery shall be made unless and until the person requesting such issue has paid to the Corporation the amount of any such tax or has established, to the satisfaction of the Corporation, that such tax has been paid.

5. Reservation of Stock Issuable Upon Conversion. The Corporation shall at all times reserve and keep available, out of its authorized but unissued Common Stock, solely for the purpose of effecting the conversion of the Preferred Stock, the full number of shares of Common Stock deliverable upon the conversion of all the Preferred Stock from time to time outstanding. The Corporation shall from time to time use its best effort to obtain necessary director and stockholder approvals, in accordance with the laws of the State of Delaware, to increase the authorized amount of its Common Stock if at any time the authorized amount of its Common Stock remaining unissued shall not be sufficient to permit the conversion of all of the shares of Preferred Stock at the time outstanding, and shall take all such actions as are necessary to increase such authorized amount of Common Stock upon obtaining such approvals. Before taking any action that would cause an adjustment reducing the Series A Conversion Price, Series A-1 Conversion Price or the Series B Conversion Price below the then-par value of the shares of Common Stock issuable upon the conversion of the Preferred Stock, the Corporation will take any corporate action that may, in the opinion of its counsel, be necessary in order that the Corporation may validly and legally issue fully paid and nonassessable shares of Common Stock at such adjusted Conversion Price.

6. Adjustment for Reclassification, Exchange and Substitution. If the Common Stock issuable upon the conversion of the Preferred Stock shall be changed into the same or a different number of shares of any class or classes of stock, whether by capital reorganization, reclassification, or otherwise (other than a subdivision or combination of shares or stock dividend provided for in Subsection E.1), then and in each such event the holder of each share of Preferred Stock shall have the right thereafter to convert such share into the kind and amount of shares of stock and other securities and property receivable upon such reorganization, reclassification, or other change, by holders of the number of shares of Common Stock into which such shares of Preferred Stock might have been converted immediately prior to such reorganization, reclassification, or change.

7. Reorganizations, Mergers or Consolidations. In case of any consolidation or merger of the Corporation with or into another corporation (other than a consolidation, merger or sale treated as a Liquidating Event pursuant to Subsection B.4 above), each share of Preferred Stock shall thereafter be convertible into the kind and amount of shares of stock or other securities or property to which a holder of the number of shares of Common Stock of the Corporation deliverable upon conversion of the Preferred Stock would have been entitled upon such consolidation, merger or sale; and in such case, appropriate adjustment (as determined in good faith by the Board of Directors of the Corporation) shall be made in the application of the provisions of Sections D and E with respect to the rights and interest thereafter of the holders of Preferred Stock, to the end that the provisions set forth in Sections D and E shall thereafter be applicable, as nearly as reasonably may be, in relation to any shares of stock or other property thereafter deliverable upon the conversion of the Preferred Stock.

8. Listing of Shares Issuable Upon Conversion. If any shares of Common Stock to be reserved for the purpose of conversion of shares of Preferred Stock require registration or listing with, or approval of, any governmental authority, stock exchange or other regulatory body under any federal or state law or regulation or otherwise, before such shares may be validly issued or delivered upon conversion, the Corporation will in good faith and as expeditiously as possible endeavor to secure such registration, listing or approval, as the case may be.

9. Valid Issuance. All shares of Common Stock that may be issued upon conversion of the shares of Preferred Stock will, upon issuance by the Corporation, be validly issued, fully paid and nonassessable and free from all taxes, liens and charges with respect to the issuance thereof.

10. No Dilution or Impairment. The Corporation will not, by amendment of this Restated Certificate or through any reorganization, transfer of assets, consolidation, merger, dissolution, issue or sale of securities or any other voluntary action, avoid or seek to avoid the observance or performance of any of the terms to be observed or performed hereunder by the Corporation, but will at all times in good faith assist in the carrying out of all of the provisions of Sections D and E and in the taking of all such action as may be necessary or appropriate in order to protect the conversion rights of the holders of Preferred Stock against impairment.

E. Adjustment of Conversion Prices.

The Conversion Prices from time to time in effect shall be subject to adjustment from time to time as follows.

1. Stock Splits, Dividends and Combinations. In case the Corporation shall at any time subdivide the outstanding shares of Common Stock or shall issue a dividend in Common Stock on its outstanding Common Stock without a corresponding subdivision of or dividend on each series of Preferred Stock, the Conversion Price in effect for each non-participating series of Preferred Stock immediately prior to such subdivision or the issuance of such dividend shall be proportionately decreased, and in case the Corporation shall at any time combine the outstanding shares of Common Stock into a lesser number of shares of Common Stock without a corresponding combination of each series of Preferred Stock, the Conversion Price in effect for each non-participating series of Preferred Stock immediately prior to such combination shall be proportionately increased, concurrently with the effectiveness of such subdivision, dividend or combination, as the case may be.

2. Noncash Dividends, Stock Purchase Rights, Capital Reorganizations and Dissolutions. In case:

(a) the Corporation shall take a record of the holders of its Common Stock for the purpose of entitling them to receive a dividend or any other distribution, payable otherwise than in cash; or

(b) the Corporation shall take a record of the holders of its Common Stock for the purpose of entitling them to subscribe for or purchase any shares of stock of any class or to receive any other rights; or

(c) of any capital reorganization of the Corporation, reclassification of the capital stock of the Corporation (other than a subdivision or combination of its outstanding shares of Common Stock), consolidation or merger of the Corporation with or into another Corporation that is not a Liquidating Event or conveyance of all or substantially all of the assets of the Corporation to another corporation that is not a Liquidating Event;

then, and in any such case, the Corporation shall cause to be mailed to the transfer agent for the Preferred Stock (if any) and to the holders of record of the outstanding Preferred Stock, at least ten (10) days prior to the date hereinafter specified, a notice stating the date on which (i) a record is to be taken for the purpose of such dividend, distribution or rights, or (ii) such reclassification, reorganization, consolidation, merger, conveyance, dissolution, liquidation or winding up is to take place and the date, if any is to be fixed, as of which holders of Common Stock of record shall be entitled to exchange their shares of Common Stock for securities or other property deliverable upon such reclassification, reorganization, consolidation, merger, conveyance, dissolution, liquidation or winding up.

3. Issuances at Less Than the Conversion Price. Upon the issuance or sale by the Corporation of:

(a) Common Stock for a consideration per share less than a Conversion Price in effect immediately prior to the time of such issue or sale;

or

(b) any Stock Purchase Rights (as hereinafter defined) where the consideration per share for which shares of Common Stock may at any time thereafter be issuable upon exercise thereof (or, in the case of Stock Purchase Rights exercisable for the purchase of Convertible Securities (as hereinafter defined), upon the subsequent conversion or exchange of such Convertible Securities) shall be less than a Conversion Price in effect immediately prior to the time of the issue or sale of such Stock Purchase Rights; or

(c) any Convertible Securities where the consideration per share for which shares of Common Stock may at any time thereafter be issuable pursuant to the terms of such Convertible Securities shall be less than a Conversion Price in effect immediately prior to the time of the issue or sale of such Convertible Securities;

other than an issuance of Common Stock pursuant to Subsections E.1 or E.6 hereof (any such issuance shall be referred to hereinafter as a "**Dilutive Issuance**"), then, forthwith upon such issue or sale, each then effective Conversion Price which exceeds the consideration per share received, shall be reduced concurrently with such issue in order to increase the number of shares of Common Stock into which the Preferred Stock, as applicable, is convertible to a price (calculated to the nearest cent) determined by the following formula:

$$CP1 = CP * \frac{N + C}{N + AS}$$

where:

CP1 = the Conversion Price as so adjusted;

CP = the former Conversion Price immediately prior to the Dilutive Issuance;

N = the number of shares of Common Stock outstanding immediately prior to such issuance (or deemed issuance) assuming exercise or conversion of all outstanding Convertible Securities and Stock Purchase Rights;

- C = the number of shares of Common Stock that the aggregate consideration received or deemed to be received by the Corporation for the total number of additional securities so issued or deemed to be issued would purchase if the purchase price per share were equal to CP; and
- AS = the number of shares of Common Stock so issued or deemed to be issued.

Notwithstanding the foregoing, the applicable Conversion Price shall not at such time be reduced if such reduction would be an amount less than \$0.01, but any such amount shall be carried forward and deduction with respect thereto made at the time of and together with any subsequent reduction that, together with such amount and any other amount or amounts so carried forward, shall aggregate \$0.01 or more.

4. Defined Terms. For purposes of this Section E, the following provisions will be applicable.

(a) “**Convertible Securities**” shall mean evidences of indebtedness, shares of stock (including, without limitation, the Preferred Stock) or other securities that are convertible into or exchangeable for, with or without payment of additional consideration, shares of Common Stock.

(b) “**Stock Purchase Rights**” shall mean any warrants, options or other rights to subscribe for, purchase or otherwise acquire any shares of Common Stock or any Convertible Securities.

(c) The Common Stock underlying the Convertible Securities and Stock Purchase Rights shall be deemed outstanding and issued or sold at the time of the issue or sale of the Convertible Security or Stock Purchase Right.

5. Determination of Consideration. The “consideration” actually received by the Corporation for the issuance, sale, grant or assumption of shares of Common Stock, Stock Purchase Rights or Convertible Securities, irrespective of the accounting treatment of such consideration, shall be valued as follows:

(a) in the case of cash, the net amount received by the Corporation after deduction of any accrued interest or dividends and before deducting any expenses paid or incurred and any underwriting commissions or concessions paid or allowed by the Corporation in connection with such issue or sale;

(b) in the case of consideration other than cash, the fair market value of such consideration, which shall not include the value of any Convertible Securities being converted or exchanged, as determined by the Board of Directors of the Corporation in good faith, after deducting any accrued interest or dividends; and

(c) with respect to the issuance of Stock Purchase Rights and Convertible Securities, the total consideration, if any, received by the Corporation as consideration for the issuance of the Stock Purchase Rights or the Convertible Securities, as the case may be, plus the minimum aggregate amount of additional consideration, if any, payable to the Corporation upon the exercise of such Stock Purchase Rights or upon the conversion or exchange of such Convertible Securities, as the case may be, in each case after deducting any accrued interest or dividends.

In the event of any change in (i) the consideration, if any, payable upon exercise of any Stock Purchase Rights or upon the conversion or exchange of any Convertible Securities, or (ii) the rate at which any Convertible Securities are convertible into or exchangeable for shares of Common Stock, the applicable Conversion Price, as computed upon the original issue thereof shall forthwith be readjusted to the Conversion Price that would have been in effect at such time had such Stock Purchase Rights or Convertible Securities provided for such changed purchase price, consideration or conversion rate, as the case may be, at the time initially granted, issued or sold. On the expiration of any Stock Purchase Rights not exercised or of any right to convert or exchange under any Convertible Securities not exercised, the Conversion Price then in effect shall forthwith be increased to the Conversion Price that would have been in effect at the time of such expiration had such Stock Purchase Rights or Convertible Securities never been issued. No readjustment of the Conversion Price pursuant to this paragraph shall (A) increase the Conversion Price by an amount in excess of the adjustment originally made to the Conversion Price in respect of the issue, sale or grant of the applicable Stock Purchase Rights or Convertible Securities, or (B) require any adjustment to the amount paid or number of shares of Common Stock received by any holder of Preferred Stock upon any conversion of any share of Preferred Stock prior to the date upon which such readjustment to the Conversion Price shall occur.

6. Exclusions for Adjustment for Issuances at Less Than the Conversion Price. Anything herein to the contrary notwithstanding, the Corporation shall not be required to make any adjustment of any Conversion Price in the case of: (a) Common Stock or Stock Purchase Rights issued or issuable to employees, officers, consultants or directors of the Corporation pursuant to any incentive plan, agreement or arrangement approved by the Board of Directors of the Corporation, which approval must include the Series B Director then serving; (b) securities issued in a Qualified Public Offering; (c) securities issued upon conversion of the Convertible Securities or the exercise of Stock Purchase Rights; or (d) Common Stock or Preferred Stock issued by way of dividend or other comparable distribution on the Preferred Stock (each of (a) through (d), "**Excluded Securities**").

7. Certificate of Adjustment. Upon the occurrence of each adjustment or readjustment of any Conversion Price pursuant to this Section E, the Corporation at its expense shall promptly compute such adjustment or readjustment in accordance with the terms thereof, and prepare and furnish to each holder of Preferred Stock affected thereby a certificate setting forth such adjustment or readjustment and showing in detail the facts upon which such adjustment or readjustment is based. The Corporation shall, upon the written notice at any time of any affected holder of Preferred Stock furnish or cause to be furnished to such holder a like certificate setting forth (a) such adjustment or readjustment, (b) the Conversion Price at the time in effect, and (c) the number of shares of Common Stock and the amount, if any, of other property that at the time would be received upon the conversion of such holder's shares.

F. Mandatory Conversion.

1. Mandatory Conversion Triggers. Each share of Preferred Stock shall automatically be converted into shares of Common Stock at the then-applicable conversion rate upon the occurrence of a Qualified Public Offering. "**Qualified Public Offering**" means a closing of an underwritten public offering pursuant to an effective registration statement under the Securities Act of 1933, as amended, covering the offer and sale of Common Stock of the Corporation to the public where the Corporation receives proceeds of not less than \$50,000,000 (net of underwriters discounts and commissions), and the price per share to the public yields a price per share to the public of not less than (a) if occurring on or prior to February 20, 2015, 1.50 times the Series B Original Price or (b) if occurring after February 20, 2015, 3.0 times the Series B Original Price. In addition, each share of Preferred Stock

shall automatically be converted into shares of Common Stock at the then applicable conversion rate upon the affirmative vote of the Requisite Preferred Holders, voting separately as a single class on an as-converted to Common Stock basis. All holders of record of shares of Preferred Stock will be given at least thirty (30) days prior written notice of the date fixed for mandatory conversion of the Preferred Stock and the event causing the mandatory conversion of the Preferred Stock into Common Stock. Such notice shall be sent by first class mail, postage prepaid, to each holder of record of Preferred Stock at such holder's address as shown in the records of the Corporation.

2. Mechanics of Mandatory Conversion. On or before the date so fixed for conversion, each holder of shares of the Preferred Stock shall surrender the certificate or certificates for all such shares to the Corporation at the place designated in such notice and shall thereafter receive certificates for the number of shares of Common Stock to which such holder is entitled. All certificates evidencing shares of Preferred Stock which are required to be surrendered for conversion on such date shall, from and after such date, be deemed to have been retired and cancelled and the shares of Preferred Stock represented thereby shall be deemed converted into shares of Common Stock for all purposes, notwithstanding the failure of the holder or holders thereof to surrender such certificates or warrants on or prior to such date. The mechanics for conversion and other provisions relating to conversion of Preferred Stock into Common Stock set forth elsewhere in this Restated Certificate shall apply to the mandatory conversion of the Preferred Stock.

G. Redemption

1. Redemption. Shares of Preferred Stock shall be redeemed by the Corporation at a price equal to the Series A Original Price, Series A-1 Original Price or Series B Original Price, as applicable, plus all declared but unpaid dividends as of the Redemption Date (the "**Redemption Price**"), in three (3) equal annual installments commencing not more than sixty (60) days after receipt by the Corporation at any time on or after the third anniversary of the first sale by the Corporation of the Series B Preferred Stock, from the Requisite Preferred Holders, voting together as a separate class on an as-converted to Common Stock basis, of written notice requesting redemption of all shares of Preferred Stock (a "**Redemption Request**"). Upon receipt of a Redemption Request, the Corporation shall apply all of its assets to any such redemption, and to no other corporate purpose, except to the extent prohibited by Delaware law governing distributions to stockholders. The date of each installment shall be referred to as a "**Redemption Date**."

2. Redemption Mechanics.

(a) Notice. The Corporation shall send written notice of the redemption (the "**Redemption Notice**") to each holder of record of Preferred Stock not less than 30 days prior to the Redemption Date. The Redemption Notice shall state:

(i) the number of shares of Preferred Stock held by the holder that the Corporation shall redeem on the Redemption Date specified in the Redemption Notice;

(ii) the Redemption Date and the Redemption Price;

(iii) that the holder is to surrender to the Corporation, in the manner and at the place designated, his, her or its certificate or certificates representing the shares of Preferred Stock to be redeemed.

(b) Number of Shares to Be Redeemed. On each Redemption Date, the Corporation shall redeem, on a pro rata basis in accordance with the number of shares of Preferred Stock, to be redeemed from each holder, that number of shares of Preferred Stock, determined by dividing (i) the total number of shares of each series of Preferred Stock to be redeemed immediately prior to such Redemption Date by (ii) the number of applicable remaining Redemption Dates (including the Redemption Date to which such calculation applies). If the redemption by the Corporation of all shares of Preferred Stock to be redeemed on such Redemption Date would be prohibited by Delaware law governing distributions to stockholders, the Corporation shall redeem a pro rata portion of each holder's redeemable shares of each series of Preferred Stock to the extent such redemption would not be prohibited by the DGCL governing distributions to stockholders, based on the respective amounts which would otherwise be payable in respect of the shares to be redeemed if the redemption of all such shares would not be prohibited by Delaware law governing distributions to stockholders, and shall redeem the remaining shares of Preferred Stock to have been redeemed as soon as practicable after the Corporation would not be prohibited from making such redemption under Delaware law governing distributions to stockholders.

(c) Exchange of Certificates; Payment. On or before the applicable Redemption Date, each holder of shares of Preferred Stock to be redeemed on the Redemption Date, unless such holder has exercised his, her or its right to convert such shares as provided in Section 4, shall surrender the certificate or certificates representing such shares (or, if such registered holder alleges that such certificate has been lost, stolen or destroyed, a lost certificate affidavit and agreement reasonably acceptable to the Corporation to indemnify the Corporation against any claim that may be made against the Corporation on account of the alleged loss, theft or destruction of such certificate) to the Corporation, in the manner and at the place designated in the Redemption Notice, and thereupon the Redemption Price for such shares shall be payable to the order of the person whose name appears on such certificate or certificates as the owner thereof. In the event less than all of the shares of Preferred Stock represented by a certificate are redeemed, a new certificate representing the unredeemed shares of Preferred Stock shall promptly be issued to such holder.

3. Rights Subsequent to Redemption. If the Redemption Notice shall have been duly given, and if on the Redemption Date the Redemption Price payable upon redemption of the shares of Preferred Stock to be redeemed on such Redemption Date is paid or tendered for payment or deposited with an independent payment agent so as to be available therefor in a timely manner, then notwithstanding that the certificates evidencing any of the shares of Preferred Stock so called for redemption shall not have been surrendered, dividends with respect to such shares of Preferred Stock shall cease to accrue after such Redemption Date and all rights with respect to such shares shall forthwith after the Redemption Date terminate, except only the right of the holders to receive the Redemption Price without interest upon surrender of their certificate or certificates therefor.

4. Redeemed or Otherwise Acquired Shares. Any shares of Preferred Stock that are redeemed or otherwise acquired by the Corporation or any of its subsidiaries shall be automatically and immediately cancelled and retired and shall not be reissued, sold or transferred. Neither the Corporation nor any of its subsidiaries may exercise any voting or other rights granted to the holders of Preferred Stock following redemption.

ARTICLE VI

The voting, dividend and liquidation rights of the holders of the Common Stock are subject to and qualified by the rights, powers and preferences of the holders of Preferred Stock set forth herein. The holders of the Common Stock are entitled to one vote for each share of Common Stock held by them at all meetings of stockholders (and for all written actions of stockholders in lieu of meetings).

ARTICLE VII

The number of directors of the Corporation, which constitute the whole Board of Directors of the Corporation, may be fixed by the Bylaws of the Corporation. Elections of directors may be, but shall not be required to be, by written ballot.

ARTICLE VIII

In furtherance and not in limitation of the powers conferred by the laws of the State of Delaware, the Board of Directors of the Corporation is expressly authorized to make, alter and repeal the Bylaws of the Corporation, subject to the power of the stockholders of the Corporation to alter or repeal any bylaw whether adopted by them or otherwise.

ARTICLE IX

The Corporation is to have perpetual existence.

ARTICLE X

Notwithstanding the provisions of Section 242 of the Delaware General Corporation Law but subject to obtaining any other vote expressly provided for herein, the number of authorized shares of Common Stock may be increased or decreased (but not below the number of shares of Common Stock then outstanding) by an affirmative vote of the holders of a majority of the outstanding capital stock of the Corporation.

ARTICLE XI

To the fullest extent permitted by the Delaware General Corporation Law as the same exists or as may hereafter be amended, no present or former director of the Corporation shall be personally liable to the Corporation or its stockholders for monetary damages for breach of fiduciary duty as a director. Neither any amendment nor repeal of this Article, nor the adoption of any provision of this Restated Certificate inconsistent with this Article, shall eliminate or reduce the effect of this Article in respect of any matter occurring, or any cause of action, suit or claim that, but for this Article, would accrue or arise, prior to such amendment, repeal or adoption of an inconsistent provision.

ARTICLE XII

The Corporation shall have the power to indemnify any person who was or is a party or is threatened to be made a party to, or testifies in, any threatened, pending or completed action, suit or proceeding, whether civil, criminal, administrative or investigative in nature, by reason of the fact such person is or was a director, officer or employee or agent of the Corporation, or is or was serving at the request of the Corporation as a director, officer, employee or agent of another corporation, partnership, joint venture, employee benefit plan, trust or other enterprise, against expenses (including attorney's

fees), judgments, fines and amounts paid in settlement actually and reasonably incurred by such person in connection with such action, suit or proceeding to the full extent permitted by law, and the Corporation may adopt bylaws or enter into agreements with any such person for the purpose of providing for such indemnification.

ARTICLE XIII

All provisions relating to any exchange, reclassification or cancellation of issued shares are set forth in this Restated Certificate.

ARTICLE XIV

Any shares of Preferred Stock redeemed, purchased, converted or otherwise acquired by the Corporation shall be deemed retired and shall be cancelled and may not under any circumstances thereafter be reissued or otherwise disposed of by the Corporation.

ARTICLE XV

Any of the rights, powers, preferences and other terms of a series of Preferred Stock set forth herein may be waived on behalf of all holders of such series of Preferred Stock only by the affirmative written consent or affirmative vote of (a) in the case of Series A Preferred Stock or Series A-1 Preferred Stock, the holders of a majority of the then outstanding shares of Series A/A-1 Preferred Stock or (b) in the case of Series B Preferred Stock, the Requisite Preferred Holders, in each case voting together as a separate class on an as-converted to Common Stock basis.

ARTICLE XVI

The Corporation renounces, to the fullest extent permitted by law, any interest or expectancy of the Corporation in, or in being offered an opportunity to participate in, any Excluded Opportunity. An “**Excluded Opportunity**” is any matter, transaction or interest that is presented to, or acquired, created or developed by, or which otherwise comes into the possession of, (i) any director of the Corporation who is not an employee of the Corporation or any of its subsidiaries, or (ii) any holder of Preferred Stock or any partner, member, director, stockholder, employee or agent of any such holder, other than someone who is an employee of the Corporation or any of its subsidiaries (collectively, “**Covered Persons**”), unless such matter, transaction or interest is presented to, or acquired, created or developed by, or otherwise comes into the possession of, a Covered Person expressly and solely in such Covered Person’s capacity as a director of the Corporation.

* * * * *

CLEARSIDE BIOMEDICAL, INC.

**SECOND AMENDED AND RESTATED
INVESTOR RIGHTS AGREEMENT**

August 29, 2014

SECOND AMENDED AND RESTATED
INVESTOR RIGHTS AGREEMENT

This Second Amended and Restated Investor Rights Agreement (this “*Agreement*”) is entered into as of the 29th day of August, 2014, by and among Clearside Biomedical, Inc., a Delaware corporation (the “*Company*”), and the holders of shares of the Company’s Preferred Stock from time to time outstanding (the “*Investor Stock*”) listed on Exhibit A attached hereto (the “*Investors*”).

RECITAL

WHEREAS, certain of the Investors (the “*Prior Investors*”) are holders of shares of the Company’s Series A Preferred Stock and Series A-1 Preferred Stock and are party to an Amended and Restated Investor Rights Agreement dated as of January 31, 2013, as amended from time to time (the “*Prior Agreement*”);

WHEREAS, in connection with the issuance and sale of shares of Series B Preferred Stock to certain of the Investors pursuant to that certain Series B Preferred Stock and Warrant Purchase Agreement, dated as of the date hereof, by and among the Company and the Investors (the “*Purchase Agreement*”), the parties to the Prior Agreement desire to amend and restate the Prior Agreement in its entirety, to accept the rights, obligations and covenants hereof in lieu of their rights, obligations and covenants under the Prior Agreement.

AGREEMENT

NOW, THEREFORE, in consideration of the mutual agreements, covenants and conditions contained herein, the Company and the Investors hereby agree as follows.

Section 1. RESTRICTIONS ON TRANSFER

1.1 Restrictive Legend. Each certificate representing (a) the Investor Stock, (b) the Common Stock of the Company (the “*Common Stock*”) issued upon conversion of the Investor Stock, and (c) any other securities issued in respect of the Investor Stock or Common Stock issued upon conversion of the Investor Stock upon any stock split, stock dividend, recapitalization, merger, consolidation or similar event, shall (unless otherwise permitted by the provisions of Section 1.2 below) be stamped or otherwise imprinted with a legend in substantially the following form (in addition to any legend required under applicable state securities laws).

