

AGENUS INC
Form 10-K
March 16, 2018

UNITED STATES

SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

Form 10-K

ANNUAL REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934
For the fiscal year ended December 31, 2017

or

TRANSITION REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF
1934

For the transition period from _____ to _____

Commission File Number: 000-29089

Agenus Inc.

(exact name of registrant as specified in its charter)

Delaware 06-1562417
(State or other jurisdiction of (I.R.S. Employer
incorporation or organization) Identification No.)

3 Forbes Road, Lexington, Massachusetts 02421

(Address of principal executive offices, including zip code)

Registrant's telephone number, including area code:

(781) 674-4400

Securities registered pursuant to Section 12(b) of the Act:

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Common Stock, \$.01 Par Value	The NASDAQ Capital Market
(Title of each class)	(Name of each exchange on which registered)

Securities registered pursuant to Section 12(g) of the Act:

None

Indicate by check mark if the registrant is a well-known seasoned issuer, as defined in Rule 405 of the Securities Act. Yes No

Indicate by check mark if the registrant is not required to file reports pursuant to Section 13 or Section 15(d) of the Act. Yes No

Indicate by check mark whether the registrant (1) has filed all reports required to be filed by Section 13 or 15(d) of the Securities Exchange Act of 1934 during the preceding 12 months (or for such shorter period that the registrant was required to file such reports), and (2) has been subject to such filing requirements for the past 90 days. Yes No

Indicate by check mark whether the registrant has submitted electronically and posted on its corporate website, if any, every Interactive Data File required to be submitted and posted pursuant to Rule 405 of Regulations S-T (§232.405 of this chapter) during the preceding 12 months (or for such shorter period that the registrant was required to submit and post such files).

Indicate by check mark if disclosure of delinquent filers pursuant to Item 405 of Regulation S-K (§229.405 of this chapter) is not contained herein, and will not be contained, to the best of registrant's knowledge, in definitive proxy or information statements incorporated by reference in Part III of this Form 10-K or any amendment to this Form 10-K.

Indicate by check mark whether the registrant is a large accelerated filer, an accelerated filer, a non-accelerated filer, a smaller reporting company, or an emerging growth company. See the definitions of "large accelerated filer," "accelerated filer," "smaller reporting company" and "emerging growth company" in Rule 12b-2 of the Exchange Act.

<input type="checkbox"/>	Large accelerated filer	<input type="checkbox"/>	Accelerated filer
<input type="checkbox"/>	Non-accelerated filer	(Do not check if a smaller reporting company)	<input checked="" type="checkbox"/> Smaller reporting company
<input type="checkbox"/>	Emerging growth company		

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act.

Indicate by check mark whether the registrant is a shell company (as defined in Rule 12b-2 of the Exchange Act). Yes No

The aggregate market value of voting stock held by non-affiliates of the registrant as of June 30, 2017 was: \$306.4 million. There were 102,556,797 shares of the registrant's Common Stock outstanding as of February 28, 2018.

DOCUMENTS INCORPORATED BY REFERENCE

None.

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Note Regarding Forward-Looking Statements

This Annual Report on Form 10-K and other written and oral statements the Company makes from time to time contain forward-looking statements. You can identify these forward-looking statements by the fact they use words such as “could,” “expect,” “anticipate,” “estimate,” “target,” “may,” “project,” “guidance,” “intend,” “plan,” “believe,” “will,” “opportunity,” “future” and other words and terms of similar meaning. Forward-looking statements include discussion of future operating or financial performance. You also can identify forward-looking statements by the fact that they do not relate strictly to historical or current facts. Forward-looking statements involve risks and uncertainties that could delay, divert or change any of them, and could cause actual outcomes to differ materially. These statements relate to, among other things, our business strategy, our research and development, our product development efforts, our ability to commercialize our product candidates, the activities of our licensees, our prospects for initiating partnerships or collaborations, the timing of the introduction of products, the effect of new accounting pronouncements, our future operating results and our potential profitability, availability of additional capital as well as our plans, objectives, expectations, and intentions.

Although we believe we have been prudent in our plans and assumptions, no assurance can be given that any goal or plan set forth in forward-looking statements can be achieved, and readers are cautioned not to place undue reliance on such statements, which speak only as of the date of this report. We undertake no obligation to release publicly any revisions to forward-looking statements as a result of new information, future events or otherwise.

The risks identified in this Annual Report on Form 10-K, including, without limitation, the risks set forth in Part I-Item 1A. “Risk Factors,” could cause actual results to differ materially from forward-looking statements contained in this Annual Report on Form 10-K. We encourage you to read those descriptions carefully. Such statements should be evaluated in light of all the information contained in this document.

AIM[™], ASV[™], AgenTus[™], Agenus[™], AutoSynVax[™], Oncophage[®], PSV[™], PhosphoSynVax[™], Prophage[™], Retrocyte Display[™], SECANT[®] and Stimulon[®] are trademarks of Agenus Inc. and its subsidiaries. All rights reserved.

PART I

Item 1. Business

Our Business

We are a clinical-stage immuno-oncology (“I-O”) company dedicated to becoming a leader in the discovery and development of innovative combination therapies and committed to bringing effective medicines to patients with cancer. Our business is designed to drive success in I-O through speed, innovation, and effective combination therapies. We have assembled fully integrated capabilities from novel target discovery, antibody generation, cell line development, and good manufacturing practice (“GMP”) manufacturing together with a comprehensive portfolio consisting of antibody-based therapeutics, adjuvants and cancer vaccine platforms. We leverage our immune biology platforms to identify effective combination therapies for development and have developed productive partnerships to advance our innovation.

We believe the next generation of cancer treatment will build on clinically validated antibodies targeting CTLA-4 and PD-1 combined with novel immuno-modulatory agents designed to address underlying tumor immune-escape mechanisms.