THE SECURITIES REPRESENTED BY THIS CERTIFICATE HAVE NOT BEEN REGISTERED UNDER THE SECURITIES ACT OF 1933, AS AMENDED, OR APPLICABLE STATE SECURITIES LAWS. THESE SECURITIES HAVE BEEN ACQUIRED FOR INVESTMENT AND NOT WITH A VIEW TO DISTRIBUTION OR RESALE, AND MAY NOT BE SOLD, MORTGAGED, PLEDGED, HYPOTHECATED OR OTHERWISE TRANSFERRED WITHOUT AN EFFECTIVE REGISTRATION STATEMENT FOR SUCH SECURITIES UNDER THE SECURITIES ACT OF 1933, AS AMENDED, AND ANY APPLICABLE STATE SECURITIES LAWS, OR THE AVAILABILITY OF AN EXEMPTION FROM THE REGISTRATION PROVISIONS OF THE SECURITIES ACT OF 1933, AS

AMENDED, AND APPLICABLE STATE SECURITIES LAWS. COPIES OF THE INVESTOR RIGHTS AGREEMENT PROVIDING FOR RESTRICTIONS ON TRANSFER OF THESE SECURITIES MAY BE OBTAINED UPON WRITTEN REQUEST BY THE HOLDER OF RECORD OF THIS CERTIFICATE TO THE SECRETARY OF THE CORPORATION AT THE PRINCIPAL EXECUTIVE OFFICES OF THE CORPORATION.

Each stockholder consents to the Company's making a notation on its records and giving instructions to any transfer agent of the Investor Stock or the Common Stock in order to implement the restrictions on transfer established in this Section 1. Such legend shall be removed by the Company from any certificate at such time as the holder of the shares represented by the certificate satisfies the requirements of Rule 144(d) under the Securities Act of 1933, as amended (the "**1933 Act**").

1.2 Notice of Proposed Transfers. The holder of each certificate representing company capital stock (the "**Company Stock**") by acceptance thereof agrees to comply in all respects with the provisions of this Section 1.2. Prior to any proposed sale, assignment, transfer or pledge of any Company Stock, unless there is in effect a registration statement under the 1933 Act and any applicable state securities laws covering the proposed transfer, the holder thereof shall give written notice to the Company of such stockholder's intention to effect such transfer, sale, assignment or pledge. Each such notice shall describe the manner and circumstances of the proposed transfer, sale, assignment or pledge in sufficient detail and, if reasonably requested by the Company, shall be accompanied at such holder's expense by a written opinion of legal counsel who shall, and whose legal opinion shall, be reasonably satisfactory to the Company addressed to the Company, to the effect that the proposed transfer of the Company Stock may be effected without such registration. Each certificate evidencing the Company Stock transferred as above provided shall bear, except if such transfer is made pursuant to Rule 144, the appropriate restrictive legend set forth in Section 1.1 above, except that such certificate shall not bear such restrictive legend if in the opinion of counsel for such holder and the Company such legend is not required in order to establish compliance with any provisions of the 1933 Act or any applicable state securities laws. Notwithstanding the foregoing, no such opinion of counsel shall be necessary for a transfer by a stockholder which is (a) a partnership transferring to its partners or former partners in accordance with the partnership interests, (b) a limited liability company transferring to its members or former members in accordance with their interest in the limited liability company, or (c) a corporation transferring to its stockholders in accordance with their interest in the corporation; provided that in each case the transferee will be subject to the terms of this Agreement to the same extent as if he or she were an original stockholder hereunder.

Section 2. PREEMPTIVE RIGHTS

2.1 Certain Definitions. As used in this Section 2:

(a) The term "**Eligible Holder**" shall mean a holder of Investor Stock, that holds (either individually or collectively with its affiliates) at least 500,000 shares of the outstanding Preferred Stock of the Company (subject to adjustment in the case of a stock split, stock dividend, recapitalization, merger, consolidation or similar event).

(b) The term "**New Securities**" shall mean any capital stock of the Company, whether now authorized or not, and rights, options or warrants to purchase such capital stock, and securities of any type whatsoever that are, or may become, convertible into such capital stock; provided that the term "New Securities" does not include the following issuances: (i) Excluded Securities (as such term is defined in the Company's Fourth Amended and Restated Certificate of Incorporation (as it may be amended from time to time, the "**Charter**"); or (ii) the securities sold or to be sold in the future pursuant to the Purchase Agreement.

(c) The term “**Pro Rata Share**” shall mean the ratio, (i) the numerator of which is the number of shares of Common Stock held by such Eligible Holder, or issuable to such Eligible Holder upon the conversion of shares of Investor Stock held by such Eligible Holder, on the date of the Company’s written notice pursuant to Section 2.3 hereof, and (ii) the denominator of which is the number of shares of Common Stock outstanding, assuming for this purpose conversion or exercise of all securities convertible into or exercisable for Common Stock of the Company.

2.2 **Preemptive Rights.** The Company hereby grants to each Eligible Holder, subject to the terms and conditions specified in this Section 2, the right of first refusal to purchase, on the terms and conditions set forth in the Company’s notice pursuant to Section 2.3 hereof, up to its Pro Rata Share of all New Securities that the Company may, from time to time, propose to sell and issue.

2.3 **Required Notices.** In the event the Company proposes to undertake an issuance of New Securities, it shall give each Eligible Holder written notice of its intention, describing the type of New Securities, the price and the general terms upon which the Company proposes to issue the same. Each Eligible Holder shall have thirty (30) days from the date of any such notice to exercise its preemptive right under Section 2.2 hereof to purchase such New Securities for the price and upon the general terms specified in the notice by giving written notice to the Company and stating therein the quantity of New Securities to be purchased.

2.4 **Company’s Right to Sell.** If not all of the Eligible Holders elect to purchase their Pro Rata Share of the New Securities, then the Company shall promptly notify in writing the Eligible Holders who do so elect and shall offer such Eligible Holders the right to acquire such unsubscribed securities on a *pro rata* basis. The Eligible Holders shall have five (5) days after receipt of such notice to notify the Company of its election to purchase all or a portion of such unsubscribed securities. The Company shall have ninety (90) days after the thirty (30) day period described in Section 2.3 hereof to sell all such New Securities respecting which the Eligible Holders’ preemptive rights hereunder were not exercised, at a price and upon terms no more favorable in any material respect to the purchasers thereof than specified in the Company’s notice. In the event the Company has not sold all such New Securities within such ninety (90) day period, the Company shall not thereafter issue or sell any New Securities without first offering such New Securities to the Eligible Holders in the manner provided herein.

2.5 **Assignment of Preemptive Rights.** The rights contained in this Section 2 may be assigned or otherwise conveyed to transferees or assignees of Eligible Holders; provided that (i) such transfer is effected in compliance with Section 1.2 hereof, (ii) such transferee (A) is a current or former principal, manager, member, limited partner, general partner, stockholder, subsidiary, officer or affiliate of such transferor of the capital stock, (B) is a family member of the transferor or a trust or other similar entity for the benefit of the transferor or a family member of the transferor, or (C) acquires at least 250,000 shares of the transferor’s capital stock (as adjusted for stock splits, stock dividends, recapitalizations and other combinations), (iii) such transferee agrees to be subject to all restrictions set forth in this Agreement, (iv) the Company is provided with written notice of such transfer and (v) such transferee is an accredited investor or otherwise eligible under applicable securities law to acquire the New Securities without the same being registered under the 1933 Act or any similar law.

2.6 **Expiration of Right.** The rights granted under this Section 2 shall not apply to, and shall expire upon, the first to occur of (a) the effectiveness of a registration statement for the sale of the Company’s shares of Common Stock in a firm commitment underwritten public offering registered under the 1933 Act (a “**Qualified Public Offering**”), or (b) a merger, consolidation, sale or reorganization that constitutes a Liquidating Event under the Charter.

Section 3. COMPANY COVENANTS

3.1 Affirmative Covenants.

(a) Financial Statements and Information. The Company will cause to be furnished to each Major Investor (as defined below) the following reports (prepared in accordance with United States generally accepted accounting principles applied on a consistent basis, except that the unaudited reports may not contain notes or reserves and will be subject to year-end adjustment), provided, however, that the Company shall not be obligated pursuant to this Section 3.1(a) to provide financial information to any person whom the Company reasonably believes is a competitor of the Company; provided, further, that the Company agrees and acknowledges that Santen Pharmaceutical Co., Ltd. ("**Santen**") shall not be deemed a competitor for purposes of Sections 3.1(a) and 3.1(b). As used herein, the term "**Major Investor**" means any Investor owning (either individually or collectively with its affiliates) not less than 500,000 of the outstanding shares of the Company's Preferred Stock and each transferee who holds no less than that number of shares of the Company's Preferred Stock.

(i) As soon as practicable, but in any event within one hundred twenty (120) days after the end of each fiscal year of the Company, the Company will furnish to each Major Investor (A) reviewed or audited consolidated balance sheets of the Company and its subsidiaries, if any, as at the end of such fiscal year, and reviewed or audited consolidated statements of income and losses, stockholders' equity and cash flows of the Company and its subsidiaries, if any, for such fiscal year, setting forth in each case in comparative form the figures for the previous fiscal year, if any, all in reasonable detail and accompanied by a report and opinion thereon by independent auditors selected by the Company's Board of Directors (the "**Board**"), and (B) a copy of such auditors' management letter prepared in connection therewith, if any.

(ii) As soon as practicable after the end of each quarter, but in any event within thirty (30) days after the end of each such quarter, the Company will furnish to each Major Investor the unaudited consolidated balance sheets of the Company and its subsidiaries, if any, as of the end of such quarter, an up-to-date capitalization table and its unaudited consolidated statements of income and losses, stockholders' equity and cash flows for such quarter, setting forth in each case in comparative form the figures for the corresponding period of the preceding fiscal year, all in reasonable detail except that such financial statements may not contain notes or reserves and will be subject to year-end adjustment.

(iii) The Company will furnish to each Major Investor with reasonable promptness, such other information respecting the business, properties or the condition or operations, financial or other, of the Company or any subsidiary as any Major Investor may from time to time reasonably request or as determined by the Board.

(iv) As soon as practicable, but in any event no less than thirty (30) days before the beginning of each fiscal year, the Company will furnish to each Major Investor an annual operating plan and budget for the following fiscal year (which budget and plan shall include capital and operating expense budgets, cash flow projections, profit and loss projections and projected balance sheets for such year on a monthly basis), accompanied by a report from the CEO detailing the assumptions underlying the budget and any other information necessary to make such budget and plan accurate and not misleading, and, as soon as practicable after the adoption thereof, copies of any revisions to such annual operating plan.

(b) Inspection. The Company shall permit each Major Investor and its transferee(s) (provided such transfer is effected in compliance with Section 1.2 hereof), its attorney or its other representative, after executing a confidentiality agreement reasonably acceptable to the Board, to visit and inspect the Company's properties, to examine the Company's books of account and other records, to make copies or extracts therefrom and to discuss the Company's affairs, finances and accounts with its officers, management, employees and independent auditors all at such reasonable times during the Company's normal business hours and as often as such Major Investor or transferee may reasonably request; provided, however, that the Company shall not be obligated pursuant to this Section 3.1(b) to provide trade secrets or confidential information or to provide information to any person whom the Company reasonably believes is a competitor of the Company; provided, further, that such Investor shall bear any out-of-pocket costs or expenses of such investigations or inquiries.

(c) Confidentiality. Unless otherwise expressly set forth in another agreement between an Investor and the Company, each Investor agrees not to use Confidential Information (as hereinafter defined) of the Company for its own use or for any purpose except to evaluate and enforce its equity investment in the Company. Except as permitted under this Section, each Investor agrees to use all reasonable efforts not to disclose such Confidential Information to any third parties. Each Investor shall undertake to treat such Confidential Information in a manner consistent with the treatment of its own information of such proprietary nature and agrees that it shall protect the confidentiality of and use all reasonable efforts to prevent disclosure of the Confidential Information to prevent it from falling into the public domain or the possession of unauthorized persons. Each transferee of any Investor who receives Confidential Information shall agree to be bound by such provisions. For purposes of this Section, "**Confidential Information**" means any information, trade secrets, data, or know-how, including, but not limited to, the Company's patent applications, test or clinical data, licenses, research, products, services, development, inventions, consultants' or advisors' identities, samples, processes, designs, engineering, marketing, finances, or business partners disclosed by the Company either directly or indirectly in writing, orally or by drawings or inspection of samples. Confidential Information does not include information, technical data or know-how which (i) is in the Investor's possession at the time of disclosure as shown by Investor's files and records immediately prior to the time of disclosure; (ii) before or after it has been disclosed to the Investor, it is part of the public knowledge or literature, not as a result of any action or inaction of the Investor; (iii) is approved for release by written authorization of Company; or (iv) is rightfully disclosed to Investor by a third party without restriction. The provisions of this Section shall not apply (a) to the extent that an Investor is required to disclose Confidential Information pursuant to any law, statute, rule or regulation or any order of any court or judicial process or pursuant to any direction, request or requirement (whether or not having the force of law but if not having the force of law being of a type with which institutional investors in the relevant jurisdiction are accustomed to comply) of any self-regulating organization or any governmental, fiscal, monetary or other authority; (b) to the disclosure of Confidential Information to an Investor's employees, counsel, accountants or other professional advisors; (c) to the extent that an Investor needs to disclose Confidential Information for the protection of any of such Investor's rights or interest against the Company, whether under this Agreement or otherwise; or (d) to the disclosure of Confidential Information to a prospective transferee of securities which agrees to be bound by the provisions of this Section 3.1(c) in connection with the receipt of such Confidential Information.

(d) Reservation of Common Stock. The Company will at all times reserve and keep available, solely for issuance and delivery upon the conversion of the Investor Stock, all shares of Common Stock issuable upon such conversion.

(e) Board Matters. The reasonable out-of-pocket expenses of members of the Board associated with attending meetings or business related to the Company will be borne by the Company, and all directors will be treated identically with regard to expense reimbursement related to their service as members of the Board. The Board shall meet at least quarterly, unless otherwise agreed to by a vote of a majority of the Board.

(f) Observer Rights.

(i) For so long as it or its affiliates (i) hold less than two (2) seats on the Board; and (ii) own at least 500,000 shares of the Preferred Stock of the Company, Hatteras shall have the right to designate one person to receive notice of all meetings of the Board, to attend any such meeting as a nonvoting observer and to comment for the record at any such meeting (for purposes of this Section 3.1(f)(i), the term “meeting” shall be deemed to include all “executive sessions” and any other similar meeting of all or part of the Board or any committee thereof); provided, however, that such observer shall agree to hold in confidence and trust and to act in a fiduciary manner with respect to all information so provided; and provided further, that the Company reserves the right to withhold any information and to exclude such observer from any meeting or portion thereof if access to such information or attendance at such meeting could adversely affect the attorney-client privilege between the Company and its counsel or result in disclosure of trade secrets or a conflict of interest, or if such Investor or its representative is a competitor of the Company. Each observer so appointed as provided above shall sign a confidentiality agreement reasonably acceptable to the Board of the Company prior to his or her first attendance to his or her first meeting of the Board.

(ii) Until the later of January 31, 2016 or such time as Santen does not hold at least 5% of the Company’s capital stock (on an as converted, fully diluted basis), Santen shall have the right to designate one person to receive notice of all meetings of the Board, to attend any such meeting as a nonvoting observer, to comment for the record at any such meeting and to receive copies of all notices, minutes, consents, and other materials that it provides, at the same time such materials are provided, to its members of the Board; provided, however, that such observer shall agree to hold in confidence and trust and to act in a fiduciary manner with respect to all information so provided; and provided further, that the Company reserves the right to withhold any information and to exclude such observer from any meeting or portion thereof if (a) access to such information or attendance at such meeting could adversely affect the attorney-client privilege between the Company and its counsel, (b) access to such information or attendance at such meeting could result in disclosure of trade secrets or a conflict of interest, (c) such Investor or its representative is a competitor of the Company or (d) such information or materials relate to the use of the Company’s technology with any proprietary pharmaceutical molecule or formulation of a third party, including preclinical and clinical data and the terms of any agreement related to the combined use of Company technology with any such third party molecule or formulation. The rights in this Section 3.1(f)(ii) are personal to Santen and shall in no event be assignable to any person or entity without the express consent of the Company.

(iii) For so long he or his affiliates own at least 300,000 shares of the capital stock of the Company (subject to adjustment for stock splits, consolidations, reclassifications, etc.), Mark Prausnitz shall have the right to receive notice of all meetings of the Board, to attend any such meeting as a nonvoting observer, to comment for the record at any such meeting and to receive copies of all notices, minutes, consents, and other materials that it provides, at the same time such materials are provided, to its members of the Board, provided that the Company reserves the right to withhold any information if access to such information could adversely affect the attorney-client privilege between the Company and its counsel. Mr. Prausnitz shall treat all information received by the Company in accordance with the terms set forth in Section 3.1(c) above.

(iv) For so long as MGC Venture Partners 2013, L.P. and its affiliates (collectively, “**MGC**”) own at least 250,000 shares of the Preferred Stock of the Company (subject to adjustment for stock splits, consolidations, reclassifications, etc.), MGC shall have the right to designate

one person to receive notice of all meetings of the Board, to attend any such meeting as a nonvoting observer, to comment for the record at any such meeting and to receive copies of all notices, minutes, consents, and other materials that it provides, at the same time such materials are provided, to its members of the Board; provided, however, that such observer shall agree to hold in confidence and trust and to act in a fiduciary manner with respect to all information so provided; and provided further, that the Company reserves the right to withhold any information and to exclude such observer from any meeting or portion thereof if access to such information or attendance at such meeting could adversely affect the attorney-client privilege between the Company and its counsel or result in disclosure of trade secrets or a conflict of interest, or if such Investor or its representative is a competitor of the Company.

(v) For so long as RMI Investments S.a.r.l. and its affiliates (collectively, “**RMI**”) own at least 250,000 shares of the Preferred Stock of the Company (subject to adjustment for stock splits, consolidations, reclassifications, etc.), RMI shall have the right to designate one person to receive notice of all meetings of the Board, to attend any such meeting as a nonvoting observer, to comment for the record at any such meeting and to receive copies of all notices, minutes, consents, and other materials that it provides, at the same time such materials are provided, to its members of the Board; provided, however, that such observer shall agree to hold in confidence and trust and to act in a fiduciary manner with respect to all information so provided; and provided further, that the Company reserves the right to withhold any information and to exclude such observer from any meeting or portion thereof if access to such information or attendance at such meeting could adversely affect the attorney-client privilege between the Company and its counsel or result in disclosure of trade secrets or a conflict of interest, or if such Investor or its representative is a competitor of the Company.

(g) Committees of the Board. The Series B Director shall be included as a member of each committee of the Board currently existing or hereafter established by the Board.

(h) Directors and Officers Insurance. The Company will maintain a Directors and Officers insurance policy in an amount of at least five (5) million dollars (\$5,000,000).

(i) Successor Indemnification. If the Company or any of its successors or assignees consolidates with or merges into any other Person and is not the continuing or surviving corporation or entity of such consolidation or merger, then to the extent necessary, proper provision shall be made so that the successors and assignees of the Company assume the obligations of the Company with respect to indemnification of members of the Board as in effect immediately before such transaction, whether such obligations are contained in the Company’s Bylaws, its Certificate of Incorporation, an indemnification agreement between the Company and each member of the Board or elsewhere, as the case may be.

(j) Key Person Insurance. The Company hereby covenants and agrees that it shall use commercially reasonable efforts to secure and maintain “key man” life and disability insurance on Daniel White in an amount acceptable to the Board. The proceeds of such policy shall be payable to the Company.

(k) Employee Agreements. The Company has previously caused or will cause (i) each person now or hereafter employed by it or by any subsidiary (or engaged by the Company or any subsidiary as a consultant/independent contractor) with access to confidential information and/or trade secrets to enter into a nondisclosure and proprietary rights assignment agreement; and (ii) each Key Employee to enter into a one (1) year noncompetition and nonsolicitation agreement, substantially in the form approved by the Board. For purposes of this Agreement, the term “**Key Employee**” shall mean each of Henry Edelhauser, Samirkumar Patel, Mark Prausnitz, Daniel H. White and Vladmir Zarnitsyn, each executive-level employee (including, division director and vice president-level positions) as well as any employee who, either alone or in concert with others, develops, invents, programs, or designs any material intellectual property of the Company.

3.2 Negative Covenants. During the term of this Agreement and so long as the holders of Series B Preferred Stock are entitled to elect a Series B Director (as such term is defined in the Charter) (each member of the Board so elected by the holders of Series B Preferred Stock, a “**Series B Director**”) pursuant to the terms of the Charter, and notwithstanding anything to the contrary contained in any other agreements, the following actions of the Company shall require the prior approval of the Board (which approval must specifically include the affirmative vote, consent or approval of the Series B Director then serving):

(a) make any loan or advance to, or own any stock or other securities of, any subsidiary or other corporation, partnership or other entity unless it is wholly owned by the Company;

(b) make any loan or advance to any person, including, any employee or director, except advances and similar expenditures in the ordinary course of business or under the terms of an employee stock or option plan approved by the Board;

(c) guarantee any indebtedness except for trade accounts of the Company or any subsidiary arising in the ordinary course of business;

(d) make any investment inconsistent with any investment policy approved by the Board;

(e) incur any aggregate indebtedness in excess of US\$100,000 that is not already included in a Board-approved budget, other than trade credit incurred in the ordinary course of business;

(f) enter into or be a party to any transaction with any director, officer or employee of the Company or any “associate” (as defined in Rule 12b-2 promulgated under the Exchange Act) of any such person, or transactions made in the ordinary course of business and pursuant to reasonable requirements of the Company’s business and upon fair and reasonable terms that are approved by a majority of the Board;

(g) hire, fire, or change the compensation of the executive officers, including approving any option grants to such executive officers;

(h) change the principal business of the Company, enter new lines of business, or exit the current line of business;

(i) sell, assign, license, sublicense, pledge or encumber material technology or intellectual property, other than licenses granted in the ordinary course of business; or

(j) enter into any corporate strategic relationship involving the payment, contribution or assignment by the Company or to the Company of assets greater than US\$500,000 other than in the ordinary course of business.

3.3 Expiration of Covenants. The covenants set forth in this Section 3 shall expire and be of no further force or effect upon the effectiveness of a Qualified Public Offering (as defined in Section 2.6 hereof). After such time, the Investors shall be entitled to receive such annual and quarterly reports as the Company shall distribute to its stockholders generally.

Section 4. REGISTRATION RIGHTS

4.1 Certain Definitions. For purposes of this Section 4:

(a) “**1934 Act**” means the Securities Exchange Act of 1934, as amended, and the rules and regulations promulgated thereunder.

(b) “**Damages**” means any loss, damage, claim or liability (joint or several) to which a party hereto may become subject under the 1933 Act, the 1934 Act, or other federal or state law, insofar as such loss, damage, claim or liability (or any action in respect thereof) arises out of or is based upon: (i) any untrue statement or alleged untrue statement of a material fact contained in any registration statement of the Company, including any preliminary prospectus or final prospectus contained therein or any amendments or supplements thereto; (ii) an omission or alleged omission to state therein a material fact required to be stated therein, or necessary to make the statements therein not misleading; or (iii) any violation or alleged violation by the indemnifying party (or any of its agents or Affiliates) of the 1933 Act, the 1934 Act, any state securities law, or any rule or regulation promulgated under the 1933 Act, the 1934 Act, or any state securities law.

(c) “**Excluded Registration**” means (i) a registration relating to the sale of securities to employees of the Company or a subsidiary pursuant to a stock option, stock purchase, or similar plan; (ii) a registration relating to an SEC Rule 145 transaction; (iii) a registration on any form that does not include substantially the same information as would be required to be included in a registration statement covering the sale of the Registrable Securities; or (iv) a registration in which the only Common Stock being registered is Common Stock issuable upon conversion of debt securities that are also being registered.

(d) “**Form S-1**” means such form under the Securities Act as in effect on the date hereof or any successor registration form under the Securities Act subsequently adopted by the SEC.

(e) “**Form S-3**” means such form under the Securities Act as in effect on the date hereof or any registration form under the Securities Act subsequently adopted by the SEC that permits incorporation of substantial information by reference to other documents filed by the Company with the SEC.

(f) “**Holder**” means any holder of Registrable Securities who is a party to this Agreement.

(g) “**Immediate Family Member**” means a child, stepchild, grandchild, parent, stepparent, grandparent, spouse, sibling, mother-in-law, father-in-law, son-in-law, daughter-in-law, brother-in-law, or sister-in-law, including, adoptive relationships, of a natural person referred to herein.

(h) “**Initiating Holders**” means, collectively, Holders who properly initiate a registration request under this Agreement.

(i) “**IPO**” means the Company’s first underwritten public offering of its Common Stock under the 1933 Act.

(j) “**Person**” means any individual, corporation, partnership, trust, limited liability company, association or other entity.

(k) “**Registrable Securities**” means (i) the Common Stock issuable or issued upon conversion of the Company’s Preferred Stock; (ii) any Common Stock, or any Common Stock issued or issuable (directly or indirectly) upon conversion and/or exercise of any other securities of the Company,

acquired by the Investors after the date hereof; and (iii) any Common Stock issued as (or issuable upon the conversion or exercise of any warrant, right, or other security that is issued as) a dividend or other distribution with respect to, or in exchange for or in replacement of, the shares referenced in clauses (i) and (ii) above.

(l) “**Registrable Securities then outstanding**” means the number of shares determined by adding the number of shares of outstanding Common Stock that are Registrable Securities and the number of shares of Common Stock issuable (directly or indirectly) pursuant to then exercisable and/or convertible securities that are Registrable Securities.

(m) “**SEC**” means the Securities and Exchange Commission.

(n) “**Selling Expenses**” means all underwriting discounts, selling commissions, and stock transfer taxes applicable to the sale of Registrable Securities, and fees and disbursements of counsel for any Holder, except for the fees and disbursements of the Selling Holder Counsel borne and paid by the Company as provided in Subsection 4.7.

4.2 Demand Registration.

(a) If at any time after the earlier of (i) three (3) years after the date of this Agreement or (ii) one hundred eighty (180) days after the effective date of the registration statement for the IPO, the Company receives a request from Holders holding forty percent (40%) of the Registrable Securities then outstanding that the Company file a Form S-1 registration statement with respect to the Registrable Securities then outstanding if the anticipated aggregate offering price, net of Selling Expenses, would exceed \$10 million, then the Company shall (x) within ten (10) days after the date such request is given, give notice thereof (the “**Demand Notice**”) to all Holders other than the Initiating Holders; and (y) as soon as practicable, and in any event within sixty (60) days after the date such request is given by the Initiating Holders, file a Form S-1 registration statement under the Securities Act covering all Registrable Securities that the Initiating Holders requested to be registered and any additional Registrable Securities requested to be included in such registration by any other Holders, as specified by notice given by each such Holder to the Company within twenty (20) days of the date the Demand Notice is given, and in each case, subject to the limitations of Subsections 4.2(c) and 4.4.

(b) If at any time when it is eligible to use a Form S-3 registration statement, the Company receives a request from Holders of at least twenty percent (20%) of the Registrable Securities then outstanding that the Company file a Form S-3 registration statement with respect to outstanding Registrable Securities of such Holders having an anticipated aggregate offering price, net of Selling Expenses, of at least \$1 million, then the Company shall (i) within ten (10) days after the date such request is given, give a Demand Notice to all Holders other than the Initiating Holders; and (ii) as soon as practicable, and in any event within forty-five (45) days after the date such request is given by the Initiating Holders, file a Form S-3 registration statement under the Securities Act covering all Registrable Securities requested to be included in such registration by any other Holders, as specified by notice given by each such Holder to the Company within twenty (20) days of the date the Demand Notice is given, and in each case, subject to the limitations of Subsections 4.2(c) and 4.4.

(c) Notwithstanding the foregoing obligations, if the Company furnishes to Holders requesting a registration pursuant to this Subsection 4.2 a certificate signed by the Company’s chief executive officer stating that in the good faith judgment of the Company’s Board it would be materially detrimental to the Company and its stockholders for such registration statement to either become effective or remain effective for as long as such registration statement otherwise would be required to remain effective, because such action would (i) materially interfere with a significant acquisition, corporate

reorganization, or other similar transaction involving the Company; (ii) require premature disclosure of material information that the Company has a bona fide business purpose for preserving as confidential; or (iii) render the Company unable to comply with requirements under the 1933 Act or 1934 Act, then the Company shall have the right to defer taking action with respect to such filing, and any time periods with respect to filing or effectiveness thereof shall be tolled correspondingly, for a period of not more than ninety (90) days after the request of the Initiating Holders is given; provided, however, that the Company may not invoke this right more than twice in any twelve (12) month period; and provided further that the Company shall not register any securities for its own account or that of any other stockholder during such ninety (90) day period other than an Excluded Registration.

(d) The Company shall not be obligated to effect, or to take any action to effect, any registration pursuant to Subsection 4.2(a): (i) during the period that is sixty (60) days before the Company's good faith estimate of the date of filing of, and ending on a date that is one hundred eighty (180) days after the effective date of, a Company-initiated registration, provided that the Company is actively employing in good faith commercially reasonable efforts to cause such registration statement to become effective; (ii) after the Company has effected two registrations pursuant to Subsection 4.2(a); or (iii) if the Initiating Holders propose to dispose of shares of Registrable Securities that may be immediately registered on Form S-3 pursuant to a request made pursuant to Subsection 4.2(b). The Company shall not be obligated to effect, or to take any action to effect, any registration pursuant to Subsection 4.2(b): (i) during the period that is thirty (30) days before the Company's good faith estimate of the date of filing of, and ending on a date that is ninety (90) days after the effective date of, a Company-initiated registration, provided that the Company is actively employing in good faith commercially reasonable efforts to cause such registration statement to become effective; or (ii) if the Company has effected two registrations pursuant to Subsection 4.2(b) within the twelve (12) month period immediately preceding the date of such request. A registration shall not be counted as "effected" for purposes of this Subsection 4.2(d) until such time as the applicable registration statement has been declared effective by the SEC, unless the Initiating Holders withdraw their request for such registration, elect not to pay the registration expenses therefor, and forfeit their right to one demand registration statement pursuant to Subsection 2.6, in which case such withdrawn registration statement shall be counted as "effected" for purposes of this Subsection 4.2(d).

4.3 Company Registration. If the Company proposes to register (including, for this purpose, a registration effected by the Company for stockholders other than the Holders) any of its securities under the 1933 Act in connection with the public offering of such securities solely for cash (other than in an Excluded Registration), the Company shall, at such time, promptly give each Holder notice of such registration. Upon the request of each Holder given within twenty (20) days after such notice is given by the Company, the Company shall, subject to the provisions of Subsection 4.4, cause to be registered all of the Registrable Securities that each such Holder has requested to be included in such registration. The Company shall have the right to terminate or withdraw any registration initiated by it under this Subsection 4.3 before the effective date of such registration, whether or not any Holder has elected to include Registrable Securities in such registration. The expenses (other than Selling Expenses) of such withdrawn registration shall be borne by the Company in accordance with Subsection 4.7.

4.4 Underwriting Requirements.

(a) If, pursuant to Subsection 4.2, the Initiating Holders intend to distribute the Registrable Securities covered by their request by means of an underwriting, they shall so advise the Company as a part of their request made pursuant to Subsection 4.2, and the Company shall include such information in the Demand Notice. The underwriter(s) will be selected by the Company and shall be reasonably acceptable to a majority in interest of the Initiating Holders. In such event, the right of any Holder to include such Holder's Registrable Securities in such registration shall be conditioned upon such

Holder's participation in such underwriting and the inclusion of such Holder's Registrable Securities in the underwriting to the extent provided herein. All Holders proposing to distribute their securities through such underwriting shall (together with the Company as provided in Subsection 4.5(e)) enter into an underwriting agreement in customary form with the underwriter(s) selected for such underwriting. Notwithstanding any other provision of this Subsection 4.4, if the managing underwriter(s) advise(s) the Initiating Holders in writing that marketing factors require a limitation on the number of shares to be underwritten, then the Initiating Holders shall so advise all Holders of Registrable Securities that otherwise would be underwritten pursuant hereto, and the number of Registrable Securities that may be included in the underwriting shall be allocated among such Holders of Registrable Securities, including the Initiating Holders, in proportion (as nearly as practicable) to the number of Registrable Securities owned by each Holder or in such other proportion as shall mutually be agreed to by all such selling Holders; provided, however, that the number of Registrable Securities held by the Holders to be included in such underwriting shall not be reduced unless all other securities are first entirely excluded from the underwriting. To facilitate the allocation of shares in accordance with the above provisions, the Company or the underwriters may round the number of shares allocated to any Holder to the nearest one hundred (100) shares.

(b) In connection with any offering involving an underwriting of shares of the Company's capital stock pursuant to Subsection 4.3, the Company shall not be required to include any of the Holders' Registrable Securities in such underwriting unless the Holders accept the terms of the underwriting as agreed upon between the Company and its underwriters, and then only in such quantity as the underwriters in their sole discretion determine will not jeopardize the success of the offering by the Company. If the total number of securities, including Registrable Securities, requested by stockholders to be included in such offering exceeds the number of securities to be sold (other than by the Company) that the underwriters in their reasonable discretion determine is compatible with the success of the offering, then the Company shall be required to include in the offering only that number of such securities, including Registrable Securities, which the underwriters and the Company in their sole discretion determine will not jeopardize the success of the offering. If the underwriters determine that less than all of the Registrable Securities requested to be registered can be included in such offering, then the Registrable Securities that are included in such offering shall be allocated among the selling Holders in proportion (as nearly as practicable to) the number of Registrable Securities owned by each selling Holder or in such other proportions as shall mutually be agreed to by all such selling Holders. To facilitate the allocation of shares in accordance with the above provisions, the Company or the underwriters may round the number of shares allocated to any Holder to the nearest one hundred (100) shares. Notwithstanding the foregoing, in no event shall (i) the number of Registrable Securities included in the offering be reduced unless all other securities (other than securities to be sold by the Company) are first entirely excluded from the offering, or (ii) the number of Registrable Securities included in the offering be reduced below twenty percent (20%) of the total number of securities included in such offering, unless such offering is the IPO, in which case the selling Holders may be excluded further if the underwriters make the determination described above and no other stockholder's securities are included in such offering. For purposes of the provision in this Subsection 4.4(b) concerning apportionment, for any selling Holder that is a partnership, limited liability company, or corporation, the partners, members, retired partners, retired members, shareholders, and Affiliates of such Holder, or the estates and Immediate Family Members of any such partners, retired partners, members, and retired members and any trusts for the benefit of any of the foregoing Persons, shall be deemed to be a single "selling Holder," and any pro rata reduction with respect to such "selling Holder" shall be based upon the aggregate number of Registrable Securities owned by all Persons included in such "selling Holder," as defined in this sentence.

(c) For purposes of Subsection 4.2, a registration shall not be counted as "effected" if, as a result of an exercise of the underwriter's cutback provisions in Subsection 4.4(a), fewer than fifty percent (50%) of the total number of Registrable Securities that Holders have requested to be included in such registration statement are actually included.

4.5 **Obligations of the Company.** Whenever required under this Section 4 to effect the registration of any Registrable Securities, the Company shall, as expeditiously as reasonably possible:

(a) prepare and file with the SEC a registration statement with respect to such Registrable Securities and use its commercially reasonable efforts to cause such registration statement to become effective and, upon the request of the Holders of a majority of the Registrable Securities registered thereunder, keep such registration statement effective for a period of up to one hundred twenty (120) days or, if earlier, until the distribution contemplated in the registration statement has been completed; provided, however, that (i) such one hundred twenty (120) day period shall be extended for a period of time equal to the period the Holder refrains, at the request of an underwriter of Common Stock (or other securities) of the Company, from selling any securities included in such registration, and (ii) in the case of any registration of Registrable Securities on Form S-3 that are intended to be offered on a continuous or delayed basis, subject to compliance with applicable SEC rules, such one hundred twenty (120) day period shall be extended for up to sixty (60) days, if necessary, to keep the registration statement effective until all such Registrable Securities are sold;

(b) prepare and file with the SEC such amendments and supplements to such registration statement, and the prospectus used in connection with such registration statement, as may be necessary to comply with the Securities Act in order to enable the disposition of all securities covered by such registration statement;

(c) furnish to the selling Holders such numbers of copies of a prospectus, including a preliminary prospectus, as required by the 1933 Act, and such other documents as the Holders may reasonably request in order to facilitate their disposition of their Registrable Securities;

(d) use its commercially reasonable efforts to register and qualify the securities covered by such registration statement under such other securities or blue-sky laws of such jurisdictions as shall be reasonably requested by the selling Holders; provided that the Company shall not be required to qualify to do business or to file a general consent to service of process in any such states or jurisdictions, unless the Company is already subject to service in such jurisdiction and except as may be required by the 1933 Act;

(e) in the event of any underwritten public offering, enter into and perform its obligations under an underwriting agreement, in usual and customary form, with the underwriter(s) of such offering;

(f) use its commercially reasonable efforts to cause all such Registrable Securities covered by such registration statement to be listed on a national securities exchange or trading system and each securities exchange and trading system (if any) on which similar securities issued by the Company are then listed;

(g) provide a transfer agent and registrar for all Registrable Securities registered pursuant to this Agreement and provide a CUSIP number for all such Registrable Securities, in each case not later than the effective date of such registration;

(h) promptly make available for inspection by the selling Holders, any managing underwriter(s) participating in any disposition pursuant to such registration statement, and any attorney or accountant or other agent retained by any such underwriter or selected by the selling Holders, all financial

and other records, pertinent corporate documents, and properties of the Company, and cause the Company's officers, directors, employees, and independent accountants to supply all information reasonably requested by any such seller, underwriter, attorney, accountant, or agent, in each case, as necessary or advisable to verify the accuracy of the information in such registration statement and to conduct appropriate due diligence in connection therewith;

(i) notify each selling Holder, promptly after the Company receives notice thereof, of the time when such registration statement has been declared effective or a supplement to any prospectus forming a part of such registration statement has been filed; and

(j) after such registration statement becomes effective, notify each selling Holder of any request by the SEC that the Company amend or supplement such registration statement or prospectus.

In addition, the Company shall ensure that, at all times after any registration statement covering a public offering of securities of the Company under the Securities Act shall have become effective, its insider trading policy shall provide that the Company's directors may implement a trading program under Rule 10b5-1 of the Exchange Act.

4.6 Furnish Information. It shall be a condition precedent to the obligations of the Company to take any action pursuant to this Section 4 with respect to the Registrable Securities of any selling Holder that such Holder shall furnish to the Company such information regarding itself, the Registrable Securities held by it, and the intended method of disposition of such securities as is reasonably required to effect the registration of such Holder's Registrable Securities.

4.7 Expenses of Registration. All expenses (other than Selling Expenses) incurred in connection with registrations, filings, or qualifications pursuant to Section 4, including all registration, filing, and qualification fees; printers' and accounting fees; fees and disbursements of counsel for the Company; and the reasonable fees and disbursements (not to exceed \$30,000), of one counsel for the selling Holders ("**Selling Holder Counsel**"), shall be borne and paid by the Company; provided, however, that the Company shall not be required to pay for any expenses of any registration proceeding begun pursuant to Subsection 4.2 if the registration request is subsequently withdrawn at the request of the Holders of a majority of the Registrable Securities to be registered (in which case all selling Holders shall bear such expenses pro rata based upon the number of Registrable Securities that were to be included in the withdrawn registration), unless the Holders of a majority of the Registrable Securities agree to forfeit their right to one registration pursuant to Subsections 4.2(a) or 4.2(b), as the case may be; provided further that if, at the time of such withdrawal, the Holders shall have learned of a material adverse change in the condition, business, or prospects of the Company from that known to the Holders at the time of their request and have withdrawn the request with reasonable promptness after learning of such information then the Holders shall not be required to pay any of such expenses and shall not forfeit their right to one registration pursuant to Subsections 4.2(a) or 4.2(b). All Selling Expenses relating to Registrable Securities registered pursuant to this Section 4 shall be borne and paid by the Holders pro rata on the basis of the number of Registrable Securities registered on their behalf.

4.8 Delay of Registration. No Holder shall have any right to obtain or seek an injunction restraining or otherwise delaying any registration pursuant to this Agreement as the result of any controversy that might arise with respect to the interpretation or implementation of this Section 4.

4.9 Indemnification. If any Registrable Securities are included in a registration statement under this Section 4:

(a) To the extent permitted by law, the Company will indemnify and hold harmless each selling Holder, and the partners, members, officers, directors, and shareholders of each such Holder; legal counsel and accountants for each such Holder; any underwriter (as defined in the 1933 Act) for each such Holder; and each Person, if any, who controls such Holder or underwriter within the meaning of the 1933 Act or the 1934 Act, against any Damages, and the Company will pay to each such Holder, underwriter, controlling Person, or other aforementioned Person any legal or other expenses reasonably incurred thereby in connection with investigating or defending any claim or proceeding from which Damages may result, as such expenses are incurred; provided, however, that the indemnity agreement contained in this Subsection 4.9(a) shall not apply to amounts paid in settlement of any such claim or proceeding if such settlement is effected without the consent of the Company, which consent shall not be unreasonably withheld, nor shall the Company be liable for any Damages to the extent that they arise out of or are based upon actions or omissions made in reliance upon and in conformity with written information furnished by or on behalf of any such Holder, underwriter, controlling Person, or other aforementioned Person expressly for use in connection with such registration.

(b) To the extent permitted by law, each selling Holder, severally and not jointly, will indemnify and hold harmless the Company, and each of its directors, each of its officers who has signed the registration statement, each Person (if any), who controls the Company within the meaning of the 1933 Act, legal counsel and accountants for the Company, any underwriter (as defined in the 1933 Act), any other Holder selling securities in such registration statement, and any controlling Person of any such underwriter or other Holder, against any Damages, in each case only to the extent that such Damages arise out of or are based upon actions or omissions made in reliance upon and in conformity with written information furnished by or on behalf of such selling Holder expressly for use in connection with such registration; and each such selling Holder will pay to the Company and each other aforementioned Person any legal or other expenses reasonably incurred thereby in connection with investigating or defending any claim or proceeding from which Damages may result, as such expenses are incurred; provided, however, that the indemnity agreement contained in this Subsection 4.9(b) shall not apply to amounts paid in settlement of any such claim or proceeding if such settlement is effected without the consent of the Holder, which consent shall not be unreasonably withheld; and provided further that in no event shall the aggregate amounts payable by any Holder by way of indemnity or contribution under Subsections 4.9(b) and 4.9(d) exceed the proceeds from the offering received by such Holder (net of any Selling Expenses paid by such Holder), except in the case of fraud or willful misconduct by such Holder.

(c) Promptly after receipt by an indemnified party under this Subsection 4.9 of notice of the commencement of any action (including any governmental action) for which a party may be entitled to indemnification hereunder, such indemnified party will, if a claim in respect thereof is to be made against any indemnifying party under this Subsection 4.9, give the indemnifying party notice of the commencement thereof. The indemnifying party shall have the right to participate in such action and, to the extent the indemnifying party so desires, participate jointly with any other indemnifying party to which notice has been given, and to assume the defense thereof with counsel mutually satisfactory to the parties; provided, however, that an indemnified party (together with all other indemnified parties that may be represented without conflict by one counsel) shall have the right to retain one separate counsel, with the fees and expenses to be paid by the indemnifying party, if representation of such indemnified party by the counsel retained by the indemnifying party would be inappropriate due to actual or potential differing interests between such indemnified party and any other party represented by such counsel in such action. The failure to give notice to the indemnifying party within a reasonable time of the commencement of any such action shall relieve such indemnifying party of any liability to the indemnified party under this Subsection 4.9, to the extent that such failure materially prejudices the indemnifying party's ability to defend such action. The failure to give notice to the indemnifying party will not relieve it of any liability that it may have to any indemnified party otherwise than under this Subsection 4.9.

(d) To provide for just and equitable contribution to joint liability under the 1933 Act in any case in which either: (i) any party otherwise entitled to indemnification hereunder makes a claim for indemnification pursuant to this Subsection 4.9 but it is judicially determined (by the entry of a final judgment or decree by a court of competent jurisdiction and the expiration of time to appeal or the denial of the last right of appeal) that such indemnification may not be enforced in such case, notwithstanding the fact that this Subsection 4.9 provides for indemnification in such case, or (ii) contribution under the 1933 Act may be required on the part of any party hereto for which indemnification is provided under this Subsection 4.9, then, and in each such case, such parties will contribute to the aggregate losses, claims, damages, liabilities, or expenses to which they may be subject (after contribution from others) in such proportion as is appropriate to reflect the relative fault of each of the indemnifying party and the indemnified party in connection with the statements, omissions, or other actions that resulted in such loss, claim, damage, liability, or expense, as well as to reflect any other relevant equitable considerations. The relative fault of the indemnifying party and of the indemnified party shall be determined by reference to, among other things, whether the untrue or allegedly untrue statement of a material fact, or the omission or alleged omission of a material fact, relates to information supplied by the indemnifying party or by the indemnified party and the parties' relative intent, knowledge, access to information, and opportunity to correct or prevent such statement or omission; provided, however, that, in any such case (x) no Holder will be required to contribute any amount in excess of the public offering price of all such Registrable Securities offered and sold by such Holder pursuant to such registration statement, and (y) no Person guilty of fraudulent misrepresentation (within the meaning of Section 11(f) of the 1933 Act) will be entitled to contribution from any Person who was not guilty of such fraudulent misrepresentation; and provided further that in no event shall a Holder's liability pursuant to this Subsection 4.9(d), when combined with the amounts paid or payable by such Holder pursuant to Subsection 4.9(b), exceed the proceeds from the offering received by such Holder (net of any Selling Expenses paid by such Holder), except in the case of willful misconduct or fraud by such Holder.

(e) Notwithstanding the foregoing, to the extent that the provisions on indemnification and contribution contained in the underwriting agreement entered into in connection with the underwritten public offering are in conflict with the foregoing provisions, the provisions in the underwriting agreement shall control.

(f) Unless otherwise superseded by an underwriting agreement entered into in connection with the underwritten public offering, the obligations of the Company and Holders under this Subsection 4.9 shall survive the completion of any offering of Registrable Securities in a registration under this Section 4, and otherwise shall survive the termination of this Agreement.

4.10 Reports Under 1934 Act. With a view to making available to the Holders the benefits of SEC Rule 144 and any other rule or regulation of the SEC that may at any time permit a Holder to sell securities of the Company to the public without registration or pursuant to a registration on Form S-3, the Company shall:

(a) make and keep available adequate current public information, as those terms are understood and defined in SEC Rule 144, at all times after the effective date of the registration statement filed by the Company for the IPO;

(b) use commercially reasonable efforts to file with the SEC in a timely manner all reports and other documents required of the Company under the 1933 Act and the 1934 Act (at any time after the Company has become subject to such reporting requirements); and

(c) furnish to any Holder, so long as the Holder owns any Registrable Securities, forthwith upon request (i) to the extent accurate, a written statement by the Company that it has complied

with the reporting requirements of SEC Rule 144 (at any time after ninety (90) days after the effective date of the registration statement filed by the Company for the IPO), the 1933 Act, and the 1934 Act (at any time after the Company has become subject to such reporting requirements), or that it qualifies as a registrant whose securities may be resold pursuant to Form S-3 (at any time after the Company so qualifies); and (ii) such other information as may be reasonably requested in availing any Holder of any rule or regulation of the SEC that permits the selling of any such securities without registration (at any time after the Company has become subject to the reporting requirements under the 1934 Act) or pursuant to Form S-3 (at any time after the Company so qualifies to use such form).

4.11 Limitations on Subsequent Registration Rights. From and after the date of this Agreement, the Company shall not, without the prior written consent of the Holders of a majority of the Registrable Securities then outstanding, enter into any agreement with any holder or prospective holder of any securities of the Company that would provide to such holder the right to include securities in any registration on other than either a pro rata basis with respect to the Registrable Securities or on a subordinate basis after all Holders have had the opportunity to include in the registration and offering all shares of Registrable Securities that they wish to so include; provided that this limitation shall not apply to any additional Investor who becomes a party to this Agreement.

4.12 "Market Stand off" Agreement. Each Holder hereby agrees that it will not, without the prior written consent of the managing underwriter, during the period commencing on the date of the final prospectus relating to the registration by the Company of shares of its Common Stock or any other equity securities under the 1933 Act on a registration statement on Form S-1, and ending on the date specified by the Company and the managing underwriter (such period not to exceed one hundred eighty (180) days in the case of the IPO, or such other period as may be requested by the Company or an underwriter to accommodate regulatory restrictions on (1) the publication or other distribution of research reports, and (2) analyst recommendations and opinions, including, but not limited to, the restrictions contained in FINRA Rule 2711(f)(4) or NYSE Rule 472(f)(4), or any successor provisions or amendments thereto), (i) lend; offer; pledge; sell; contract to sell; sell any option or contract to purchase; purchase any option or contract to sell; grant any option, right, or warrant to purchase; or otherwise transfer or dispose of, directly or indirectly, any shares of Common Stock or any securities convertible into or exercisable or exchangeable (directly or indirectly) for Common Stock (whether such shares or any such securities are then owned by the Holder or are thereafter acquired) or (ii) enter into any swap or other arrangement that transfers to another, in whole or in part, any of the economic consequences of ownership of such securities, whether any such transaction described in clause (i) or (ii) above is to be settled by delivery of Common Stock or other securities, in cash, or otherwise. The foregoing provisions of this Subsection 4.12 shall apply only to the IPO, shall not apply to the sale of any shares to an underwriter pursuant to an underwriting agreement, or the transfer of any shares to any trust for the direct or indirect benefit of the Holder or the immediate family of the Holder, provided that the trustee of the trust agrees to be bound in writing by the restrictions set forth herein, and provided further that any such transfer shall not involve a disposition for value, and shall be applicable to the Holders only if all officers, directors and shareholders individually owning more than one percent (1%) of the Company's outstanding Common Stock (after giving effect to conversion into Common Stock of all outstanding Preferred Stock) are subject to the same restrictions. The underwriters in connection with such registration are intended third party beneficiaries of this Subsection 4.12 and shall have the right, power and authority to enforce the provisions hereof as though they were a party hereto. Each Holder further agrees to execute such agreements as may be reasonably requested by the underwriters in connection with such registration that are consistent with this Subsection 4.12 or that are necessary to give further effect thereto. Any discretionary waiver or termination of the restrictions of any or all of such agreements by the Company or the underwriters shall apply pro rata to all Holders subject to such agreements, based on the number of shares subject to such agreements.

4.13 Termination of Registration Rights. The right of any Holder to request registration or inclusion of Registrable Securities in any registration pursuant to Subsections 4.2 or 4.3 shall terminate upon the earliest to occur of:

- (a) the closing of a Liquidating Event, as such term is defined in the Charter; and
- (b) the fifth (5th) anniversary of the IPO.

Section 5. MISCELLANEOUS

5.1 Governing Law. This Agreement shall be governed by the laws of the State of Delaware without regard to conflicts of law provisions.

5.2 Successors and Assigns. Except as otherwise expressly provided herein, the provisions hereof shall inure to the benefit of, and be binding upon, the successors, assigns, heirs, executors and administrators of the parties hereto.

5.3 Entire Agreement. This Agreement constitutes the full and entire understanding and agreement among the parties with regard to the subjects hereof. Nothing in this Agreement, express or implied, is intended to confer upon any party, other than the parties hereto and their successors and assigns, any rights, remedies, obligations or liabilities under or by reason of this Agreement, except as expressly provided herein.

5.4 Severability. Any invalidity, illegality or limitation of the enforceability with respect to any stockholder of any one or more of the provisions of this Agreement, or any part thereof, whether arising by reason of the law of any such person's domicile or otherwise, shall in no way affect or impair the validity, legality or enforceability of this Agreement with respect to any other stockholder. In case any provision of this Agreement shall be invalid, illegal or unenforceable, it shall to the extent practicable, be modified so as to make it valid, legal and enforceable and to retain as nearly as practicable the intent of the parties, and the validity, legality, and enforceability of the remaining provisions shall not in any way be affected or impaired thereby.

5.5 Amendment and Waiver. Except as otherwise expressly provided herein, any term of this Agreement may be amended and the observance of any term of this Agreement may be waived (either generally or in a particular instance, either retroactively or prospectively and either for a specified period of time or indefinitely) with the written consent of the Company and the Investors, or their transferees, holding a majority of the shares of Investor Stock and voting together as a single group (treated as if converted at the conversion rate then in effect and including, for such purposes, shares of Common Stock into which any shares of Investor Stock shall have been converted that are held by a stockholder); provided, however, that no such amendment or waiver shall reduce the aforesaid percentage of Investor Stock and Common Stock issued upon conversion thereof, the holders of which are required to consent to any waiver or supplemental agreement, without the consent of the holders of all of such Investor Stock and Common Stock. Notwithstanding anything to the contrary in this Section 5.5, this Agreement may not be amended, modified or terminated and the observance of any term hereunder may not be waived with respect to any Investor without the written consent of such Investor unless such amendment, modification, termination or waiver applies to all Investors in the same fashion. Notwithstanding anything to the contrary in this Section 5.5, Section 3.1(f)(ii) of this Agreement may not be amended without the consent of Santen. Notwithstanding anything to the contrary in this Section 5.5, Investors purchasing Series B Preferred Stock of the Company pursuant to the Purchase Agreement may join as a party to this Agreement and Exhibit A hereto may be amended without the written consent of the Investors. Any amendment or waiver effected in accordance with this Section 5.5 shall be binding upon

the Company, each Investor and each transferee of the Company Stock. Upon the effectuation of each such amendment or waiver, the Company shall promptly give written notice thereof to the Investors who have not previously consented thereto in writing.

5.6 Delays or Omissions. No delay or omission to exercise any right, power or remedy accruing to the Company, the Investors, or any transferees upon any breach, default or noncompliance of the Investors or any transferee or the Company under this Agreement, shall impair any such right, power or remedy, nor shall it be construed to be a waiver of any such breach, default or noncompliance, or any acquiescence therein, or of any similar breach, default or noncompliance thereafter occurring. It is further agreed that any waiver, permit, consent or approval of any kind or character on the part of the Company or the Investors of any breach, default or noncompliance under this Agreement or any waiver on the Company's or the Investors' part of any provisions or conditions of this Agreement must be in writing and shall be effective only to the extent specifically set forth in such writing and that all remedies, either under this Agreement, by law, or otherwise afforded to the Company and the Investors, shall be cumulative and not alternative.

5.7 Notices, etc. Any notice required or permitted under this Agreement shall be given in writing and shall be deemed effectively given and received: (a) upon personal delivery to the party to be notified; (b) upon delivery by confirmed facsimile transmission if received by the recipient before 5:00 p.m. local time on a business day, and if not, then the next business day; (c) if to a U.S. resident, five (5) days after deposit with the United States Post Office, by registered or certified mail, postage prepaid; or (d) if to a U.S. resident, one (1) business day after deposit with a nationally recognized overnight courier service (or if to a non-U.S. resident, two (2) business days after deposit with an internationally recognized overnight courier service, specifying international priority delivery), and addressed:

(a) if to the Company, at:

Clearside Biomedical, Inc.
1220 Old Alpharetta Rd., Suite 300
Alpharetta, GA 30005
Attn: Chief Executive Officer

With a copy to:

Hutchison PLLC
3110 Edwards Mill Road, Suite 300
Raleigh, North Carolina 27612
Attn: William N. Wofford
Telephone: (919) 829-9600
Facsimile: (919) 829-9696

or at such other address as the Company shall have furnished to the Investors in writing;

(b) if to the Investors, at the addresses of such Investors specified on Exhibit A hereto, or at such other addresses as the Investors shall have furnished to the Company in writing; and

(c) if to a stockholder other than the Investors, at such stockholder's address as shall have been furnished to the Company in writing.

5.8 Attorneys' Fees. In the event that any suit or action is instituted under or in relation to this Agreement, including without limitation to enforce any provision in this Agreement, the prevailing

party in such dispute shall be entitled to recover from the losing party all fees, costs and expenses of enforcing any right of such prevailing party under or with respect to this Agreement, including without limitation, such reasonable fees and expenses of attorneys and accountants, which shall include, without limitation, all fees, costs and expenses of appeals.

5.9 Aggregation of Stock. All shares of Company Stock held or acquired by affiliated entities or persons or persons or entities under common management or control shall be aggregated together for the purpose of determining the availability of any rights under this Agreement.

5.10 Titles and Subtitles. The titles of the sections and subsections of this Agreement are for convenience of reference only and are not to be considered in construing this Agreement.

5.11 Counterparts. This Agreement may be executed in any number of counterparts, each of which shall be deemed an original, but all of which together shall constitute one instrument.

[The next page is the signature page.]

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

COMPANY:

Clearside Biomedical, Inc.

By: /s/ Daniel H. White

Name: Daniel H. White

Title: President and CEO

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

RMI Investments S.a.r.l.

By: /s/ Vladimir Gurdus

Name: Vladimir Gurdus

Title: Category A Director

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Daniel H. White
Daniel H. White

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

Daniel H. White IRA

By: /s/ Daniel H. White

Name: Daniel H. White

Title: Trustee

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

GRA Venture Fund, LLC

By: /s/ Diana Murphy

Name: Diana Murphy

Title: Managing Director

INVESTOR:

GRA Venture Fund (T.E.), LLC

By: /s/ Diana Murphy

Name: Diana Murphy

Title: Managing Director

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

Hatteras Venture Affiliates III, LP

By: Hatteras Venture Advisors III, LLC, its General Partner

By: /s/ Clay B. Thorp

Name: Clay B. Thorp

Title: Manager

INVESTOR:

Hatteras Venture Partners III, LP

By: Hatteras Venture Advisors III, LLC, its General Partner

By: /s/ Clay B. Thorp

Name: Clay B. Thorp

Title: Manager

INVESTOR:

Hatteras Venture Partners IV SBIC, LP

By: Hatteras Venture Advisors IV SBIC, LLC, its General Partner

By: /s/ Clay B. Thorp

Name: Clay B. Thorp

Title: Manager

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

H&M Holdings, LLC

By: /s/ Willam N. Wofford
Name: William N. Wofford
Title: Manager

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

KFBSF Private Equity Fund II, LP

By: /s/ David Stevens
Name: David Stevens
Title: Manager

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

MGC Venture Partners 2013, L.P.

By: /s/ Joe C. Cook III
Name: Joe C. Cook III
Title: Managing Member

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

Santen Pharmaceutical Co., Ltd.

By: /s/ Andrew Liu

Name: Andrew Liu

Title: Head of Corporate Planning

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Jerry Cagle
Jerry Cagle

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Charles A. Deignan
Charles A. Deignan

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Sanjay N. Rao
Sanjay N. Rao

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Tripti Prasad Thakur

Tripti Prasad Thakur

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Amar Bansal

Amar Bansal

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

Biren M. Patel Revocable Trust

By: /s/ Biren M. Patel
Name: Biren M. Patel
Title: Trustee/Owner

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Byron Smith

Byron Smith

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Joseph Cook Jr.

Joseph Cook, Jr.

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

Farview Management, LP

By: /s/ Joe C. Cook Jr.
Name: Joe C. Cook Jr.
Title: General Partner

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Jaya Hariprasad
Jaya Hariprasad

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

MMIC Investment Holdings, Inc.

By: /s/ Jason T. Sandner

Name: Jason T. Sandner

Title: CFO

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

The JWR, Jr. Family Trust

By: /s/ Donald G. Perry, Jr.

Name: Donald G. Perry, Jr.

Title: Trustee

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Joe W. Rogers, Jr.

Joe W. Rogers, Jr.

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Wyatt Thomas Johnson

Wyatt Thomas Johnson

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

The Starr Moore 2007 Revocable Trust

By: /s/ Frances G. Rogers

Name: Frances G. Rogers

Title: Trustee

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

SUNIL RAICHAND REVOCABLE TRUST DATED MAY 7, 2014 SUNIL RAICHAND TRUSTEE

By: /s/ Sunil Raichand
Name: Sunil Raichand
Title: Trustee

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Sidney Jain
Sidney Jain

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

By: /s/ Daniel Kiernan

Daniel Kiernan

IN WITNESS WHEREOF, this Second Amended and Restated Investor Rights Agreement has been duly executed and delivered by the parties as of the date first above written.

INVESTOR:

Georgia Research Alliance, Inc.

By: /s/ C. Michael Cassidy
Name: C. Michael Cassidy
Title: President

EXHIBIT A

SCHEDULE OF INVESTORS

<u>Investor</u>	<u>Address</u>
Bansal, Amar	1187 CleAnder Court Naperville, IL 60540
Biren M. Patel Revocable Trust	16 Baker Ln. Naperville, IL 60565
Cagle, Jerry	6309 Greenway Rd. Fort Worth, TX 76116
Cook, Jr., Joseph	1600 Division St Suite 580 Nashville, TN 37203 Attn: Joseph Cook, Jr.
Deignan, Charlie	1860 Broadwell Oaks Drive Alpharetta, GA 30004
Farview Management, LP	1600 Division St Suite 580 Nashville, TN 37203 Attn: Joseph Cook, Jr.
Georgia Research Alliance, Inc.	191 Peachtree Street, NE Suite 849 Atlanta, GA 30303 Attention: Ashley Cornelison and Diana Murphy
GRA Venture Fund (T.E.), LLC	c/o Georgia Research Alliance, Inc. 191 Peachtree Street, NE Suite 849 Atlanta, GA 30303 Attention: Ashley Cornelison and Diana Murphy
GRA Venture Fund, LLC	c/o Georgia Research Alliance, Inc. 191 Peachtree Street, NE Suite 849 Atlanta, GA 30303 Attention: Ashley Cornelison and Diana Murphy
H&M Holdings, LLC	3110 Edwards Mill Road, Suite 300 Raleigh, NC 27612 Attn: William N. Wofford
Hariprasad, Jaya	4626 South Woodlawn Avenue Chicago, IL 60653
Hatteras NC Fund	280 S. Mangum St., Suite 350 Durham, NC 27701 Attn: Clay Thorp
Hatteras Venture Affiliates III, LP	280 S. Mangum St., Suite 350 Durham, NC 27701 Attn: Clay Thorp

Hatteras Venture Partners III, LP

280 S. Mangum St., Suite 350
Durham, NC 27701
Attn: Clay Thorp

Hatteras Venture Partners IV SBIC, LP

280 S. Mangum St., Suite 350
Durham, NC 27701
Attn: Clay Thorp

Jain, Sidney

3386 Harvest Ridge Road
Geneva, IL 60134

Johnson, Wyatt Thomas

3290 Northside Parkway, Suite 375
Atlanta, GA 30327

KFBSF Private Equity Fund II, LP

University of North Carolina
Center for Entrepreneurial Studies
Kenan-Flagler Business School
CB# 3440 Kenan Center
Chapel Hill, NC 27599-3440
Attn: Tamala M. Grissett

Kiernan, Daniel F.

100 Banks Ave #1311
Rockville Centre, NY 11570

MGC Venture Partners 2013, L.P.

1600 Division St
Suite 580
Nashville, TN 37203

MMIC Investment Holdings, Inc.

Attn: Joseph Cook, Jr.
c/o Medical Mutual Insurance Company of North Carolina
700 Spring Forest Road Suite 400
Raleigh, North Carolina 27609
Attn: Jason Sandner, CFO

Rao, Sanjay

16 Kimberley Circle
Oak Brook, IL. 60523

RMI Investments S.a.r.l.

7, Rue Robert Stümper
L-2557 Luxembourg
Attn: Vladimir Gurdus

Rogers, Jr., Joe W.

3290 Northside Parkway, Suite 375
Atlanta, GA 30327

Santen Pharmaceutical Co., Ltd.

4-20, Ofukacho
Kita-ku3Osaka 530-8552, Japan
Attn: -General Manager, Finance & Accounting Group
Facsimile: 81-6-6321-7332

With a copy to:
Head of Global Business Development
Facsimile: 81-6-6359-3832

With a second copy to:
Santen, Inc.
2100 Powell Street, Suite 1600
Emeryville, California 94608
Attn: Yusuf Ali

Smith, Byron

1600 Division St
Suite 580
Nashville, TN 37203

Starr Moore 2007 Revocable Trust

Attn: Byron Smith
3290 Northside Parkway, Suite 375
Atlanta, GA 30327

Sunil Raichand Revocable Trust Dated May 7, 2014 Sunil Raichand Trustee

3012 Lincoln Road
Oak Brook IL 60523

Thakur, Tripti Prasad

1728 Waller Street
San Francisco, CA 94117

The JWR Jr Family Trust

3290 Northside Parkway, Suite 375
Atlanta, GA 30327

White, Daniel H.

212 Rivergate Dr.
Sawanee, GA 30024

White, Daniel H. (IRA)

Millennium Trust Company
c/o Daniel White
2001 Spring Road, Suite 700
Oak Brook, IL 60523

LEASE AGREEMENT

THIS LEASE AGREEMENT (the "Lease"), made and entered into as of March 14, 2012 (the "date of this Lease") by and between Clearside Biomedical, Inc., a Delaware corporation (hereinafter referred to as "Tenant"), and McDonald Ventures XI, LLC, a Georgia limited liability company (hereinafter referred to as "Landlord");

WITNESSETH

1. **PREMISES.** For and in consideration of the obligation of Tenant to pay rent as herein provided, and in consideration of the other terms, provisions and covenants hereof, Landlord hereby demises and leases to Tenant, and Tenant hereby leases from Landlord certain premises (Suite 300) being approximately 8,823 square feet of space (the "Premises") shown on Exhibit "B", within the building known as Windward Chase 300, (the "Building"), located at 1220 Old Alpharetta Road, Alpharetta, Forsyth County, Georgia, described on Exhibit "A" attached hereto and incorporated herein by reference, together with all rights, privileges, easements, and appurtenances belonging to or in any way pertaining to said Premises (hereinafter collectively the "Property").

TO HAVE AND TO HOLD the Premises for the Demised Term, as hereinafter defined.

2. **TERM.** The Term of this Lease (hereinafter referred to as the "Demised Term") shall be for a period commencing on the Commencement Date, as hereinafter defined, and ending thirty (30) full calendar months thereafter, unless sooner terminated as provided in this Lease; provided, however, that, in the event the Commencement Date is not the first day of a calendar month, the Demised Term shall extend for the remainder of the calendar month in which the Commencement Date occurs plus said number of months.

The "Commencement Date" shall be the earlier of: (i) April 1, 2012; or (ii) the date upon which the buildings and other improvements erected and to be erected upon the Premises shall have been substantially completed in accordance with the plans and specifications described on Exhibit "B" attached hereto and incorporated herein by reference (the "Plans"). In the event of delays caused by Tenant due to changes to the Plans or delayed approval of final construction drawings, the Commencement Date shall be the date upon which Landlord would have achieved substantial completion in the absence of Tenant's delays.

Landlord shall provide the improvements more particularly described on Exhibit "B" attached hereto. Landlord shall over perform the improvements described on the Plans, excluding those improvements to be installed by Tenant. Tenant shall have the right to enter the Premises during the final stages of construction for purposes of installing its phone and data cabling and security systems, if applicable. Upon substantial completion, Landlord shall notify Tenant in writing that the improvements are ready for occupancy. In the event that said buildings and other improvements have not in fact been substantially completed as aforesaid, Tenant shall notify Landlord in writing of its objections. Landlord shall have a reasonable time after delivery of such notice in which to take such corrective action as may be necessary, and shall notify Tenant in writing as soon as it deems such corrective action has been completed so that said buildings and other improvements are completed and ready for occupancy, subject to punch list items. Taking of possession by Tenant shall be deemed conclusively to establish that said buildings and other improvements have been completed in accordance with the plans and specifications and are in good and satisfactory condition. In the event of any dispute as to substantial completion of work performed or required to be performed by Landlord, or the date of substantial completion of such work, the certificate of the Landlord's contractor shall be conclusive. Tenant acknowledges that no representations as to the condition of the Premises have been made by Landlord, unless such are expressly set forth in this Lease. After the Commencement Date, Tenant shall, upon demand, execute and deliver to Landlord a letter of acceptance of delivery of the Premises.

3. **BASE RENT.**

A. Tenant agrees to pay Landlord rent for the Premises ("Base Rent"), in advance, without demand, deduction or set off, for the Demised Term at the rate of Six Thousand Six Hundred Eighteen and No/100s (\$6,618.00) per month. The first monthly installment of Base Rent shall be due and payable on the date hereof and a like monthly installment of Base Rent shall be due and payable on or before the first day of each calendar month thereafter during the Demised Term, except that the rent payment for any fractional calendar month at the commencement or end of the Demised Term shall be prorated on the basis of a thirty-day month.



B. Landlord and Tenant agree that the Base Rent set forth in Paragraph 3.A. above shall increase by three percent (3%) at the beginning of the thirteenth (13th) full calendar month of the Demised Term, which adjusted rent amount shall remain in effect for the next twelve (12) consecutive months and shall increase by three percent (3%) each twelve (12) months thereafter for the balance of the Demised Term; it being the express intention of the parties that in the event the Commencement Date is not the first day of a calendar month, the anniversary date of the rent adjustment hereunder shall be the first day of the first full calendar month. Whenever Base Rent is escalated under this Lease based on a percentage increase, the resulting escalated Base Rent amount shall be rounded up or down to the nearest whole dollar.

4. SECURITY DEPOSIT. Tenant agrees to deposit with Landlord on the date hereof, in addition to the rent specified in Paragraph 3, the sum off Six Thousand Six Hundred Eighteen and No/100s (\$6,618.00), which sum shall be held by Landlord, without obligation for interest (except as may be required by law), as security for the performance of Tenant's covenants and obligations under this Lease, it being expressly understood and agreed that such deposit is not an advance rent payment or a measure of Landlord's damages in case of Tenant's default. Upon the occurrence of any event of default by Tenant under this Lease, Landlord may (but without obligation to do so), from time to time, without prejudice to any other remedy provided herein or provided by law or in equity, use this security deposit to the extent necessary to make good any arrears of rent or other payments due Landlord hereunder, and any other damage, injury, expense or liability caused by such event of default. Tenant shall pay Landlord, on demand, the amount of the security deposit so applied in order to restore the security deposit to its original amount. This security deposit shall be deemed the property of Landlord, but any remaining balance of such deposit shall be returned by Landlord to Tenant within thirty (30) days after termination of this Lease and date that all of Tenant's obligations under this fulfilled.

5. USE. The Premises shall be used only for general office purposes, engineering to include (a) visual inspection – requires microscope and special light as well as computer and digital camera; (b) mechanical inspection – requires force-displacement station and video camera; (c) micro processing including polishing, sharpening; (d) assembly of devices and tools; (e) electrical testing – requires oscilloscopes and amplifiers; (f) light manufacturing; (g) research and development; and for the purpose of receiving, storing, shipping and selling (other than retail) products, materials and merchandise made and/or distributed by Tenant and for such other lawful purposes as may be incidental thereto. In the event, Tenant's uses should require any permits, variances, or other requirements as in order to occupy and use the Premises, Tenant shall be responsible for securing them at its sole cost and expense. Outside storage, including without limitation, trucks and other vehicles, is prohibited without Landlord's prior written consent. Tenant shall at its own cost and expense, obtain any and all other licenses and permits necessary for any such use. Tenant shall comply with all governmental laws, ordinances and regulations applicable to the use of the Premises, and shall promptly comply with all governmental orders and directives for the correction, prevention and abatement of nuisances in or upon, or connected with, the Premises, all at Tenant's sole expense. Tenant shall not permit any objectionable or unpleasant odors, smoke, dust, gas, noise or vibrations to emanate from the Premises, nor take any other action which would constitute a nuisance or would disturb or endanger any other tenants of the building or buildings in which the Premises are situated or unreasonably interfere with their use of their respective premises. Without Landlord's prior written consent, Tenant shall not receive, store or otherwise handle any product, material or merchandise which is explosive or highly flammable. Tenant will not permit the Premises to be used for any purpose or in any manner (including without limitation, any method of storage) which would render the insurance thereon void or the insurance risk more hazardous or cause the State Board of Insurance or other insurance authority to disallow any sprinkler credits. Tenant shall not use the Premises for the generation, storage, transportation or disposal of dangerous, toxic or hazardous materials, chemicals, wastes or similar substances.

6. TAXES.

A. Landlord agrees to pay before delinquency, all taxes, assessments and governmental charges of any kind and nature whatsoever (hereinafter collectively referred to as "taxes") lawfully levied or assessed against the Property. Tenant agrees to pay to Landlord, as additional rent, upon demand, the amount of Tenant's projected "proportionate share" of the taxes assessed against the Property. Tenant's "proportionate share",



as used in this Lease, shall mean a fraction, the numerator of which is the gross square footage contained in the Premises and the denominator of which is the gross square footage contained in the building or buildings located on the Property.

B. If at any time during the Term of this Lease, the present method of taxation shall be changed so that in lieu of the whole or any part of any taxes, assessments or governmental charges, levied, assessed or imposed on real estate and the improvements thereon, there shall be charged, levied, assessed or imposed on Landlord a capital levy or other tax directly on the rents received therefrom and/or a franchise tax, assessment, levy or charge measured by or based, in whole or in part, upon such rents for the present or any future building or buildings on the Property, then all such taxes, assessments, levies or charges, or the part thereof so measured or based, shall be deemed to be included within the term "taxes" for the purposes hereof.

C. The Landlord shall have the right (but no obligation) to employ a tax-consulting firm to attempt to assure a fair tax burden on the building or buildings on the Property within the applicable tax jurisdiction. Tenant shall pay to Landlord upon demand from time to time, as additional rent, the amount of Tenant's "proportionate share" (as defined in subparagraph 6.A herein) of the cost of such service, which is normally a contingency fee based on a percentage of tax savings generated by the tax firm.

D. Any payment to be made pursuant to this Paragraph 6 shall be prorated in the event any portion of the Demised Term is not within a full real estate tax year.

7. LANDLORD'S REPAIRS. Landlord, at its expense, shall maintain only the roof, foundation, and the structural soundness of the exterior walls of the Premises in good repair, reasonable wear and tear excepted. Tenant shall repair and pay for any damage caused by the negligence of Tenant, or Tenant's employees, agents or invitees, or caused by Tenant's default hereunder. The term "walls" as used herein shall not include windows, glass or plate glass, doors, special storefronts or office entries. Tenant shall immediately give Landlord written notice of defects or need for repairs, after which Landlord shall have reasonable opportunity to repair same or cure such defects. Landlord's liability with respect to any defects, repairs or maintenance for which Landlord is responsible under any of the provisions of this Lease shall be limited to the cost of such repairs or maintenance or the curing of such defects.

8. TENANT'S REPAIRS.

A. Tenant shall at its own cost and expense keep and maintain all parts of the Premises (except those for which Landlord is expressly responsible under the terms of this Lease) in good condition, promptly making all necessary repairs and replacements, including but not limited to, windows, glass and plate glass, doors, any special office entry, interior walls and finish work, floors and floor covering, heating and air condition systems, dock boards, truck doors, dock bumpers, plumbing work and fixtures, termite and pest extermination, and regular removal of trash and debris.

Tenant shall provide Landlord with prior notice of any repair to be undertaken by Tenant costing in excess of \$1,000 (in Tenant's reasonable estimation) and such other information as Landlord may reasonably request with respect to such repair, except such notice shall not be required if immediate repair is necessary for security or safety reasons.

B. Tenant shall not damage any wall or disturb the integrity and support provided by any wall and shall, at its sole cost and expense, promptly repair any damage or injury to any wall caused by Tenant or its employees, agents or invitees.

C. Tenant shall, at its own cost and expense, enter into a quarterly preventative maintenance/service contract with one of Landlord's preferred licensed HVAC contractors for servicing all heating and air conditioning systems and equipment, within the Premises over the term of the lease. In the event Landlord has not received a copy of Tenant's service contract described herein within 30 days after the commencement date of the Lease or Tenant fails to maintain the contract during the Term, Landlord may, at its option, enter into a service contract on behalf of Tenant, and Tenant shall reimburse Landlord, within 10 days notice from Landlord, for



the cost of such service contract. Tenant shall keep accurate and complete records of the performance of all scheduled maintenance under such contract and shall provide copies thereof to Landlord from time to time upon request by Landlord. The service contract must i) include all services suggested by the licensed contractor to keep the units in good repair, and ii) comply with any warranties (if applicable). Notwithstanding the foregoing, Landlord will warrant, for the first six (6) months of the Demised Term ("Warranty Period"), that all heating, cooling, ventilation, air conditioning systems are in good working order. Landlord will assume responsibility for repairs and replacements during the Warranty Period, by but not routine maintenance. During the Warranty Period and throughout the term of the Lease, Tenant will be responsible for any required maintenance to all systems in compliance with any warranties as recommended as part of the preventative maintenance agreement. Upon expiration of the Warranty Period, Tenant will be responsible for repairs, replacements, and maintenance of the HVAC units as required herein. Upon expiration of the Warranty Period, Landlord will agree to cap HVAC repairs or replacement at \$3,000.00 per unit per lease year for the balance of the Demised Term. Tenant will not be allowed to apply the cost of the regular maintenance contract towards the cap. Tenant shall notify Landlord in writing of any repairs or replacements needed that exceed the above referenced cap prior to such repairs and replacements being made. Landlord shall have the right to approve the contractor who is to perform such work or use a contractor of Landlord's choosing. The cap will be null and void if Tenant has not i) kept a quarterly maintenance contract with a licensed contractor, approved by Landlord, in place over the term of the lease to maintain the applicable equipment per the manufacturer's recommendations and ii) performed the maintenance recommended by the licensed contractor to keep the units in good repairs, iii) complied with any warranties (if applicable), (iv) notified Landlord in writing prior to such repairs and replacements being made, and (v) kept accurate and complete records of the performance of all scheduled maintenance under the service contract and provided copies thereof to Landlord from time to time upon request by Landlord.

9. COMMON AREA MAINTENANCE.

A. Tenant shall pay to Landlord as additional rent a common area operating and maintenance fee ("CAM") equal to Tenant's "proportionate share" (as defined in subparagraph 6.A. herein) of the cost and expense for the operation and maintenance of the common areas of the building and park in which the Premises are located, including, but not limited to, the mowing of grass, care of shrubs, general landscaping, common sewage line plumbing, maintenance of building, parking areas, entrances, driveways, and management thereof. If Tenant or any other particular tenant of the building can be clearly identified as being responsible for obstructions or stoppage of the common sanitary sewage line, then Tenant, if Tenant is responsible, or such other responsible tenant, shall pay the entire cost thereof, upon demand, as additional rent. Payment shall be made on the first day of each month based on the projected cost of such maintenance. At the end of each year, Landlord shall determine the actual costs of such maintenance. Any additional costs due from Tenant based on the actual costs shall be promptly paid by Tenant. Any savings will be credited against the following year's payments.

B. In addition to the CAM costs above, Tenant shall pay to Landlord a management fee equal to three percent (3%) of all rents paid in accordance with the Lease on a monthly basis, which payment shall be made on the first day of each month and shall be capped at three percent (3%) during the term of the Lease.

10. TENANT IMPROVEMENTS TO PREMISES. Notwithstanding Landlord's obligations under Exhibit "B", Tenant shall not make any alterations, additions or improvements to the Premises, exterior or interior (the "Tenant Alterations"), without the prior written consent of Landlord, except for unattached movable furniture and equipment which may be installed without drilling, cutting or otherwise defacing, damaging or overloading the Premises. In the event, Tenant desires to make an alterations to the Premises, Tenant shall submit to Landlord for approval: i) scope of work of the requested alterations, along with, ii) design drawings, and iii) the proposed contractors to perform as reasonably acceptable to Landlord. Upon Approval, Tenant shall prepare, if applicable, for review and approval for Landlord: i) all architectural, ii) mechanical, iii) electrical, iv) plumbing, v) and partition drawings. Once the work is completed, Tenant will provide Landlord with certificate of completion or equivalent documentation from governing authority and provide Landlord with all drawings. Without limiting the generality of the foregoing, Tenant acknowledges that floor striping of any kind constitutes a Tenant Alteration requiring the prior written approval of Landlord. The Tenant Alterations shall be deemed for all purposes a part of the real property of the Building immediately upon the installation thereof and shall remain on the Premises as



Landlord's property at the expiration or earlier termination of the term hereof without compensation to Tenant, unless Landlord elects by notice to Tenant to have Tenant remove such Tenant Alterations, in which event, Tenant, at its sole cost and expense, shall promptly remove such Tenant Alterations and restore the Premises to its condition prior to the installation of such Tenant Alterations, normal wear and tear excepted (it being agreed that any damages resulting from the removal of any Tenant Alterations shall not constitute normal wear and tear). Tenant may not use or penetrate the roof of the Premises for any purpose whatsoever without Landlord's prior written consent. All construction work done by Tenant in the Premises shall be performed in a good and workmanlike manner, in compliance with all governmental requirements, and at such times and in such manner as will cause a minimum of interference with other construction in progress and with the transaction of business in the building or buildings in which the Premises are located.

11. SIGNAGE. Landlord shall install, at its expense, Tenant's name and suite number on the building directory sign and Tenant may install its name and logo on the door to the Premises. Tenant shall not install any signs visible from outside the Premises except with the prior written consent of Landlord. Any permitted signs shall be maintained in compliance with applicable governmental rules and regulations governing such signs. Tenant shall be responsible to Landlord for any damage caused by the installation, use or maintenance of said signs. Tenant agrees, upon removal of said signs, to repair all damage (including discoloration) incident thereto.

12. RIGHT OF ENTRY INSPECTION. Landlord and Landlord's agents and representatives shall have the right to enter and inspect the Premises at any reasonable time during business hours, upon reasonable advance notice (except in cases of emergency), to ascertain the condition of the Premises, to make such repairs as may be required or permitted to be made by Landlord under the terms of this Lease, or to show the Premises to prospective purchasers or tenants. Landlord shall have the right to place or erect on the Property, but not directly in the Premises a suitable sign indicating the Premises are available for rent (during the last six (6) months of the Demised Term

Tenant shall give written notice to Landlord at least thirty (30) days prior to vacating the Premises and shall arrange to meet with Landlord for a joint inspection of the Premises prior to vacating. In the event of Tenant's failure to give such notice or arrange a joint inspection, Landlord's inspection at or after Tenant's vacating the Premises shall be conclusively deemed correct for purposes of determining Tenant's responsibility for repairs and restoration.

13. UTILITIES. Landlord agrees to provide at its cost water, electricity and telephone service connections into the Premises; but Tenant shall pay for all water, gas, heat, light, power, telephone, sewer, sprinkler charges and other utilities and services used in or from the Premises, together with any taxes, penalties, surcharges or the like pertaining thereto and any maintenance charges for utilities and shall furnish all electric light bulbs and tubes. If any such services are not separately metered to Tenant, Tenant shall pay a reasonable proportion as determined by Landlord of all charges jointly metered with other premises. Landlord shall in no event be liable for any interruption or failure of utility services on the Premises;.

14. ASSIGNMENT AND SUBLETTING.

A. Tenant shall not, directly or indirectly, have the right to assign this Lease or to sublet the whole or any part of the Premises without the prior written consent of Landlord, which shall not be unreasonably withheld, conditioned or delayed. Consent to any assignment or sublease shall not be deemed a waiver of the right of Landlord to approve or disapprove a further assignment or subletting. Notwithstanding any permitted assignment or subletting, Tenant shall at all times remain directly, primarily and fully responsible and liable for the payment of the rent herein specified and for compliance with all of its other obligations under the terms, provisions and covenants of this Lease. Upon the occurrence of an "event of default", as hereinafter defined, if the Premises or any part thereof are then assigned or sublet, Landlord, in addition to any other remedies herein provided, or provided by law, may at its option collect directly from such assignee or subtenant all rents becoming due to Tenant under such assignment or sublease and apply such rent against any sums due to Landlord from Tenant hereunder, and no such collection shall be construed to constitute a novation or a release of Tenant from the further performance of Tenant's obligations hereunder. For purposes of this Paragraph 14, each of the following events shall be deemed an assignment:

- (i) if Tenant is a partnership, a dissolution of the partnership or a change in ownership, legal or beneficial, of 50% or more of the partnership interests, whether by withdrawal or admission, voluntary or by operation of law;



- (ii) if Tenant is a corporation, the dissolution, consolidation or merger of Tenant or the sale or transfer of more than 50% of the voting shares of Tenant;
- (iii) distribution or sale of over 50% of the value of Tenant's assets (net of undistributed consideration received); or
- (iv) any other change of effective control of Tenant.

Notwithstanding (ii) & (iii) the above, Tenant shall have the right to assign this Lease in the event of sale or transfer of more than 50% of the voting shares or assets of Tenant without Landlord's consent, provided however, Tenant must provide Landlord written notice of its election to assign this Lease and execute those reasonable instruments recognizing such transfer. In the event of assignment, Tenant shall not be released from its obligations and shall remain liable for all obligations of the Lease under the Demised Term including any Renewal Options as provided for in Section 32.

B. In the event that Tenant assigns this Lease or sublets the Premises or any part thereof, as permitted herein, and at any time receives rent and/or other consideration which exceeds that which Tenant would at that time be obligated to pay Landlord, Tenant shall pay to Landlord 50% of the gross excess in such rent as such rent is received by Tenant and 50% of any other consideration received by Tenant from such assignee or subtenant. In addition, should Landlord agree to an assignment or sublease agreement, Tenant will pay to Landlord on demand a sum equal to all Landlord's costs, including reasonable attorney's fees, incurred in connection with such assignment or transfer not to exceed \$2,500.00. If an assignment or subletting is approved, Tenant shall be entitled to deduct from any excess proceeds described in this subparagraph 14.B. its reasonable expenses incurred in connection with such assignment or subletting.

15. INSURANCE.

A. Landlord agrees to maintain standard all risk insurance covering the building or buildings of which the Premises are a part in an amount not less than 80% (or such greater percentage as may be necessary to comply with the provisions of any co-insurance clauses of the policy) of the "replacement cost" thereof as such term is defined in the Replacement Cost Endorsement to be attached thereto, insuring against the perils of Fire, Lightning and Extended Coverage, such coverage and endorsements to be as defined, provided and limited in the standard bureau forms prescribed by the insurance regulatory authority for the State in which the Premises are situated for use by insurance companies admitted in such state for the writing of such insurance on risks located within such state. Subject to the provisions of this Paragraph 15, such insurance shall be for the sole benefit of Landlord and under its sole control.

Tenant agrees to pay to Landlord, as additional rent, the amount of Tenant's "proportionate share" of the cost of Landlord's insurance coverage on the building, all costs and premiums of all insurance, including but not limited to fire, casualty, business interruption, boiler and machinery (equipment breakdown) and commercial general liability insurance applicable to the building, the common areas and the operation thereof and Landlord's personal property used in connection therewith. Said payments shall be made to Landlord within ten (10) days after presentation to Tenant of Landlord's statement setting forth the amount due. Any payment to be made pursuant to this subparagraph 15.A. shall be prorated for any portion of the Demised Term that is not a full premium period under said insurance policy.

B. Tenant shall, throughout the Term of this Lease, at its cost and expense, provide and keep in force a commercial general liability insurance policy for bodily injury, property damage, and personal injury to a third party in the amount of not less than \$1,000,000 per occurrence and \$2,000,000 on an aggregate basis. Tenant shall be solely responsible for keeping insured, to the extent Tenant elects, its own personal property located on the Premises. All such insurances shall provide that the coverage there under shall be primary and noncontributing with



other coverage maintained by any additional insured affiliates and subsidiaries, including w/o limitation McDonald Development Company and McDonald Investments, Ltd. Tenant agrees to provide to Landlord a copy of the insurance policy endorsement confirming that the Landlord, McDonald Development Company, and McDonald Investments, Ltd. Area are additionally insured.

All insurance provided by Tenant as required by this subparagraph 15.B. shall name, as additional insured, Landlord and any mortgagees or deed to secure debt holders of the Premises, and be carried by such responsible companies and in such form satisfactory to Landlord.

Tenant agrees to deliver to Landlord on or before the Commencement Date the original policy of insurance required by this subparagraph 15.B. or certificate thereof and evidence of payment of premium. Prior to the expiration of each such policy, Tenant shall deliver to Landlord the new original policy or certificate for renewal insurance and evidence of payment of premium. In the event Landlord has not received (on or before the Commencement Date) evidence of Tenant's compliance with the insurance coverage's required by this subparagraph 15.B., Landlord may, but shall not be required to, secure coverage on behalf of Tenant in the amounts required by this subparagraph 15.B with companies satisfactory to Landlord. Tenant shall pay the costs of such coverage directly, or, if paid by Landlord, reimburse Landlord as additional rent, within ten (10) days of notice, for all costs incurred by Landlord in securing such coverage.

Tenant shall not violate or knowingly permit to be violated any of the conditions or provisions of any policy required by this subparagraph 15.B.

Each insurance policy (including renewal insurance) or certificates thereof issued by the insurer shall contain an agreement by the insurer that should any of the policies be cancelled before the expiration date thereof, notice will be delivered in accordance with the policy provisions, and in no event shall such policies be canceled by Tenant without Landlord's prior written consent.

C. Tenant and Landlord shall cooperate in connection with the collection of any insurance monies that may be due in the event of loss. Tenant and Landlord shall execute and deliver such proofs of loss and other instruments that may be required for the purpose of obtaining the recovery of any such insurance monies.

D. Any insurance provided for in this Paragraph 15 may be effected by a policy or policies of blanket insurance; provided, however, that the amount of the total insurance allocated to the Premises shall be such as to furnish in protection the equivalent of separate policies in the amount herein required, and provided further that in all other respects, any such policy or policies shall comply with the other provisions of this Lease. In any such case, it shall not be necessary to deliver the original of any such blanket policy, but rather a certified duplicate of such policy or certificate thereof.

16. DAMAGE OR DESTRUCTION; MUTUAL WAIVER OF SUBROGATION.

A. If the Premises should be damaged or destroyed by fire, tornado or other casualty, Tenant shall give written notice thereof to Landlord as soon as practicable.

B. If the Premises should be totally destroyed by fire, tornado or other casualty, or if they should be so damaged thereby that rebuilding or repairs cannot, in Landlord's estimation, be completed within one hundred fifty (150) days after the date upon which Landlord is notified by Tenant of such damage, this Lease shall terminate and the rent shall be abated during the unexpired portion of this Lease, effective upon the date of the occurrence of such damage.

C. If 80% or more of the gross square footage of the building in which the Premises are located is damaged or destroyed by any peril or casualty, Landlord shall have the right to terminate this Lease. Subject to the foregoing, if the Premises should be damaged or destroyed by any peril covered by the insurance to be provided by Landlord under subparagraph 15.A., but only to the extent that rebuilding or repairs can in Landlord's estimation be completed one hundred fifty (150) days after the date upon which the Landlord is notified by Tenant of such damage, this Lease shall not terminate, and Landlord shall at its sole cost and expense thereupon proceed



with reasonable diligence to rebuild and repair such buildings to substantially the condition in which they existed prior to such damage, except that Landlord shall not be required to rebuild, repair or replace any part of the partitions, fixtures, addition and other improvements which may have been placed in, on or about the Premises by Tenant. If the Premises are untenable in whole or in part following such damage, the rent payable hereunder during the period in which they are untenable shall be reduced to such extent Tenant is able to conduct its business. In the event that Landlord should fail to complete such repairs and rebuilding within one hundred fifty (150) days after the date upon which Landlord is notified by Tenant of such damage, Tenant may at its option terminate this Lease by delivering written notice of termination to Landlord as Tenant's exclusive remedy, whereupon all rights and obligations hereunder shall cease and terminate as of the date of such notice.

D. Notwithstanding anything herein to the contrary, in the event the holder of any indebtedness secured by a mortgage or deed to secure debt covering the Premises requires that the insurance proceeds be applied to such indebtedness, then Landlord shall have the right to terminate this Lease by delivering written notice of termination to Tenant within fifteen (15) days after such requirement is made by any such holder, whereupon all rights and obligations hereunder shall cease and terminate.

E. Each of Landlord and Tenant hereby releases the other from any loss or damage to property caused by fire or any other perils insured through or under them by way of subrogation or otherwise for any loss or damage to property caused by fire or any other perils insured in policies of insurance covering such property, even if such loss or damage shall have been caused by the fault or negligence of the other party, or anyone for whom such party may be responsible; provided, however, that this release shall be applicable and in force and effect only with respect to loss or damage occurring during such times as the releasor's policies shall contain a clause or endorsement to the effect that any such release shall not adversely affect or impair said policies or prejudice the right of the releasor to recover thereunder and then only to the extent of the insurance proceeds payable under such policies. Each of the Landlord and Tenant agrees that it will request its insurance carriers to include in its policies such a clause or endorsement. If extra cost shall be charged therefore, each party shall advise the other thereof and of the amount of the extra cost, and the other party, at its election, may pay the same, but shall not be obligated to do so.

17. LIABILITY; MUTUAL INDEMNIFICATION.

A. Landlord shall not be liable to Tenant or Tenant's employees, agents, invitees, patrons or visitors, or to any other person whomsoever, for any injury to person or damage to property on or about the Premises, resulting from and/or caused in part or whole by, the negligence or misconduct of Tenant, its employees, agents, invitees, patrons or visitors, or of any other person entering upon the Premises, or caused by the buildings and improvements located on the Premises becoming out of repair, use, generation, storage or disposal of toxic or hazardous materials or substances on or about the Premises, or caused by leakage of gas, oil, water or steam or by electricity emanating from the Premises, or due to any cause whatsoever. Tenant hereby covenants and agrees that it will at all times indemnify and hold safe and harmless the property, the Landlord (including without limitation, the trustee and beneficiaries if Landlord is a trust), Landlord's agents and employees from any loss, liability, claims, suits, costs, expenses, including without limitation, attorney's fees and damages, both real and alleged, arising out of any such damage or injury, except injury to persons or damage to property the sole cause of which is the negligence of Landlord or the failure of Landlord to repair any part of the Premises which Landlord is obligated to repair and maintain hereunder within a reasonable time after the receipt of written notice from Tenant of needed repairs.

B. Tenant shall not be liable to Landlord or Landlord's employees, agents, invitees, patrons or visitors, or to any other person whomsoever, for any injury to person or damage to property on or about the Building, resulting from and/or caused in part or whole by, the negligence or misconduct of Landlord, its employees, agents, invitees, patrons or visitors, or caused by the Building (other than the Premises) becoming out of repair, or caused by the use, generation, storage or disposal of toxic or hazardous materials or substances on or about the Building (other than the Premises), or caused by leakage of gas, oil, water or steam or by electricity emanating from the Building, or due to any cause whatsoever Subject to the waiver of subrogation provision in Section 16(E) of this Lease, Landlord hereby covenants and agrees that it will at all times indemnify and hold safe and harmless the Tenant (including without limitation, the trustee and beneficiaries if Tenant is a trust), Tenant's agents and employees from any loss, liability, claims, suits, costs, expenses, including without limitation, attorney's fees and damages arising out of any such damage or injury, except injury to persons or damage to property the cause of



which is the negligence of Tenant or the failure of Tenant to repair any part of the Premises which Tenant is obligated to repair and maintain hereunder within a reasonable time after the receipt of written notice from Landlord of needed repairs.

18. CONDEMNATION.

A. If 80% or more of the gross square footage of the building in which the Premises are located should be taken for public or quasi-public use under governmental law, ordinance or regulation by right of eminent domain, or by private purchase in lieu thereof, Landlord shall have the right to terminate this Lease and the rent shall be abated effective on the date of Landlord's election to so terminate. Additionally, if the whole or any substantial part of the Premises should be taken for any public or quasi-public use under governmental law, ordinance or regulation, or by right of eminent domain, or by private purchase in lieu thereof and the taking would prevent or materially interfere with the use of the Premises for the purpose for which that are being used, this Lease shall terminate and the rent shall be abated during the unexpired portion of this Lease, effective on the date of the physical taking of the Premises.

B. If part of the Premises shall be taken for any public or quasi-public use under any governmental law, ordinance or regulation, or by right of eminent domain, or by private purchase in lieu thereof, and this Lease is not terminated as provided in subparagraph 18A., this Lease shall not terminate but the rent payable hereunder during the unexpired portion of this Lease shall be reduced to such extent as may be fair and reasonable under all of the circumstances.

C. In the event of any such taking or private purchase in lieu thereof, Landlord and Tenant shall each be entitled to receive and retain such separate awards and/or portion of lump sum awards as may be allocated to their respective interests in any condemnation proceedings.

19. HOLDING OVER. Tenant shall, at the termination of this Lease by lapse of time or otherwise, deliver immediate possession of the Premises to Landlord. If Landlord agrees in writing that Tenant may hold over after the expiration or termination of this Lease, unless the parties hereto otherwise agree in writing on the terms of such holding over, the hold over tenancy shall be subject to termination by Landlord at any time upon not less than thirty (30) days advance written notice, or by Tenant at any time upon not less than thirty (30) days advance written notice, and all of the other terms and provisions of this Lease shall be applicable during that period, except that Tenant shall pay Landlord from time to time, upon demand, as rent for the period of any such hold over, an amount equal to one hundred fifty percent (150%) of the rent in effect on the termination date, computed on a daily basis for each day of the hold over period. No holding over by Tenant, whether with or without consent of Landlord, shall operate to extend this Lease except as otherwise expressly provided. The preceding provisions of this Paragraph 19 shall not be construed as Landlord's consent for Tenant to hold over.

20. QUIET ENJOYMENT. Landlord represents and warrants that it has full right and authority to enter into this Lease and that Tenant, upon paying the rent herein set forth and performing its other covenants agreements herein set forth, shall peaceably and quietly have, hold and enjoy the Premises for the Demised Term hereof, subject to the terms and provisions of this Lease.

21. EVENTS OF DEFAULT. The following events shall each be deemed an event of default by Tenant under this Lease:

A. Tenant shall fail to pay any installment of the rent herein required when due, or any payment with respect to taxes hereunder when due, or any other payment or reimbursement to Landlord required herein when due, and such failure shall continue for a period of five (5) days from the date of written notice from Landlord, provided, however, that Landlord shall not be required to give such notice more than twice in a twelve (12) month period.

B. Tenant shall become insolvent, or shall make a transfer to defraud creditors, or shall make an assignment for the benefit of creditors.



C. Tenant shall file a petition under any section or chapter of the National Bankruptcy Act, as amended, or under any similar law or statute of the United States or any State thereof, or Tenant shall be adjudged bankrupt or insolvent in proceedings filed against Tenant thereunder.

D. A receiver or trustee shall be appointed for all or substantially all of the assets of Tenant.

E. Tenant shall desert, abandon or vacate any substantial portion of the Premises.

F. Tenant shall fail to deliver an estoppel certificate to Landlord within the time frame set forth in Paragraph 24.

G. Tenant shall fail to comply with any term, provision or covenant of this Lease (other than as set forth in clauses A through F in this Paragraph 21), and shall not cure such failure within twenty (20) days after written notice thereof to Tenant; except however, in the event such cure is not available within twenty (20) days and Tenant continuously works to cure, then it shall have an extension until the compliance date.

22. REMEDIES. Upon the occurrence of any of the events of default described in Paragraph 21 hereof, Landlord shall have the option to pursue any one or more of the following remedies without notice or demand whatsoever:

A. Immediately or at any time thereafter terminate this Lease, and this Lease shall be deemed to have been terminated upon giving of notice of such termination pursuant to Paragraph 26 hereof. Upon such termination, Landlord shall have the right to recover from Tenant, as liquidated damages, the following:

(1) the worth, at the time of the award, of the unpaid rent that has been earned at the time of termination of this Lease: and

(2) the worth, at the time of the award, of the amount by which the unpaid rent that would have been earned after the date of termination of this Lease until the time of the award exceeds the net amount of rent that could have been reasonably obtained by Landlord using reasonable diligence to relet the Premises; and

(3) the worth, at the time of the award, of the amount by which the unpaid rent for the balance of the Lease term (as extended), if applicable) after the time of the award exceeds the net amount of rent that reasonably could have been obtained by Landlord using reasonable diligence to relet the Premises; and

(4) any other amount and court costs necessary to compensate Landlord for all detriment directly caused by Tenant's failure to perform its obligations under this Lease.

Any such payment shall constitute liquidated damages to Landlord, Landlord and Tenant acknowledging and agreeing that it is difficult to determine the actual damages Landlord would suffer by virtue of an event of default and that the agreed-upon liquidated damages are not punitive or a penalty and are just, fair, and reasonable, all in accordance with O.C.G.A. sec. 13-6-7.

The following words and phrases as used above in this Paragraph 22.A. shall have the following meanings:

(i) the "worth at the time of the award" as used in Paragraph 22.A. (1) and (2) shall be computed by allowing interest at the lesser of (A) eight percent (8%) per annum or (B) the maximum rate permitted by law.

(ii) the "worth at the time of the award" as used in Paragraph 22.A (3) shall be computed by discounting the amount at the discount rate of six percent (6%) per annum;

(iii) the term "time of the award" shall mean either the date upon which Tenant pays to Landlord the amount recoverable by Landlord as set forth above or the date of entry of any determination, order, or judgment of any court, whichever first occurs: and



(iv) the term "net amount of rent" means the gross rent payable under a lease reletting the Premises on market terms for the remaining Term of this Lease from and after the time of the award, less the amortized portion (such amortization being on a straightline basis over the term of the subject lease) attributable to the remaining Term of this Lease from and after the time of the award of brokerage commissions and fees, design fees, attorney's fees, improvement allowances, rent concessions, improvement costs, and other economic concessions and costs made or incurred in connection with a reletting of the Premises to a third party on market terms.

B. Enter upon and take possession of the Premises and expel or remove Tenant and any other person who may be occupying the Premises or any part thereof, by force if necessary (to the extent permitted by law), without being liable for prosecution or any claim for damages thereof, and relet the Premises and receive the rent therefore; and Tenant agrees to pay to the Landlord on demand any deficiency that may arise by reason of such reletting. In the event Landlord is successful in reletting the Premises at a rent in excess of that agreed to be paid by Tenant pursuant to the terms of this Lease, Landlord and Tenant each mutually agree that Tenant shall not be entitled, under any circumstances, to such excess rent, and Tenant does hereby specifically waive any claim to such excess rent.

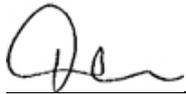
C. Enter upon the Premises, by force if necessary (to the extent permitted by law), without being liable for prosecution or any claim for damages therefore, and do whatever Tenant is obligated to do under the terms of this Lease; and Tenant agrees to reimburse Landlord on demand for any expenses which Landlord may incur in thus effecting compliance with Tenant's obligations under this Lease, and Tenant further agrees that Landlord shall not be liable for any damages resulting to the Tenant from such action, whether caused by the negligence of Landlord or otherwise.

D. Landlord shall have all other rights and remedies provided by law or in equity.

In the event Tenant fails to pay any installment of rent hereunder within five (5) days after such installment is due, to help defray the additional cost to Landlord for processing such late payments, Tenant shall pay to Landlord a late charge in an amount equal to five percent (5%) of such installment; and the failure to pay such amount within ten (10) days shall be an event of default hereunder. The provision for such late charge shall be in addition to all Landlord's other rights and remedies hereunder or at law and shall not be construed as liquidated damages or as limiting Landlord's remedies in any manner.

Pursuit of any of the foregoing remedies shall not preclude pursuit of any of the other remedies herein provided or any other remedies provided by law, nor shall pursuit of any remedy herein provided constitute a forfeiture or waiver of any rent due to Landlord hereunder or any damages accruing to Landlord by reason of the violation of any of the terms, provisions and covenants herein contained. Notwithstanding the foregoing, if Landlord elects to terminate this Lease pursuant to Paragraph 22.A. hereof, Landlord's remedies thereunder constituting liquidated damages shall be Landlord's exclusive remedies hereunder respecting Landlord's breach of contract damages resulting from the termination of this Lease. No act or thing done by the Landlord or its agents during the Demised Term shall be deemed a termination of this Lease or an acceptance of the surrender of the Premises, and no agreement to terminate this Lease or accept a surrender of the Premises shall be valid unless in writing signed by Landlord. No waiver by Landlord of any violation or breach of any of the terms, provisions and covenants herein contained shall be deemed or construed to constitute a waiver of any other violation or breach of any of the terms, provisions and covenants herein contained. Landlord's acceptance of the payment of rent or other payments hereunder after the occurrence of an event of default shall not be construed as a waiver of such default, unless Landlord so notifies Tenant in writing. Forbearance by Landlord to enforce one or more of the remedies herein provided upon an event of default shall not be deemed or construed to constitute a waiver of such default or of Landlord's right to enforce any such remedies with respect to such default or any subsequent default. If, on account of any breach or default by Tenant in Tenant's obligations under the terms and conditions of this Lease, it shall become necessary or appropriate for Landlord to employ or consult with an attorney concerning or to enforce or defend any of Landlord's rights or remedies hereunder, Tenant agrees to pay any reasonable attorney's fees so incurred.

Tenant agrees to indemnify and hold Landlord harmless from any and all losses, costs, expenses (including, without limitation, attorney's fees), liabilities, causes of action, suits, claims, and damages arising out of, or in connection with any violation or breach of, or failure of Tenant to fully and completely observe, satisfy, perform and comply with, the terms and conditions of this Lease.



23. LANDLORD'S LIEN. [Intentionally Deleted]

24. MORTGAGES, GROUND LEASES AND ESTOPPEL CERTIFICATES.

A. Tenant hereby agrees and accepts that this Lease is and shall be subject and subordinate to any mortgage(s) and/or deeds to secure debt (collectively referred to as the "Mortgage") now or any time hereafter constituting a lien or charge upon the Premises or the improvements situated thereon; provided, however, that if the holder of any such Mortgage elects to have Tenant's interest in this Lease superior to any such instrument, then by notice to Tenant from such holder, this Lease shall be deemed superior to such lien, whether this Lease was executed before or after said Mortgage. If the holder of the Mortgage or any successor in interest shall succeed to the rights of Landlord under this Lease through a foreclosure sale or a sale in lieu of foreclosure, Tenant will attorn to and recognize such successor-landlord as Tenant's landlord and such successor-landlord shall accept such attornment and recognize Tenant's rights of possession and use of the Premises in accordance with the provisions of this Lease. At the request of any holder of a Mortgage, Tenant, Landlord and such holder shall enter into a subordination, non-disturbance and attornment agreement reasonably acceptable to Landlord, Tenant and such holder, but in any event substantially consistent with the terms of this Lease. Tenant shall at any time hereafter on demand execute any instruments, releases or other documents which may be required by the holder of the Mortgage for the purpose of subjecting and subordinating this Lease to the lien of any such Mortgage.

If, in connection with obtaining financing or refinancing for the Premises, or a sale of the Premises, any lender or purchaser shall request reasonable modifications in this Lease as a condition to such financing or purchase, Tenant will not unreasonably withhold or delay or defer its consent thereto, provided that such modifications do not increase the obligations of Tenant hereunder or materially and adversely affect Tenant's rights hereunder.

B. Tenant hereby further agrees and accepts that this Lease is and shall be subject and subordinate to any ground lease now or at any time hereafter affecting the Premises. Tenant shall at any time hereafter on demand execute any instruments, releases or other documents which may be required by the ground lessor of any ground lease affecting the Premises for the purpose of subjecting and subordinating this Lease to any such ground lease.

In the event any ground lessor of a ground lease affecting the Premises requests reasonable modifications in this Lease, Tenant will not unreasonably withhold or delay or defer its consent thereto, provided that such modifications do not increase the obligations of Tenant hereunder or materially and adversely affect Tenant's rights hereunder.

C. At any time and from time to time designated by Landlord, Tenant will execute, acknowledge and deliver to Landlord, within ten (10) days after receipt of a request from Landlord, a certificate certifying (a) that this Lease is unmodified and in full force and effect (or, if there have been modifications, that this Lease is in full force and effect, as modified, and stating the date and nature of each modification), (b) the date, if any, to which rent and other sums payable hereunder have been paid, (c) that no notice has been received by Tenant of any default which has not been cured, except as to defaults specified in said certificate, and (d) such other matters as may be requested by Landlord. Any such certificate may be relied upon by any existing or prospective purchaser, investor, ground lessor, mortgagee or holder of any deed to secure debt on the Building or any part thereof.

25. MECHANIC'S LIENS. Tenant shall have no authority, express or implied, to create or place any lien or encumbrance of any kind or nature whatever upon or in any matter to bind, the interest of Landlord in the Premises or to charge the rent payable hereunder for any claim in favor of any person dealing with Tenant, including those who may furnish materials or perform labor for any construction or repairs and each such claim shall affect and each such lien shall attach to, if at all, only the leasehold interest granted to Tenant by this instrument. Tenant



covenants and agrees that it will pay or cause to be paid all sums legally due and payable by it on account of any labor performed or materials furnished in connection with any work performed on the Premises on which any lien is or can be validly and legally asserted against its leasehold interest in the Premises or the improvements thereon and that it will save and hold Landlord harmless from any and all loss, cost or expense based on or arising out of asserted claims or liens against the leasehold estate or against the right, title and interest of the Landlord in the Premises or under the terms of the lease.

26. **NOTICES.** Any notice, demands, payments or other communications required or permitted to be delivered under this Lease shall be given by personal delivery, by deposit with a courier service that provides next-business-day service, or by deposit in the United States Mail, postage prepaid, Certified or Registered Mail, addressed to the parties hereto at the respective addresses set out below, or at such other address as they have theretofore specified by written notice delivered in accordance herewith:

LANDLORD:

TENANT:

McDonald Ventures XI, LLC
c/o McDonald Development Company
3715 Northside Parkway, Bldg 200, Suite 700
Atlanta, Georgia 30327
Attn: John R. McDonald

Clearside Biomedical, Inc.
1220 Old Alpharetta Road
Suite 300
Alpharetta, Georgia 30005

All notices shall be deemed given upon personal delivery, or upon deposit with a courier service that provides next-business-day service, or deposit in the United States Mail, postage prepaid, Certified or Registered Mail, except as otherwise specifically provided in this Lease and except as to the payments of rent to Landlord which shall be effective upon receipt by Landlord.

If and when included within the term "Landlord", as used in this instrument, there are more than one person, firm or corporation, all shall jointly arrange among themselves for their joint execution of such a notice specifying some individual at some specific address for the receipt of notices and payments to Landlord; if and when included within the term "Tenant", as used in this instrument, there are more than one person, firm or corporation, all shall jointly arrange among themselves for their joint execution of such a notice specifying some individual at some specific address within the continental United States for the receipt of notices and payments to Tenant. All parties within the terms "Landlord" and "Tenant", respectively, shall be bound by notices given in accordance with the provisions of this Paragraph 26 to the same effect as if each had received such notice.

27. **RESTRICTIVE COVENANTS.** Tenant acknowledges that this Lease shall be subject and subordinate at all times to the Declaration of Easements, recorded, or to be recorded, in County, Georgia Records, as the same may be amended from time to time (hereinafter referred to as the "Declaration"), which affects the Premises. Tenant agrees to comply with all of the terms and provisions of the Declaration, and not suffer or cause any act by Tenant or any of its employees, agents or invitees, which would violate the Declaration.

28. **REAL ESTATE BROKER.** Tenant represents and warrants that the Tenant has dealt with no broker, agent or finder in connection with this Lease other than Lavista Associates, Inc. ("Broker"), which broker is acting on behalf of Tenant and shall be paid a commission by Landlord pursuant to a separate agreement, and insofar as the Tenant knows, no other brokers, agent or finder negotiated this Lease or is entitled to any commission or fee in connection herewith. Tenant agrees to indemnify, defend and hold Landlord free and harmless from and against all claims for broker's or agent's commissions or finder's fees by any person claiming to have been retained by Tenant in connection with this transaction, or any other losses, costs, expenses (including, without limitation, attorney's fees), liabilities, damages, causes of actions or suits arising out of the alleged employment or use of a broker, agent or finder by Tenant.

29. LIMITATIONS ON LANDLORD'S LIABILITY LANDLORD'S LIABILITY FOR DAMAGES OR BREACH OR NONPERFORMANCE BY LANDLORD, OR ARISING OUT OF THE SUBJECT MATTER OF THIS LEASE OR THE RELATIONSHIP CREATED HEREBY, SHALL BE LIMITED TO, AND COLLECTIBLE ONLY OUT OF, LANDLORD'S INTEREST IN THE PREMISES AND NO PERSONAL LIABILITY IS ASSUMED BY, OR SHALL AT ANY TIME BE ASSERTED AGAINST, LANDLORD OR ITS AFFILIATED CORPORATIONS, ITS AND THEIR PARTNERS, VENTURERS, DIRECTORS, SHAREHOLDERS, OFFICERS, AGENTS, SERVANTS AND EMPLOYEES, OR ANY OF ITS OR THEIR SUCCESSORS OR ASSIGNS; ALL SUCH LIABILITY, IF ANY, BEING EXPRESSLY WAIVED AND RELEASED BY TENANT. IF LANDLORD, IN VIOLATION OF THE TERMS OF THIS LEASE OR THE PROVISIONS OF LAW, WITHHOLDS, DENIES OR DELAYS ANY CONSENT WHICH TENANT IS REQUIRED TO OBTAIN HEREUNDER, TENANT MAY SEEK SPECIFIC PERFORMANCE BUT SHALL NOT BE ENTITLED TO DAMAGES THEREFORE. LANDLORD'S REVIEW, SUPERVISION, COMMENTING ON OR APPROVAL OF ANY ASPECT OF WORK TO BE DONE BY OR FOR TENANT IS SOLELY FOR LANDLORD'S PROTECTION AND, EXCEPT AS EXPRESSLY PROVIDED, CREATES NO WARRANTIES OR DUTIES TO TENANT OR TO THIRD PARTIES. LANDLORD SHALL NOT BE LIABLE IN ANY EVENT FOR INDIRECT, CONSEQUENTIAL, OR SPECULATIVE DAMAGES SUCH AS BUSINESS LOSS.

30. RIGHT TO RELOCATE. Anytime after the Twenty-fourth (24) month following the Commencement Date, or anytime after the after sixtieth (60th) month following the Commencement Date provided Tenant has exercised its Renewal Option under Section 32 (Renewal Option), Landlord shall have the option to relocate the Tenant to alternative space ("Relocated Space") in the Building or Park at any time during the Term of the Lease, which alternative space shall be substantially the same size as the Premises. In the event, Landlord is interested in relocating Tenant to a purposed Relocated Space, Landlord shall send written notice to Tenant with the location of the Relocated Space and Tenant shall have five (5) business days to decide in writing, to Landlord, if the Relocation Space is acceptable or not to Tenant. If Tenant does not send written notice to Landlord of its decision to accept or reject the purposed Relocated Space, Tenant shall have deemed to have accepted the Relocated Space. If Tenant responds in writing, that the Relocated Space is not acceptable, Landlord, at it's option, shall have five (5) business days to decide whether or not to terminate the Lease with written notice to Tenant. If Landlord elects to terminate the Lease, Landlord shall provide Tenant an \$8,000 reimbursement allowance for reasonable and actual out of pocket relocation expenses of for its actual move to another facility. If Tenant elects to accept that the Relocated Space, Landlord shall, at its option, give not less than sixty (60) days' prior written notice, to Tenant, of such relocation, which notice shall include the date on which the Tenant shall be required to relocate or move and a description of the space to which Tenant will be relocated. Landlord shall pay all reasonable and actual out-of-pocket costs and expenses of relocating Tenant. If the Premises have already been improved for Tenant's occupancy, Landlord shall improve the new premises at its expense such that they are in substantially the same condition as the Premises. In the event of such relocation, such alternative space shall, for all purposes, be deemed the Premises hereunder and this Lease shall continue in full force and effect without any change in the other terms or conditions hereof; provided however, the Base Rent and other expenses paid hereunder shall be adjusted such that the Base Rent and other expenses paid by Tenant for the new premises shall be the same on a per square foot basis as they are for the Premises.

31. MISCELLANEOUS.

A. Words of any gender used in this Lease shall be held and construed to include any other gender, and words in the singular number shall be held to include the plural, unless the context otherwise requires.

B. The terms, provisions and covenants and conditions contained in this Lease shall apply to, inure to the benefit of, and be binding upon, the parties hereto and upon their respective heirs, legal representatives, successors and permitted assigns, except as otherwise herein expressly provided. Landlord shall have the right to assign any of its rights and obligations under this Lease.

C. Tenant agrees to furnish to the Landlord promptly upon demand, a corporate resolution, proof of due authorization by partners, or other appropriate documentation evidencing the due authorization of such Tenant to enter into this Lease.



D. The captions inserted in this Lease are for convenience only and in no way define, limit or otherwise describe the scope or intent of this Lease, or any provision hereof, or in any way affect the interpretation of this Lease.

E. Time is of the essence of this Lease.

F. Tenant agrees from time to time within ten (10) days after request of Landlord, to deliver to Landlord, or Landlord's designee, an estoppel certificate stating whether this Lease is in full force and effect, the date to which rent has been paid, the unexpired Term of this Lease and such other matters pertaining to this Lease as may be requested by Landlord. It is understood and agreed that Tenant's obligation to furnish such estoppel certificates in a timely fashion is a material inducement for Landlord's execution of this Lease.

G. This Lease may not be altered, changed, or amended except by an instrument in writing signed by both parties hereto.

H. All obligations of Tenant hereunder not fully performed as of the expiration or earlier termination of the Demised Term shall survive the expiration or earlier termination of the Demised Term, including without limitation, all payment obligations with respect to taxes and insurance and all obligations concerning the condition of the Premises. Upon the expiration or earlier termination of the Demised Term, and prior to Tenant vacating the Premises, Tenant shall pay to Landlord any amount reasonably estimated by Landlord as necessary to put the Premises, including without limitation, all heating and air conditioning systems and equipment therein, in good condition and repair. Tenant shall also, prior to vacating the Premises, pay to Landlord the amount, as estimated by Landlord, of Tenant's obligation hereunder for real estate taxes and insurance premiums for the year in which the lease expires or terminates. All such amounts shall be used and held by Landlord for payment of such obligations of Tenant hereunder, with Tenant being liable for any additional costs therefore upon demand by Landlord, or being liable for any additional costs therefore upon demand by Landlord, or with any excess to be returned to Tenant after all such obligations have been determined and satisfied, as the case may be. Any security deposit held by Landlord shall be credited against the amount payable by Tenant under this Paragraph 31.H.

I. If any clause, sentence, paragraph or provision of this Lease is illegal, invalid or unenforceable under present or future laws effective during the Term of this Lease, then and in that event it is the intention of the parties hereto that the remainder of this Lease shall not be affected thereby, and it is also the intention of the parties to this Lease that in lieu of each clause, sentence, paragraph or provision of this Lease that is illegal, invalid or unenforceable, there be added as a part of this Lease contract a clause, sentence, paragraph or provision as similar in terms to such illegal, invalid or unenforceable clause or provision as may be possible and be legal, valid and enforceable.

J. Provided Landlord is the prevailing party, Tenant agrees to pay any and all attorneys' fees and expenses Landlord incurs in enforcing any of the obligations of Tenant under this Lease, or in any litigation or negotiation in which Landlord shall, by virtue of this Lease or Landlord's ownership of the Premises, become involved in, through or on account of this Lease. Provided Tenant is the prevailing party, Landlord agrees to pay any and all attorneys' fees and expenses Tenant incurs in enforcing any of the obligations of Landlord under this Lease.

K. This Lease shall create the relationship of landlord and tenant between Landlord and Tenant; no estate shall pass out of Landlord; Tenant has only a usufruct, not subject to levy and sale.

L. Neither this Lease, nor any memorandum, affidavit or other writing with respect thereto, shall be recorded by Tenant or by anyone acting through, under or on behalf of Tenant, and the recording thereof in violation of this provision shall make this Lease voidable at Landlord's election.

M. Because the Premises are on the open market and are presently being shown, this Lease shall be treated as an offer with the Premises being subject to prior lease and such offer subject to withdrawal or non-acceptance by Landlord or to other use of the Premises without notice, and this Lease shall not be valid or binding unless and until accepted by Landlord in writing and a fully executed copy delivered to both parties hereto.



N. All references in this Lease to "the date hereof" or similar references shall be deemed to refer to the last date, in point of time, on which all parties hereto have executed this Lease.

O. This Lease may be executed in counterparts, each of which shall be deemed an original, and all of which shall constitute one and the same lease agreement

P. Landlord and Tenant each hereby covenant and agree that this lease, including the exhibits hereto, sets forth all of the promises, covenants, agreements, conditions and understandings between them with respect to the subject matter hereof and supersedes all prior and contemporaneous agreements and understandings relating to such subject matter except as otherwise set forth herein. In entering into this lease, neither party has relied upon any representation, warranty, covenant or other inducement not expressly set forth in this lease.

32. RENEWAL OPTION. Provided that Tenant is not in default at the time of exercise of this option or at the expiration of the initial Demised Term, Tenant shall have the right to renew the term of the lease for a single, thirty (30) month period, upon not less than six (6) months and no more than eighteen (18) months written notice to Landlord. The base rental rate shall increase by three percent (3%) over the then escalated Base Rent at the beginning of the Renewal Period and shall increase by three percent (3%) every twelve (12) months during the Renewal Period. Additionally, Landlord as part of the Renewal Period will provide Tenant with \$2.00 per square foot allowance to make improvements to the Premises.

(Signatures on the following Page)

A handwritten signature in black ink, appearing to be a stylized 'D' followed by a flourish.

IN WITNESS WHEREOF, the parties have executed this Lease with intent to be bound hereby as of the day and year indicated above each signature block, respectively, but effective as of the day and year first above written.

EXECUTED BY LANDLORD, this 15 day of March, 2012.

LANDLORD

McDonald Ventures XI, LLC, a Georgia limited liability company

By: McDonald Industrial XI, LLC, a Georgia limited liability company, its Manager

By: McDonald Development Company, a Georgia corporation, its Manager

By: /s/ John R. McDonald
John R. McDonald, President

EXECUTED BY TENANT, this 14 day of March, 2012.

TENANT

Clearside Biomedical, Inc., a Delaware corporation

By: /s/ Daniel H. White
Title: Daniel H. White
President & CEO

Exhibit A

Site 300

ALL THAT TRACT OR PARCEL OF LAND lying and being in Land Lots 961, 984, 985, and 1032 of the 2nd District, 1st Section, Forsyth County, Georgia and being more particularly described as follows:

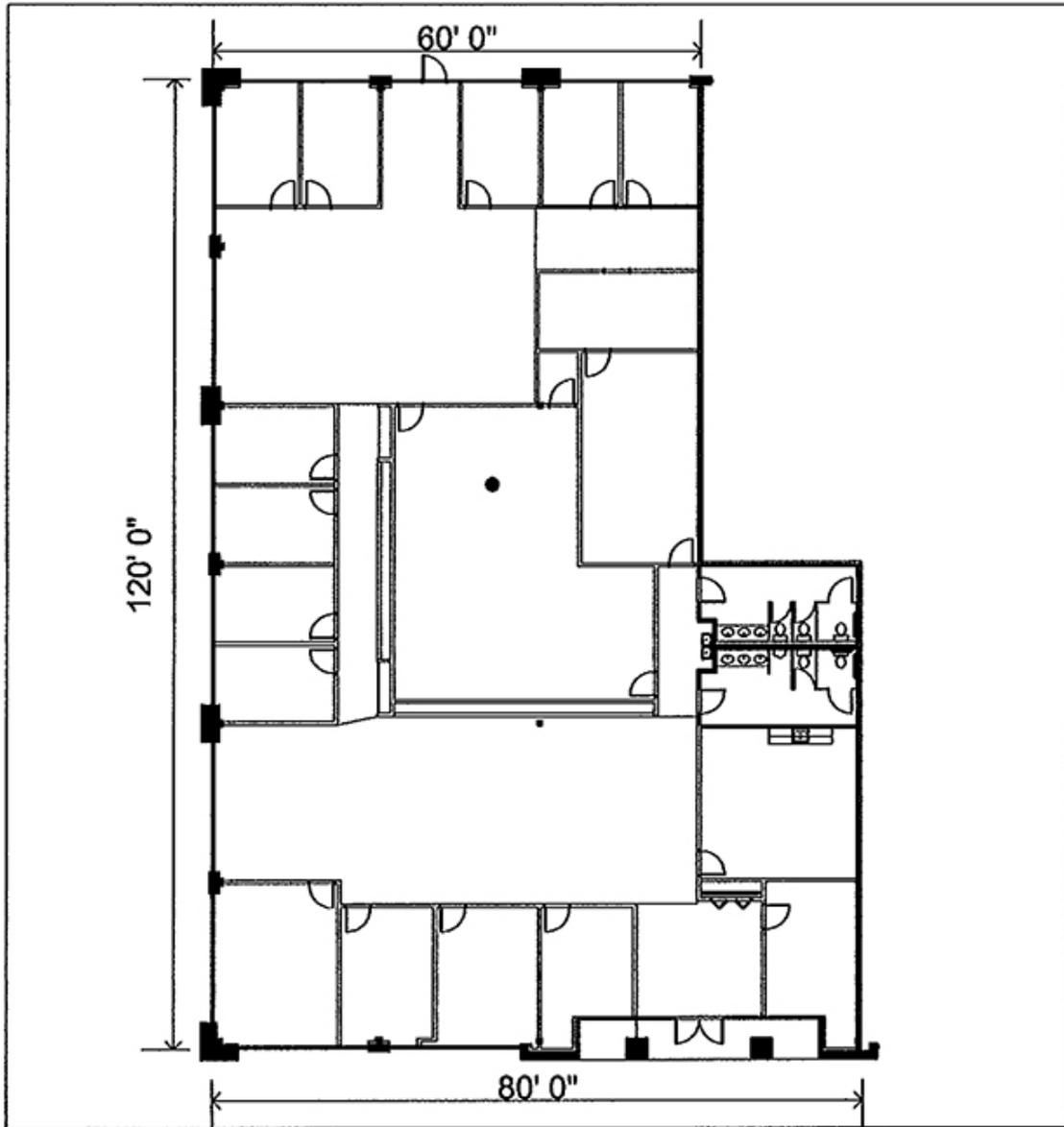
BEGINNING at an iron pin set (1/2" rebar) in the centerline of a creek, said point being 595.28 feet Easterly along the Northern right-of-way of Old Alpharetta Road (80' right-of-way) from the intersection of said right-of-way with the Eastern right-of-way of Curie Drive (right-of-way varies); thence along the centerline of said creek the following courses and distances: North 25 degrees 15 minutes 00 seconds West, a distance of 35.53 feet to a point; thence North 76 degrees 58 minutes 18 seconds West, a distance of 38.18 feet to a point; thence North 68 degrees 03 minutes 59 seconds West, a distance of 54.22 feet to a point; thence North 79 degrees 37 minutes 19 seconds West, a distance of 93.86 feet to a point; thence North 43 degrees 29 minutes 10 seconds West, a distance of 41.28 feet to a point in the centerline of a ditch; thence leaving said creek run along the centerline of said ditch the following courses and distances: North 85 degrees 37 minutes 32 seconds East a distance of 56.58 feet to a point; thence South 87 degrees 26 minutes 32 seconds East, a distance of 57.91 feet to a point; thence South 87 degrees 46 minutes 39 seconds East, a distance of 56.41 feet to a point; thence North 85 degrees 24 minutes 48 seconds East, a distance of 74.94 feet to an iron pin set (1/2" rebar); thence leaving said centerline, run North 12 degrees 55 minutes 23 seconds West, a distance of 193.67 feet to an iron pin set (1/2" rebar); thence North 03 degrees 35 minutes 37 seconds West, a distance of 239.39 feet to an iron pin set (1/2" rebar); thence North 02 degrees 43 minutes 53 seconds East, a distance of 155.00 feet to an iron pin set (1/2" rebar); thence North 09 degrees 50 minutes 59 seconds West; a distance of 530.00 feet to an iron pin set (1/2" rebar); thence North 08 degrees 40 minutes 54 seconds West, a distance of 318.37 feet to a point in the centerline of a creek, thence along the centerline of said creek the following courses and distances: North 74 degrees 51 minutes 04 seconds East, a distance of 2.98 feet to a point; thence 61 degrees 47 minutes 22 seconds East, a distance of 69.08 feet to a point; thence North 71 degrees 58 minutes 52 seconds East, a distance of 49.00 feet to a point; thence North 54 degrees 40 minutes 01 seconds East, a distance of 85.21 feet to a point; thence North 34 degrees 05 minutes 43 seconds East, a distance of 43.55 feet to a point; thence North 44 degrees 21 minutes 22 seconds East, a distance of 80.85 feet to a point; thence North 55 degrees 24 minutes 53 seconds East, a distance of 29.08 feet to a point; thence North 78 degrees 05 minutes 36 seconds East, a distance of 39.31 feet to a point; thence North 20 degrees 47 minutes 30 seconds East, a distance of 60.84 feet to a point; thence North 03 degrees 41 minutes 47 seconds West, a distance of 52.09 feet to a point; thence North 08 degrees 25 minutes 09 seconds East, a distance of 42.24 feet to a point; thence North 12 degrees 45 minutes 07 seconds West, a distance of 35.00 feet to a point; thence North 37 degrees 35 minutes 10 seconds East, a distance of 102.08 feet to a point; thence North 22 degrees 14 minutes 09 seconds West, a distance of 16.34 feet to a point; thence North 67 degrees 28 minutes 23 seconds East, a distance of 59.20 feet to a point; thence North 12 degrees 04 minutes 49 seconds East, a distance of 65.65 feet to a point; thence North 45 degrees 16 minutes 17 seconds East, a distance of 12.33 feet to a point; thence South 70 degrees 34 minutes 30 seconds East, a distance of 30.58 feet to a point; thence North 25 degrees 37 minutes 40 seconds East, a distance of 7.33 feet to a point; thence leaving said centerline of creek run South 09 degrees 41 minutes 02 seconds East, a distance of 1348.01 feet to an iron pin found (1/2" open top pipe); thence South 00 degrees 30 minutes 04 seconds West, a distance of 199.99 feet to an iron pin found (1/2" open top pipe); thence South 18 degrees 29 minutes 50 seconds West, a distance of 199.95 feet to an iron pin found (1/2" rebar); thence South 09 degrees 33 minutes 05 seconds West, a distance of 241.90 feet an iron pin set (1/2" rebar) on said Northern right-of-way line of Old Alpharetta Road; thence along said right-of-way line 447.43 feet along the arc of a curve to the left, having a chord bearing a distance of South 68 degrees 05 minutes 06 seconds West, 447.42 feet and a radius of 26,773.81 feet to the POINT OF BEGINNING.

Said tract contains 22.818 acres.

Being the same tract of land as depicted on Survey for McDonald Ventures XI, LLC, Chicago Title Insurance Company and Regions Bank, prepared by Rochester & Associates, Inc., dated March 26, 1999, last revised April 14, 1999.



Premises



Windward Chase 300 1220 Old Alpharetta Road, Suite 300, Alpharetta, Georgia 30005

Total (Building)	153,140
Total (Premises)	8,823
Warehouse	0
Office Area	8,823

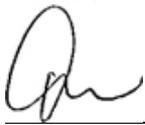


McDonald Development Company

Landlord will make the following improvements to the Premises:

- Paint walls and frames with one coat to match existing colors
- Repaint accent walls with one coat to match existing
- Wall covering to remain "as-is". Glue as needed
- Provide new vct in the restrooms, break room, and electrical room.
- Provide new cove base throughout the Premises.
- Re-Seal doors to match
- Luxury vinyl composite tile (wood look) in lobby area
- Provide new building standard carpet.
- Replace 32 T-12 lights.
- Re-Lamp 1 exit sign, re-charge 3 fire extinguishers,
- Patch sheet rock in corner of 1 office
- Demolish phone board, remove wiring,

Note: All work to be performed using Landlord's building standards attached hereto or as mutually agreed to by both Tenant and Landlord.



McDONALD DEVELOPMENT COMPANY
OUTLINE OF
STANDARD TENANT FINISH SPECIFICATIONS

1. CARPENTRY & MILLWORK

- A. Single fixture toilet rooms to have a wall-hung lavatory. Multiple fixture toilet rooms shall have plastic laminate lavatory counter-tops without base cabinet for handicap access.
- B. One coat rod and shelf in each coat closet.
- C. Two levels of adjustable shelves in storage room.
- D. Toilet partitions to be metal, in standard colors.

2. DOORS & HARDWARE

- A. Glass entrance door is existing.
- B. Interior Doors
 - 3' - 0" x 6' - 8" x 1 - 3/4" flush solid core, stain grade, birch veneer door or to match existing (New doors to have Minwax Dark Walnut #2716).
 - Door frames shall be KD hollow metal.
 - (3) 4 x 4 hinges with 26 D finish (Brushed finish).
 - Hardware shall be lever action Cal-Royal SL Series with 26D finish (Grade 2 or better).
 - Function to be compatible with room types indicated on drawings
 - Wall mounted door stops with interior wall blocking.
 - Provide closers on all bathroom doors (excluding single fixture bathrooms) and doors leading to warehouse.
- C. Labeled Doors (where required)
 - Provide labeled doors and frames with rated hinges.
 - Lever action Cal-Royal SL Series locksets.
 - Surface mounted closers.

3. PARTITIONS

- A. Demising Walls
 - Demising wall partitions shall be 1-hour rated extending from finished slab to deck (height varies) and shall be constructed with 5/8" type C gypsum wallboard screwed to both sides of 6" wide metal studs. Provide R-11 fiberglass blankets in wall cavity floor to roof deck (UL Des. U-465), painted white with black cove base.
- B. Interior Partitions
 - Typical 9' - 0" interior partitions shall extend from finished slab to finished ceiling. Construction to be of 3-5/8" wide metal studs (25 Ga.) spaced 24" o.c. with 1/2" gypsum wallboard screwed to both sides. Toilet room walls shall extend to 10'.
 - Insulation - sound attenuation blanket is to be installed in all toilet room, break room, and conference room walls.
 - All exterior tilt-up walls in office areas to receive 1/2" gypsum board on 3-5/8" wide metal studs (25 Ga.) spaced 24" o.c. with R-11 insulation.



C. Office/Warehouse Separation Wall

If not 1-hour rated: Construction to be of 6" wide metal studs with 1/2" gypsum wallboard screwed to both sides (one side to the deck, one side to 10') and R-11 insulation to deck. Warehouse side to be painted white with black cove base.

4. ACOUSTICAL CEILINGS

- A. 2' - 0" x 2' - 0" standard grid system with 5/8" non-directional fissured mineral board ceiling panels installed at 9' a.f.f. with no insulation above ceiling grid.
- B. Ceiling to be continuous over all interior partitions, except at toilet rooms.

5. FLOORING

- A. Vinyl composition tile (VCT) is to be installed in bathrooms and kitchen/break room,
- B. All other finished office areas are to be carpeted using building standard carpet.
- C. 4" rubber cove base is to be installed in all finished office areas and the warehouse side of the warehouse/office separation wall. Pre-molded tab corners to be used on all outside corners.
- D. The warehouse floor shall be sealed with Lapidolith.

6. PAINTING

- A. All walls in finished office area are to be painted with a minimum of two coats of eggshell paint.
- B. Warehouse side of office warehouse separation wall to be painted white with a minimum of two coats of eggshell paint.
- C. All door frames to be painted with oil-based enamel, semi-gloss.
- D. Toilet room walls to be painted with eggshell latex paint.

7. HVAC

- A. Heating and air conditioning in office area to be furnished by either split systems or roof top/package units to maintain a maximum 75°F with 94°F outside air temperature. Higher cooling loads imposed by equipment or heavier than normal occupancy would be an additional cost. Roof mounted equipment shall not be installed within thirty feet (30') of building perimeter unless a parapet provides adequate screening.
- B. Provide exhaust fans in toilet rooms.
- C. Heating in the warehouse is to be furnished by gas-fired unit heaters in order to maintain 50°F with 32°F outside air temperature. All vent stacks shall be provided with appropriate roof flashing that has fully soldered joints.
- D. Provide additional structural reinforcement as required for all roof mounted equipment.
- E. Warehouse ventilation provided by roof-mounted up blast fans.

8. PLUMBING

- A. Water closet and urinals in toilet rooms in quantities as indicated on drawings.
- B. Lavatory in each toilet room per drawings.
- C. 48" grab bar and 36" grab bar in handicapped toilet.
- D. Single roll toilet paper holder.
- E. Surface mounted towel dispenser.

- F. One mirror extending the full width of the lavatory counter top.
- G. Provided water heater (above ceiling in office area) sized as required to server fixtures indicated.
- H. All drinking fountains to be non-electric.

9. FIRE PROTECTION

- A. Provide sprinkler drops to accommodate the office lay-out.
- B. Provide fire extinguishers per local fire marshal's requirements.

10. ELECTRICAL

- A. Provide service as recommended by electrical contractor.
- B. Provide 3-phase, 277/480 volt electric service.
- C. Provide warehouse lighting of 20 f.c. (unracked at 36" a.f.f.) using metal halide fixtures. Switching at panel with one fixture on continuous circuit for night-light.
- D. Office lighting by 2' - 0" x 4' - 0" lay-in fluorescent fixtures with acrylic lens to maintain 50 foot candles at desk top.
- E. Provide duplex receptacles and telephone boxes as shown on the drawings (Phone and data wiring to be provided by tenant.).
- F. Exit signs and battery-pack emergency lighting as required by code.
- G. Fire alarm and security system to be provided by tenant.
- H. Provide electrical room lay-out drawing for owner's approval prior to beginning work.

11. MISCELLANEOUS

- A. Bali customizer mini blinds, 6 gauge aluminum, Snow Cap White (#386) to be placed at all storefront glass in first generation build-out. Repair and replace as necessary to match existing in renovated spaces.



McDonald Ventures XI, LLC
c/o McDonald Development Company
3715 Northside Parkway NW
Building 200, Suite 700
Atlanta, Georgia 30327

March 18, 2014

Charles A. Deignan
Chief Financial Officer
Clearside Biomedical, Inc.
1220 Old Alpharetta Road, Suite 300
Alpharetta, Georgia 30005

**Via Overnight Mail &
Email**

RE: 1220 Old Alpharetta Road, Suite 100, Alpharetta GA 30005
Renewal Notice dated November 26, 2013

Dear Charles:

We are in receipt of your letter of November 26, 2013, in which Clearside Biomedical, Inc. exercised its option to renew its Lease Agreement between McDonald Ventures XI, LLC and Clearside Biomedical, Inc. dated March 14, 2012. Please consider this letter as an acknowledgement of Clearside Biomedical, Inc.'s exercising of its Renewal Option outlined in Section 32 of the Lease Agreement.

Beginning October 1, 2014, Base Rent shall be \$7,233.00 per month (\$9.84 per square foot) and shall increase by 3% on October 151 each year thereafter. The Lease Term is extended by thirty (30) months, expiring March 31, 2017.

Sincerely,

McDonald Ventures XI, LLC, a
Georgia limited liability company

By: McDonald Industrial XI, LLC, a
Georgia limited liability company, its Manager

By: McDonald Development Company, a Georgia
corporation, its Manager

By: /s/ John R. McDonald
John R. McDonald, President

cc: Tom Davenport

FIRST AMENDMENT TO LEASE AGREEMENT

THIS FIRST AMENDMENT TO LEASE AGREEMENT (“Amendment”), made and entered into this 22nd day of August 2014, by and between Clearside Biomedical, Inc., a Delaware corporation (hereinafter referred to as “Tenant”), and McDonald Ventures XI, LLC, a Georgia limited liability company (hereinafter referred to as “Landlord”):

WITNESSETH:

WHEREAS, Landlord and Tenant entered into a lease agreement of certain premises located at 1220 Old Alpharetta Road, Suite 300, Alpharetta, Forsyth County, Georgia 30005, as evidenced by that certain Lease Agreement dated March 14, 2012 (the “Lease”); and

WHEREAS, Tenant exercised is Renewal Option via a letter to Landlord dated November 26, 2013, a copy of which is attached hereto as Exhibit “A”; and

WHEREAS, Landlord and Tenant desire to amend the Lease to set forth the terms of renewal.

NOW, THEREFORE, for and in consideration of TEN AND NO/100 DOLLARS (\$10.00), and other good and valuable consideration, the receipt, adequacy and sufficiency of which are hereby acknowledged, Landlord and Tenant hereby agree as follows:

1. Defined Terms. All capitalized terms used herein shall have the same meaning as in the Lease, unless otherwise expressly defined herein.
2. Extension of Demised Term. The Demised Term under Paragraph 2 of the Lease is hereby extended for thirty (30) full calendar months beginning October 1, 2014 (the “Renewal Commencement Date”) and ending March 31, 2017.
3. Base Rent. Commencing on the Renewal Commencement Date, Base Rent under Paragraph 3.A. of the Lease shall be Seven Thousand Two Hundred Thirty-Three and No/100s Dollars (\$7,233.00) per month.
4. Rent Increase. Base Rent set forth in Paragraph 3. above shall increase at the beginning of the thirteenth (13th) full calendar month from the Renewal Commencement Date by three percent (3%), which adjusted rent amount shall remain in effect for the next twelve (12) consecutive months and shall increase by three percent (3%) each twelve (12) months thereafter for the balance of the Demised Term.
5. Tenant Improvements. Landlord shall provide Tenant with a onetime tenant improvement allowance in the amount of \$17,646.00 (\$2.00 per square foot) (the “Allowance”) for the costs relating to construction of the Tenant Improvement Work, as hereinafter defined. Landlord shall perform the improvements shown on the space plan prepared by Loia Budde Associates dated August 7, 2014, attached hereto as Exhibit “B” (the “Tenant Improvement Work”), which shall be completed in accordance with the Standard Tenant Finish Specifications attached hereto as Exhibit “C”. The current estimate for the Tenant Improvement Work is Forty Thousand Three Hundred Forty-Two and No/100 Dollars (\$40,342.00) (the “Total Projected Cost”). Tenant shall fund the cost of any Tenant Improvement Work that exceeds the Allowance. Tenant shall pay to Landlord at execution of the Amendment the estimated overage of the Tenant Improvement Work equal to the Total Projected Cost less the Allowance, which is currently estimated to be Twenty-Two Thousand Six Hundred Ninety-Six and No/100 Dollars (\$22,696.00) (the “Estimated Tenant Overage”). Landlord and Tenant agree, that upon determination of the actual costs of the Tenant Improvement Work, the unused portion of the Estimated Tenant Overage collected from Tenant, if any, will either be returned to Tenant or applied towards Tenant’s Base Rent obligation within thirty (30) days. In the event, the Tenant Improvement Work exceeds the Estimated Tenant Overage, Tenant shall have thirty (30) days after receipt of invoice from Landlord to pay to Landlord the balance due for any cost of the Tenant Improvement Work that exceeded the Total Projected Cost. The Tenant Improvement Work shall begin after execution of the Amendment by both Tenant and Landlord and shall be completed per a mutually agreeable schedule. McDonald Construction Services shall be the General Contractor for all construction, and McDonald Development Company will oversee the construction process.

Except as expressly modified hereby, the Lease shall remain in full force and effect, Landlord and Tenant ratifying and affirming the same.

IN WITNESS WHEREOF, the parties have executed this Amendment with intent to be bound hereby as of the day and year indicated above each signature block, respectively, but effective as of the day and year first above written.

EXECUTED BY LANDLORD, this 22nd day of August, 2014.

LANDLORD

McDonald Ventures XI, LLC, a Georgia limited liability company

By: McDonald Industrial XI, LLC, a Georgia limited liability company, its Manager

By: McDonald Development Company, a Georgia corporation, its Manager

By: /s/ John R. McDonald, President
John R. McDonald, President

EXECUTED BY TENANT, this 18th day of August, 2014.

TENANT

Clearside Biomedical, Inc., a Delaware corporation

By: /s/ Daniel White
Title: President and CEO



November 26, 2013

Mr. John T. Downing
McDonald Development Company
3715 Northside Parkway, Bldg 200, Ste. 700
Atlanta, GA 30327

RE: Lease Renewal Option

Dear Mr. Downing,

Clearside Biomedical is exercising its Renewal Option as provided for in Section 32 of the lease dated March 14, 2012.

Attached please find a copy of our audited financial statements from May 26, 2011 through December 31, 2012.

If you need additional information, please do not hesitate to contact me.

Sincerely,

CLEARSIDE BIOMEDICAL, INC.

A handwritten signature in black ink, appearing to read 'Charles A. Deignan'.

Charles A. Deignan
Chief Financial Officer

CAS/sgk

Attachment (1)

1220 Old Alpharetta Road, Suite 300 - Alpharetta GA 30005
Phone 678.270.3631
www.clearsidebio.com

Exhibit "B"



Exhibit "C"

McDONALD DEVELOPMENT COMPANY
OUTLINE OF
STANDARD OFFICE FINISH SPECIFICATIONS

1. CARPENTRY & MILLWORK

- A. Single fixture toilet rooms to have a wall-hung lavatory. Multiple fixture toilet rooms shall have plastic laminate lavatory counter-tops without base cabinet for handicap access.
- B. One coat rod and shelf in each coat closet.
- C. Two levels of adjustable shelves in storage room.
- D. Toilet partitions to be metal, in standard colors.

2. DOORS & HARDWARE

- A. Glass entrance door is existing.
- B. Interior Doors
 - 3' - 0" x 6' - 8" x 1 - 3/4" flush solid core, stain grade, birch veneer door or to match existing (New doors to have Minwax Dark Walnut #2716).
 - Door frames shall be KD hollow metal.
 - (3) 4 x 4 hinges with 26 D finish (Brushed finish).
 - Hardware shall be lever action Cal-Royal SL Series with 26D finish (Grade 2 or better).
 - Function to be compatible with room types indicated on drawings
 - Wall mounted door stops with interior wall blocking.
 - Provide closers on all bathroom doors (excluding single fixture bathrooms) and doors leading to warehouse.
- C. Labeled Doors (where required)
 - Provide labeled doors and frames with rated hinges.
 - Lever action Cal-Royal SL Series locksets.
 - Surface mounted closers.

3. PARTITIONS

- A. Demising Walls
 - Demising wall partitions shall be 1-hour rated extending from finished slab to deck (height varies) and shall be constructed with 5/8" type C gypsum wallboard screwed to both sides of 6" wide metal studs Provide R-11 fiberglass blankets in wall cavity floor to roof deck (UL Des. U-465), painted white with black cove base.
- B. Interior Partitions
 - Typical 9' - 0" interior partitions shall extend from finished slab to finished ceiling. Construction to be of 3-5/8" wide metal studs (25 Ga.) spaced 24" o.c. with 1/2" gypsum wallboard screwed to both sides. Toilet room walls shall extend to 10'.
 - Insulation — sound attenuation blanket is to be installed in all toilet room, break room, and conference room walls.
 - All exterior tilt-up walls in office areas to receive 1/2" gypsum board on 3-5/8" wide metal studs (25 Ga.) spaced 24" o.c. with R-11 insulation.

4. ACOUSTICAL CEILINGS

- A. 2' - 0" x 2' - 0" standard grid system with 5/8" non-directional reveal-edge mineral board ceiling panels.
- B. Ceiling to be continuous over all interior partitions, except at toilet rooms.

5. FLOORING

- A. Vinyl composition tile (VCT) is to be installed in bathrooms and kitchen/break room.
- B. All other finished areas are to be carpeted using building standard carpet.
- C. 4" rubber cove base is to be installed in all finished areas. Pre-molded tab corners to be used on all outside corners.

6. PAINTING

- A. All walls are to be painted with a minimum of two coats of eggshell paint.
- B. All door frames to be painted with oil-based enamel, semi-gloss.
- C. Toilet room walls to be painted with eggshell latex paint.

7. HVAC

- A. Heating and air conditioning to be furnished by either split systems or roof top/package units to maintain a maximum 75°F with 94°F outside air temperature. Higher cooling loads imposed by equipment or heavier than normal occupancy would be an additional cost. Roof mounted equipment shall not be installed within thirty feet (30') of building perimeter unless a parapet provides adequate screening.
- B. Provide exhaust fans in toilet rooms.
- C. Provide additional structural reinforcement as required for all roof mounted equipment.

8. PLUMBING

- A. Water closet and urinals in toilet rooms in quantities as indicated on drawings.
- B. Lavatory in each toilet room per drawings.
- C. 48" grab bar and 36" grab bar in handicapped toilet.
- D. Single roll toilet paper holder.
- E. Surface mounted towel dispenser.
- F. One mirror extending the full width of the lavatory counter top.
- G. Provide water heater (above ceiling in office area) sized as required to serve fixtures indicated.
- H. All drinking fountains to be non-electric.

9. FIRE PROTECTION

- A. Provide sprinkler drops to accommodate the office lay-out.
- B. Provide fire extinguishers per local fire marshal's requirements.

10. ELECTRICAL

- A. Provide service as recommended by electrical contractor.
- B. Office lighting by 2' - 0" x 4' - 0" lay-in fluorescent fixtures with parabolic lens to maintain 50 foot candles at desk top.
- C. Provide duplex receptacles and telephone boxes as shown on the drawings (Phone and data wiring to be provided by tenant.).
- D. Exit signs and battery-pack emergency lighting as required by code.
- E. Fire alarm and security system to be provided by tenant.

11. MISCELLANEOUS

- A. Bali customizer mini blinds, 6 gauge aluminum, Snow Cap White (#386) to be placed at all storefront glass in first generation build-out. Repair and replace as necessary to match existing in renovated spaces.

CLEARSIDE BIOMEDICAL, INC.

2011 STOCK INCENTIVE PLAN

1. Purpose

The purpose of this 2011 Stock Incentive Plan (the “**Plan**”) of Clearside Biomedical, Inc., a Delaware corporation (the “**Company**”), is to advance the interests of the Company’s stockholders by enhancing the Company’s ability to attract, retain and motivate persons who are expected to make important contributions to the Company and by providing such persons with equity ownership opportunities and performance-based incentives that are intended to align their interests with those of the Company’s stockholders. Except where the context otherwise requires, the term “**Company**” includes the Company’s present or future parent or subsidiary corporations as defined in Sections 424(e) or (f) of the Internal Revenue Code of 1986, as amended, and any regulations promulgated thereunder (the “**Code**”) and other business ventures (including, without limitation, any joint venture or limited liability company) in which the Company has a controlling interest, as determined by the Board of Directors of the Company (the “**Board**”).

2. Eligibility

All of the Company’s employees, officers, directors, and individual consultants and advisors (each a “**Service Provider**”) are eligible to receive options, restricted stock, restricted stock units and other stock-based awards (each, an “**Award**”) under the Plan. Each person who receives an Award under the Plan is deemed a “**Participant**.”

3. Administration and Delegation

(a) Administration by Board of Directors. The Plan shall be administered by the Board. The Board shall have authority to grant Awards and to adopt, amend and repeal such administrative rules, guidelines and practices relating to the Plan as it shall deem advisable. The Board may correct any defect, supply any omission or reconcile any inconsistency in the Plan or any Award in the manner and to the extent it shall deem expedient to carry the Plan into effect and it shall be the sole and final judge of such expediency. All decisions by the Board shall be made in the Board’s sole discretion and shall be final and binding on all persons having or claiming any interest in the Plan or in any Award. No director or person acting pursuant to the authority delegated by the Board shall be liable for any action or determination relating to or under the Plan made in good faith.

(b) Appointment of Committees. To the extent permitted by applicable law, the Board may delegate any or all of its powers under the Plan to one or more committees or subcommittees of the Board (a “**Committee**”). All references in the Plan to the “**Board**” shall mean the Board or a Committee of the Board to the extent that the Board’s powers or authority under the Plan have been delegated to such Committee.

4. Stock Available for Awards.

(a) Subject to adjustment under Section 8, Awards may be made under the Plan for up to 1,119,744 shares of the common stock of the Company, \$0.001 par value per share (the “**Common Stock**”). If any Award expires or is terminated, surrendered or canceled without having been fully exercised or is forfeited in whole or in part (including as the result of shares of Common Stock subject to

such Award being repurchased by the Company at the original issuance price pursuant to a contractual repurchase right) or results in any Common Stock not being issued, the unused Common Stock covered by such Award shall again be available for the grant of Awards under the Plan. Further, shares of Common Stock tendered to the Company by a Participant to exercise an Award shall be added to the number of shares of Common Stock available for the grant of Awards under the Plan. However, in the case of Incentive Stock Options (as hereinafter defined), the foregoing provisions shall be subject to any limitations under the Code. Shares issued under the Plan may consist in whole or in part of authorized but unissued shares or treasury shares. At no time while there is any Option (as defined below) outstanding and held by a Participant who was a resident of the State of California on the date of grant of such Option, shall the total number of shares of Common Stock issuable upon exercise of all outstanding options and the total number of shares provided for under any stock bonus or similar plan or agreement of the Company exceed the applicable percentage as calculated in accordance with the conditions and exclusions of Section 260.140.45 of the California Code of Regulations, as amended (the “**California Regulations**”), based on the shares of the Company which are outstanding at the time the calculation is made unless the Plan complies with all conditions of Rule 701 of the Securities Act of 1933, as amended.

(b) Substitute Awards. In connection with a merger or consolidation of an entity with the Company or the acquisition by the Company of property or stock of an entity, the Board may grant Awards in substitution for any options or other stock or stock-based awards granted by such entity or an affiliate thereof. Substitute Awards may be granted on such terms as the Board deems appropriate in the circumstances, notwithstanding any limitations on Awards contained in the Plan. Substitute Awards shall not count against the overall share limit set forth in Section 4(a), except as may be required by reason of Section 422 and related provisions of the Code.

5. Stock Options

(a) General. The Board may grant options to purchase Common Stock (each, an “**Option**”) and determine the number of shares of Common Stock to be covered by each Option, the exercise price of each Option and the conditions and limitations applicable to the exercise of each Option, including conditions relating to applicable federal or state securities laws, as it considers necessary or advisable. An Option, or portion of an Option, which is not intended to be or fails to qualify as an Incentive Stock Option (as hereinafter defined) shall be designated a “**Nonstatutory Stock Option.**”

(b) Incentive Stock Options. An Option that the Board intends to be an “incentive stock option” as defined in Section 422 of the Code (an “**Incentive Stock Option**”) shall only be granted to employees of the Company and any other entities the employees of which are eligible to receive Incentive Stock Options under the Code, and shall be subject to and shall be construed consistently with the requirements of Section 422 of the Code. A Participant who owns more than 10% of the total combined voting power of all classes of outstanding stock of the Company shall not be eligible for the grant of an Incentive Stock Option unless (i) the exercise price is at least 110% of the Fair Market Value (as defined below) on the date the Option is granted and (ii) such Incentive Stock Option by its terms is not exercisable after the expiration of five years from the date the Option is granted. The Company shall have no liability to a Participant, or any other party, if an Option (or any part thereof) that is intended to be an Incentive Stock Option is not an Incentive Stock Option or for any action taken by the Board pursuant to Section 9(f), including without limitation the conversion of an Incentive Stock Option to a Nonstatutory Stock Option.

(c) Exercise Price. The Board shall establish the exercise price of each Option and specify such exercise price in the applicable option agreement. The exercise price shall be not less than 100% of the Fair Market Value on the date the Option is granted unless the Board specifically determines that the

exercise price is intended to be less than such Fair Market Value, in which case the option agreement shall contain provisions complying with Section 409A of the Code; provided that if the Board approves the grant of an Option with an exercise price to be determined on a future date, the exercise price shall be not less than 100% of the Fair Market Value on such future date. The term “**Fair Market Value**” shall mean, as of a given date: (i) if the Common Stock is listed on a national securities exchange, the last sale price of the Common Stock in the principal trading market for the Common Stock on such date; (ii) if the Common Stock is not listed on a national securities exchange, but is traded in the over-the counter market, the closing bid price for the Common Stock on such date, as reported by the OTC Bulletin Board or the National Quotation Bureau, Incorporated or similar publisher of such quotations; or (iii) if the Common Stock is not listed on a national securities exchange or traded in the over-the-counter market, such price as shall be determined by (or in a manner approved by) the Board in good faith and in compliance with applicable provisions of the Code and the regulations issued thereunder.

(d) Duration of Options. Each Option shall be exercisable at such times and subject to such terms and conditions as the Board may specify in the applicable option agreement.

(e) Exercise of Option. Options may be exercised by delivery to the Company of a written notice of exercise signed by the proper person or by any other form of notice (including electronic notice) approved by the Board together with payment in full as specified in Section 5(f) for the number of shares of Common Stock for which the Option is exercised. Shares of Common Stock subject to the Option will be delivered by the Company following exercise either as soon as practicable or, subject to such conditions as the Board shall specify, on a deferred basis (with the Company’s obligation to be evidenced by an instrument providing for future delivery of the deferred shares at the time or times specified by the Board).

(f) Payment Upon Exercise. Common Stock purchased upon the exercise of an Option granted under the Plan shall be paid for as follows:

(1) in cash or by check, payable to the order of the Company;

(2) except as may otherwise be provided in the applicable option agreement, by (i) delivery of an irrevocable and unconditional undertaking by a creditworthy broker to deliver promptly to the Company sufficient funds to pay the exercise price and any required tax withholding or (ii) delivery by the Participant to the Company of a copy of irrevocable and unconditional instructions to a creditworthy broker to deliver promptly to the Company cash or a check sufficient to pay the exercise price and any required tax withholding;

(3) when the Common Stock is registered under the Securities Exchange Act of 1934, as amended (the “**Exchange Act**”) and to the extent provided for in the applicable option agreement or approved by the Board, in its sole discretion, by delivery (either by actual delivery or attestation) of shares of Common Stock owned by the Participant valued at their Fair Market Value, provided (i) such method of payment is then permitted under applicable law, (ii) such Common Stock, if acquired directly from the Company, was owned by the Participant for such minimum period of time, if any, as may be established by the Board in its discretion and (iii) such Common Stock is not subject to any repurchase, forfeiture, unfulfilled vesting or other similar requirements;

(4) to the extent permitted by applicable law and provided for in the applicable option agreement or approved by the Board, in its sole discretion, by (i) delivery of a promissory note of the Participant to the Company on terms determined by the Board, or (ii) payment of such other lawful consideration as the Board may determine; or

(5) by any combination of the above permitted forms of payment.

6. Restricted Stock; Restricted Stock Units

(a) General. The Board may grant Awards entitling recipients to acquire shares of Common Stock (“**Restricted Stock**”), subject to the right of the Company to repurchase all or part of such shares at their issue price or other stated or formula price (or to require forfeiture of such shares if issued at no cost) from the recipient in the event that conditions specified by the Board in the applicable Award are not satisfied prior to the end of the applicable restriction period or periods established by the Board for such Award. Instead of granting Awards for Restricted Stock, the Board may grant Awards entitling the recipient to receive shares of Common Stock to be delivered at the time such shares of Common Stock vest (“**Restricted Stock Units**”) (Restricted Stock and Restricted Stock Units are each referred to herein as a “**Restricted Stock Award**”).

(b) Terms and Conditions. The Board shall determine the terms and conditions of a Restricted Stock Award, including the conditions for repurchase (or forfeiture) and the issue price, if any.

(c) Additional Provisions Relating to Restricted Stock.

(1) Dividends. Participants holding shares of Restricted Stock will be entitled to all ordinary cash dividends paid with respect to such shares, unless otherwise provided by the Board. If any such dividends or distributions are paid in shares, or consist of a dividend or distribution to holders of Common Stock other than an ordinary cash dividend, the shares, cash or other property will be subject to the same restrictions on transferability and forfeitability as the shares of Restricted Stock with respect to which they were paid. Each dividend payment will be made no later than the end of the calendar year in which the dividends are paid to stockholders of that class of stock or, if later, the 15th day of the third month following the date the dividends are paid to stockholders of that class of stock.

(2) Stock Certificates. The Company may require that any stock certificates issued in respect of a Restricted Stock Award shall be registered in the name of the Participant and be deposited by the Participant, together with a stock power endorsed in blank, with the Company (or its designee). After the expiration of the applicable restriction periods, upon request of a Participant or as otherwise determined by the Company, the Company (or such designee) shall deliver the certificates no longer subject to such restrictions to the Participant or if the Participant has died, to the beneficiary designated, in a manner determined by the Board, by a Participant to receive amounts due or exercise rights of the Participant in the event of the Participant’s death (the “**Designated Beneficiary**”). In the absence of an effective designation by a Participant, “Designated Beneficiary” shall mean the Participant’s then living spouse, or, if none, the Participant’s estate.

7. Other Stock-Based Awards

Other Awards of shares of Common Stock, and other Awards that are valued in whole or in part by reference to, or are otherwise based on, shares of Common Stock or other property, may be granted hereunder to Participants (“**Other Stock-Based Awards**”), including without limitation stock appreciation rights and Awards entitling recipients to receive shares of Common Stock to be delivered in the future. Such Other Stock-Based Awards shall also be available as a form of payment in the settlement of other Awards granted under the Plan or as payment in lieu of compensation to which a Participant is otherwise entitled. Other Stock-Based Awards may be paid in shares of Common Stock or cash, as the Board shall determine. Subject to the provisions of the Plan, the Board shall determine the conditions of each Other Stock-Based Award, including any purchase price applicable thereto.

8. Adjustments for Changes in Common Stock and Certain Other Events

(a) Changes in Capitalization. In the event of any stock split, reverse stock split, stock dividend, recapitalization, combination of shares, reclassification of shares, spin-off or other similar change in capitalization or event, or any dividend or distribution to holders of Common Stock other than an ordinary cash dividend, (i) the number and class of securities available under this Plan, (ii) the number and class of securities and exercise price per share of each outstanding Option, (iii) the number of shares subject to and the repurchase price per share subject to each outstanding Restricted Stock Award, and (iv) the terms of each other outstanding Award shall be equitably adjusted by the Company (or substituted Awards may be made, if applicable) in the manner determined by the Board. Without limiting the generality of the foregoing, in the event the Company effects a split of the Common Stock by means of a stock dividend and the exercise price of and the number of shares subject to an outstanding Option are adjusted as of the date of the distribution of the dividend (rather than as of the record date for such dividend), then an optionee who exercises an Option between the record date and the distribution date for such stock dividend shall be entitled to receive, on the distribution date, the stock dividend with respect to the shares of Common Stock acquired upon such Option exercise, notwithstanding the fact that such shares were not outstanding as of the close of business on the record date for such stock dividend.

(b) Change in Control

(1) Definition. Unless otherwise specifically provided in an Award agreement, a “**Change in Control**” shall be deemed to have occurred upon the first to occur of:

(i) any “person” (as such term is used in sections 13(d) and 14(d) of the Exchange Act) becoming a “beneficial owner” (as defined in Rule 13d-3 under the Exchange Act), directly or indirectly, of securities of the Company representing either (A) more than a majority of the voting power of the then outstanding securities of the Company, or (B) more than a majority of the aggregate fair market value of the then outstanding securities of the Company; provided, however, that a Change in Control shall not be deemed to occur as a result of (x) a transaction in which the Company becomes a subsidiary of another corporation and in which the stockholders of the Company, immediately prior to the transaction, will beneficially own, immediately after the transaction, shares entitling such stockholders to more than majority of all votes to which all stockholders of the parent corporation would be entitled in the election of directors, or (y) a transaction in which the person acquires newly issued securities of the Company in exchange for an investment in the Company; or

(ii) the consummation of either: (A) a merger, share exchange, consolidation or reorganization of the Company where the stockholders of the Company, immediately prior to the merger or consolidation, will not beneficially own, immediately after the merger, share exchange, consolidation or reorganization, shares entitling such stockholders to either (x) more than a majority of all votes to which all stockholders of the surviving corporation would be entitled in the election of directors, or (y) more than a majority of the aggregate fair market value of then outstanding securities of the Company; or (B) a sale or other disposition of all or substantially all of the assets of the Company.

(2) Consequences of a Change in Control on Awards Other than Restricted Stock Awards. In connection with a Change in Control, the Board may take any one or more of the following actions as to all (or any portion of) outstanding Awards other than Restricted Stock Awards on such terms as the Board determines: (i) provide that Awards shall be assumed, or substantially equivalent Awards

shall be substituted, by the acquiring or succeeding corporation (or an affiliate thereof) in compliance with the applicable provisions of the Code, including Code Sections 409A, 422 and 424, (ii) upon written notice to a Participant, provide that the Participant's unexercised Options or other unexercised Awards will terminate immediately prior to the consummation of such Change in Control unless exercised by the Participant within a specified period following the date of such notice, (iii) provide that outstanding Awards shall become exercisable, realizable or deliverable, or restrictions applicable to an Award shall lapse, in whole or in part prior to or upon such Change in Control, (iv) in the event of a Change in Control under the terms of which holders of Common Stock will receive upon consummation thereof a cash payment for each share surrendered in the Change in Control (the "**Acquisition Price**"), make or provide for a cash payment to a Participant equal to the excess, if any, of (A) the Acquisition Price times the number of shares of Common Stock subject to the Participant's Options or other Awards (to the extent the exercise price does not exceed the Acquisition Price) less (B) the aggregate exercise price of all such outstanding Options or other Awards and any applicable tax withholdings, in exchange for the termination of such Options or other Awards, (v) provide that, in connection with a liquidation or dissolution of the Company, Awards shall convert into the right to receive liquidation proceeds (if applicable, net of the exercise price thereof) and (vi) any combination of the foregoing. In taking any of the actions permitted under this Section 8(b), the Board shall not be obligated by the Plan to treat all Awards, or all Awards of the same type, identically.

For purposes of clause (i) above, an Option shall be considered assumed if, following consummation of the Change in Control, the Option confers the right to purchase, for each share of Common Stock subject to the Option immediately prior to the consummation of the Change in Control, the consideration (whether cash, securities or other property) received as a result of the Change in Control by holders of Common Stock for each share of Common Stock held immediately prior to the consummation of the Change in Control (and if holders were offered a choice of consideration, the type of consideration chosen by the holders of a majority of the outstanding shares of Common Stock); provided, however, that if the consideration received as a result of the Change in Control is not solely common stock of the acquiring or succeeding corporation (or an affiliate thereof), the Company may, with the consent of the acquiring or succeeding corporation, provide for the consideration to be received upon the exercise of Options to consist solely of common stock of the acquiring or succeeding corporation (or an affiliate thereof) with equivalent in value (as determined by the Board) to the per share consideration received by holders of outstanding shares of Common Stock as a result of the Change in Control.

(3) Consequences of a Change in Control on Restricted Stock Awards. Upon the occurrence of a Change in Control other than a liquidation or dissolution of the Company, the repurchase and other rights of the Company under each outstanding Restricted Stock Award shall inure to the benefit of the Company's successor and shall, unless the Board determines otherwise, apply to the cash, securities or other property which the Common Stock was converted into or exchanged for pursuant to such Change in Control in the same manner and to the same extent as they applied to the Common Stock subject to such Restricted Stock Award. Upon the occurrence of a Change in Control involving the liquidation or dissolution of the Company, except to the extent specifically provided to the contrary in the instrument evidencing any Restricted Stock Award or any other agreement between a Participant and the Company, all restrictions and conditions on all Restricted Stock Awards then outstanding shall automatically be deemed terminated or satisfied.

9. General Provisions Applicable to Awards

(a) Transferability of Awards. Except as the Board may otherwise expressly determine or provide in an Award, Awards shall not be sold, assigned, transferred, pledged or otherwise encumbered by the person to whom they are granted, either voluntarily or by operation of law, except by will or the

laws of descent and distribution or, other than in the case of an Incentive Stock Option, pursuant to a qualified domestic relations order, and, during the life of the Participant, shall be exercisable only by the Participant. References to a Participant, to the extent relevant in the context, shall include references to authorized transferees.

(b) Documentation. Unless otherwise expressly determined by the Board, each Incentive Stock Option shall be evidenced by a Notice of Incentive Stock Option and Incentive Stock Option Agreement substantially in the form attached as Exhibit A, each Nonstatutory Stock Option shall be evidenced by a Notice of Nonstatutory Stock Option and Nonstatutory Stock Option Agreement substantially in the form attached as Exhibit B, and each Restricted Stock Award shall be evidenced by a Summary of Restricted Stock Purchase and Restricted Stock Purchase Agreement substantially in the form attached as Exhibit C. Each Award may contain terms and conditions in addition to those set forth in the Plan.

(c) Board Discretion. Except as otherwise provided by the Plan, each Award may be made alone or in addition or in relation to any other Award. The terms of each Award need not be identical, and the Board need not treat Participants uniformly.

(d) Termination of Status. The Board shall determine the effect on an Award of the disability, death, termination of employment, authorized leave of absence or other change in the employment or other status of a Participant and the extent to which, and the period during which, the Participant, or the Participant's legal representative, conservator, guardian or Designated Beneficiary, may exercise rights under the Award.

(e) Withholding. The Participant must satisfy all applicable federal, state, and local or other income and employment tax withholding obligations before the Company will deliver stock certificates or otherwise recognize ownership of Common Stock under an Award. The Company may decide to satisfy the withholding obligations through additional withholding on salary or wages. If the Company elects not to or cannot withhold from other compensation, the Participant must pay the Company the full amount, if any, required for withholding or have a broker tender to the Company cash equal to the withholding obligations. Payment of withholding obligations is due before the Company will issue any shares on exercise or release from forfeiture of an Award or, if the Company so requires, at the same time as is payment of the exercise price unless the Company determines otherwise. If provided for in an Award or approved by the Board in its sole discretion, a Participant may satisfy such tax obligations in whole or in part by delivery of shares of Common Stock, including shares retained from the Award creating the tax obligation, valued at their Fair Market Value; provided, however, except as otherwise provided by the Board, that the total tax withholding where stock is being used to satisfy such tax obligations cannot exceed the Company's minimum statutory withholding obligations (based on minimum statutory withholding rates for federal and state tax purposes, including payroll taxes, that are applicable to such supplemental taxable income). Shares surrendered to satisfy tax withholding requirements cannot be subject to any repurchase, forfeiture, unfulfilled vesting or other similar requirements.

(f) Amendment of Award.

(1) The Board may amend, modify or terminate any outstanding Award, including but not limited to, substituting therefor another Award of the same or a different type, changing the date of exercise or realization, and converting an Incentive Stock Option to a Nonstatutory Stock Option, provided that the Participant's consent to such action shall be required unless the Board determines that the action, taking into account any related action, would not materially and adversely affect the Participant.

(2) The Board may, without stockholder approval, amend any outstanding Award granted under the Plan to provide an exercise price per share that is lower than the then-current exercise price per share of such outstanding Award provided that such amended exercise price is at least equal to the then-current Fair Market Value. The Board may also, without stockholder approval, cancel any outstanding award (whether or not granted under the Plan) and grant in substitution new Awards under the Plan covering the same or a different number of shares of Common Stock and having an exercise price per share lower than the then-current exercise price per share of the cancelled award.

(g) Conditions on Delivery of Stock. The Company will not be obligated to deliver any shares of Common Stock pursuant to the Plan or to remove restrictions from shares previously delivered under the Plan until (i) all conditions of the Award have been met or removed to the satisfaction of the Company, (ii) in the opinion of the Company's counsel, all other legal matters in connection with the issuance and delivery of such shares have been satisfied, including any applicable securities laws and any applicable stock exchange or stock market rules and regulations, and (iii) the Participant has executed and delivered to the Company such representations or agreements as the Company may consider appropriate to satisfy the requirements of any applicable laws, rules, regulations or contracts of the Company.

(h) Acceleration. The Board may at any time provide that any Award shall become immediately exercisable in full or in part, free of some or all restrictions or conditions, or otherwise realizable in full or in part, as the case may be.

10. Miscellaneous

(a) No Right To Employment or Other Status. No person shall have any claim or right to be granted an Award, and the grant of an Award shall not be construed as giving a Participant the right to continued employment or any other relationship with the Company. The Company expressly reserves the right at any time to dismiss or otherwise terminate its relationship with a Participant free from any liability or claim under the Plan, except as expressly provided in the applicable Award.

(b) No Rights As Stockholder. Subject to the provisions of the applicable Award, no Participant or Designated Beneficiary shall have any rights as a stockholder with respect to any shares of Common Stock to be distributed with respect to an Award until becoming the record holder of such shares. Notwithstanding the foregoing, in the event the Company effects a split of the Common Stock by means of a stock dividend or otherwise and the exercise price of and the number of shares subject to such Option are adjusted as of the effective date of the stock dividend or split (rather than as of the record date for such stock dividend or split), then an optionee who exercises an Option between the record date and the distribution date for such stock dividend or split shall be entitled to receive, on the distribution date, the stock dividend or split with respect to the shares of Common Stock acquired upon such Option exercise, notwithstanding the fact that such shares were not outstanding as of the close of business on the record date for such stock dividend or split.

(c) Effective Date and Term of Plan. The Plan shall become effective on the date on which it is adopted by the Board. No Awards shall be granted under the Plan after the expiration of 10 years from the earlier of (i) the date on which the Plan was adopted by the Board or (ii) the date the Plan was approved by the Company's stockholders, but Awards previously granted may extend beyond that date.

(d) Amendment of Plan. The Board may amend, suspend or terminate the Plan or any portion thereof at any time; provided, however, that if at any time the approval of the Company's stockholders is required as to any modification or amendment under Section 422 of the Code or any

successor provision with respect to Incentive Stock Options, the Board may not effect such modification or amendment without such approval. Unless otherwise specified in the amendment, any amendment to the Plan adopted in accordance with this Section 10(d) shall apply to, and be binding on the holders of, all Awards outstanding under the Plan at the time the amendment is adopted, provided the Board determines that such amendment does not materially and adversely affect the rights of Participants under the Plan.

(e) Authorization of Sub-Plans. The Board may from time to time establish one or more sub-plans under the Plan for purposes of satisfying applicable blue sky, securities or tax laws of various jurisdictions. The Board shall establish such sub-plans by adopting supplements to this Plan containing (i) such limitations on the Board's discretion under the Plan as the Board deems necessary or desirable or (ii) such additional terms and conditions not otherwise inconsistent with the Plan as the Board shall deem necessary or desirable. All supplements adopted by the Board shall be deemed to be part of the Plan, but each supplement shall apply only to Participants within the affected jurisdiction and the Company shall not be required to provide copies of any supplement to Participants in any jurisdiction which is not the subject of such supplement.

(f) Compliance with Code Section 409A. It is intended that all Awards granted hereunder be either exempt from, or issued in compliance with, Code Section 409A. The Company shall have no liability to a Participant, or any other party, if an Award that is intended to be exempt from, or compliant with, Code Section 409A is not so exempt or compliant, or for any action taken by the Board.

(g) Governing Law. The provisions of the Plan and all Awards made hereunder shall be governed by and construed in accordance with the General Corporation Law of the State of Delaware, as to matters within the scope thereof, and the internal laws of the State of North Carolina (without reference to conflict of law provisions), as to all other matters.

* * * * *

CLEARSIDE BIOMEDICAL, INC.

2011 STOCK INCENTIVE PLAN

CALIFORNIA SUPPLEMENT

Pursuant to Section 10(e) of the Plan, the Board has adopted this supplement for purposes of satisfying the requirements of Section 25102(o) of the California Corporations Code, as amended:

Any Awards granted under the Plan to a Participant who is a resident of the State of California on the date of grant (a “**California Participant**”) shall be subject to the following additional limitations, terms and conditions:

1. Additional Limitations on Awards.

(a) Generally. The terms of all Awards granted to a California Participant under Sections 5, 6 or 7 of the Plan shall comply, to the extent applicable, with Section 260.140.41 or Section 260.140.42 of the California Regulations.

(b) Maximum Duration of Options. No Options granted to California Participants shall have a term in excess of 10 years measured from the Option grant date.

(c) Minimum Exercise Period Following Termination. Unless a California Participant’s employment is terminated for cause (as defined by applicable law, the terms of any contract of employment between the Company and such Participant, or in the instrument evidencing the grant of such Participant’s Option), in the event of termination of employment of such Participant, such Participant shall have the right to exercise an Option, to the extent that he or she was otherwise entitled to exercise such Option on the date employment terminated, until the earlier of the Option expiration date or: (i) at least six months from the date of termination, if termination was caused by such Participant’s death or “**permanent and total disability**” (within the meaning of Section 22(e)(3) of the Code) and (ii) at least 30 days from the date of termination, if termination was caused other than by such Participant’s death or “permanent and total disability” (within the meaning of Section 22(e)(3) of the Code).

2. Additional Requirement to Provide Information to California Participants. Unless the Plan or agreement complies with all conditions of Rule 701 of the Securities Act of 1933, as amended (“**Rule 701**”), the Company shall provide to each California Participant and to each California Participant who acquires Common Stock pursuant to the Plan, not less frequently than annually, copies of annual financial statements (which need not be audited). The Company shall not be required to provide such statements to key employees whose duties in connection with the Company assure their access to equivalent information or when the Plan or agreement complies with all conditions of Rule 701.

3. Additional Limitations on Timing of Awards. No Award granted to a California Participant shall become exercisable, vested or realizable, as applicable to such Award, unless the Plan has been approved by the holders of at least a majority of the Company’s outstanding voting securities by the later of (i) within 12 months before or after the date the Plan was adopted by the Board or the agreement entered into; and (ii) prior to or within 12 months of the granting of any option or issuance of any security under the Plan or agreement to a California Participant.

4. Additional Restriction Regarding Recapitalizations, Stock Splits, Etc. For purposes of Section 8 of the Plan, in the event of a stock split, reverse stock split, stock dividend, recapitalization, combination,

reclassification or other distribution of the Company's securities, the number of securities allocated to each California Participant must be adjusted proportionately and without the receipt by the Company of any consideration from any California Participant.

EXHIBIT A

**Notice of Incentive Stock Option
and
Incentive Stock Option Agreement**

EXHIBIT B

**Notice of Nonstatutory Stock Option
and
Nonstatutory Stock Option Agreement**

EXHIBIT C

Summary of Restricted Stock Purchase and Restricted Stock Purchase Agreement

**FIRST AMENDMENT TO THE
CLEARSIDE BIOMEDICAL, INC. 2011 STOCK INCENTIVE PLAN**

December 29, 2011

THIS FIRST AMENDMENT to the Clearside BioMedical, Inc. 2011 Stock Incentive Plan (the "**Plan**") is effective as of the date set forth above.

WHEREAS, the Board of Directors (the "**Board**") of Clearside BioMedical, Inc., a Delaware corporation (the "**Company**"), has adopted and the stockholders of the Company have approved the Plan; and

WHEREAS, the Board has approved this amendment of the Plan in order to increase the number of shares of Common Stock of the Company issuable pursuant to awards granted under the Plan by 569,744 shares, from 550,000 to 1,119,744 shares.

NOW, THEREFORE, the Plan shall be amended as follows:

1. The first sentence of Paragraph 4(a) of the Plan shall be deleted in its entirety and the following substituted in lieu thereof:

"Subject to adjustment under Section 8, Awards may be made under the Plan for up to 1,119,744 shares of the Common Stock of the Company, \$0.001 par value per share (the "**Common Stock**)."

2. Except as amended herein, the terms and provisions of the Plan shall remain unchanged and in full force and effect.

[Remainder of Page Intentionally Left Blank]

**SECOND AMENDMENT TO THE
CLEARSIDE BIOMEDICAL, INC. 2011 STOCK INCENTIVE PLAN**

THIS SECOND AMENDMENT to the Clearside Biomedical, Inc. 2011 Stock Incentive Plan (the “**Plan**”) is effective as of January 31, 2013.

WHEREAS, the Board of Directors (the “**Board**”) of Clearside Biomedical, Inc., a Delaware corporation (the “**Company**”), has adopted, and the stockholders of the Company have approved, the Plan; and

WHEREAS, the Board and the stockholders of the Company have approved this amendment of the Plan in order to increase the number of shares of stock issuable pursuant to awards granted under the Plan by 1,120,764 shares from 1,119,744 shares to 2,240,508 shares.

NOW, THEREFORE, the Plan shall be amended as follows:

1. The first sentence of Section 4(a) of the Plan shall be deleted in its entirety and the following substituted in lieu thereof:

“Subject to adjustment under Section 8, Awards may be made under the Plan for up to 2,240,508 shares of the common stock of the Company, \$0.001 par value per share (the “**Common Stock**”).”

2. Except as amended herein, the terms and provisions of the Plan shall remain unchanged and in full force and effect.

[Remainder of Page Intentionally Left Blank]

**THIRD AMENDMENT TO THE
CLEARSIDE BIOMEDICAL, INC. 2011 STOCK INCENTIVE PLAN**

THIS THIRD AMENDMENT to the Clearside Biomedical, Inc. 2011 Stock Incentive Plan (the “**Plan**”) is effective as of August 28, 2014.

WHEREAS, the Board of Directors (the “**Board**”) of Clearside Biomedical, Inc., a Delaware corporation (the “**Company**”), has adopted, and the stockholders of the Company have approved, the Plan; and

WHEREAS, the Board and the stockholders of the Company have approved this amendment of the Plan in order to increase the number of shares of stock issuable pursuant to awards granted under the Plan by 1,098,268 shares from 2,240,508 shares to 3,338,776 shares.

NOW, THEREFORE, the Plan shall be amended as follows:

1. The first sentence of Section 4(a) of the Plan shall be deleted in its entirety and the following substituted in lieu thereof:

“Subject to adjustment under Section 8, Awards may be made under the Plan for up to 3,338,776 shares of the common stock of the Company, \$0.001 par value per share (the “**Common Stock**”).”

2. Except as amended herein, the terms and provisions of the Plan shall remain unchanged and in full force and effect.

[Remainder of Page Intentionally Left Blank]