Our pipeline of immuno-modulatory antibodies target important nodes of immune regulation, including our proprietary lead antibodies, anti-CTLA-4 and anti-PD-1. We are aiming to develop, register, and launch these products with a first potential biologics license application (“BLA”) filing as early as the second-half of 2019. We will then endeavor to expand on these agents with combinations of novel compounds designed to address tumor escape mechanisms, such as intratumoral Treg and tumor microenvironment conditioning.

In addition, for tumors not yet visible to the immune system, we are leveraging our immune educating neoantigen vaccine platform, designed to target mutationally based and biochemically based (phosphorylated) neoantigens (AutoSynVax and PhosPhoSynVax) to prime the immune system to attack tumors. These vaccines may be applicable for patients where checkpoint modulating (“CPM”) antibodies alone are not sufficient to bring about tumor control.

Advances in our understanding of the interactions between cancers, the tumor microenvironment, and the immune system have led to powerful new approaches to treat cancer. We recently formed a new subsidiary, AgenTus Therapeutics, to bring innovative living drugs to cancer patients. AgenTus is employing an integrated platform to discover and develop novel living drugs to treat a broad range of cancers.

To succeed in I-O, innovation and speed are paramount. We are a vertically integrated biotechnology company equipped with a suite of technology platforms to advance from novel target identification (Retrocyte Display, SECANT, Agenus Immunogenic Platform (“AIM”), functional genomics and ligandomics) through manufacturing for clinical trials of antibodies and vaccines.

Our common stock is currently listed on The Nasdaq Capital Market under the symbol “AGEN.”

Our Vision

We believe that harnessing innovation and speed with combinations of drugs are key to bringing effective treatments to patients with certain cancers. We have assembled fully integrated capabilities in novel target discovery, antibody

generation, cell line development, and GMP manufacturing, together with a comprehensive portfolio consisting of antibody-based therapeutics, adjuvants and cancer vaccines. We believe that a balanced pipeline of product candidates should focus on both validated targets as well as novel targets designed to address tumor escape mechanisms. CTLA-4 and PD-1 antagonists are recognized as the first clinically validated immunotherapy combination. These, in combination with innovative immuno-modulatory antibodies or immune education vaccines, could be a focal point of the next generation of I-O combinations. Therefore, we plan to develop, register, and launch, our proprietary PD-1 and CTLA-4 antibody programs aggressively through the clinic and expand with novel combination therapies designed to improve the clinical response and durability response of existing therapies.

Our Strategy

Our strategy is to combine our antibodies, vaccines, and adjuvants to develop effective combinations designed to yield best-in-class treatments for patients with cancer. We are pursuing a tiered risk profile and targeting compressed timelines for regulatory filings. We are executing on a clinical development plan with our anti-CTLA-4 (AGEN1884) and anti-PD-1 (AGEN2034) in definable patient populations. In addition, we plan to pursue select indications to further expedite market entry.

Our combination clinical trial of AGEN1884 with Keytruda in patients with first line non-small cell lung cancer (“1L NSCLC”) with high PD-L1 expression is targeting a definable population indicated for Keytruda monotherapy where combination with our CTLA-4 could potentially expand the clinical benefit beyond Keytruda on its own. Second line cervical cancer is also an indication that is responsive to PD-1 blockade and where combination with CTLA-4 could potentially expand clinical benefit, present a differentiated development path, and potentially provide a niche opportunity in certain markets. Some of our additional programs may pose moderate regulatory risk and will entail: 1) pursuit of effective I-O antibody and vaccine combinations with CTLA-4 and/or PD-

1 targeted antibodies as the backbone; and 2) advancement of our differentiated antibody programs such as our first-in-class bispecific programs, a next generation anti-CTLA-4, and a differentiated CD137 and TIGIT.

Part of our strategy is to develop and commercialize some of our product candidates by continuing our existing arrangements with collaborators and licensees and by entering into new collaborations.

Our Assets

Our I-O assets include antibody-based therapeutics, cancer vaccine platforms, and adjuvants. Our proprietary CTLA-4 and PD-1 antagonists are in clinical development; we are one of the few companies to have a proprietary anti-CTLA-4 and anti-PD-1 in clinical combinations.

To complement our portfolio of foundational CPMs, we have pre-clinical antibodies targeting novel immune-mechanisms. These include next-generation anti-CTLA-4, CD137 and anti-TIGIT antibodies, as well as undisclosed multi-specific antibodies.

We have proprietary cancer vaccine platforms which are designed to be autologous (Prophage) and individualized (AutoSynVax (“ASV”); PhosphoSynVax (“PSV”)). Our Prophage and synthetic ASV and PSV vaccine candidates are protein complexes that consist of heat shock proteins (“HSPs”) and peptides that are either tumor-derived or tailor-made based on the unique genomic fingerprint of a patient’s tumor, respectively. Our vaccine programs are being developed for intended combinations with CPMs.

Our QS-21 Stimulon adjuvant is partnered with GlaxoSmithKline plc. (“GSK”) and is a key component in multiple GSK vaccine programs. GSK's Shingrix vaccine, which contains our QS-21 Stimulon® adjuvant, is approved by the US Food and Drug Administration (“FDA”) and Health Canada and recently received the Committee for Medicinal Products for Human Use recommendation for approval.

Our Antibody Discovery Platforms and CPM Programs

Checkpoint antibodies regulate immune response against pathogens that invade the body and are achieving positive outcomes in a number of cancers that were untreatable only a few years ago. Two classes of checkpoint targets include:

1. inhibitory checkpoints that help suppress an immune response in order to prevent excessive immune reaction resulting in undesired inflammation and/or auto-immunity, and
2. stimulatory checkpoints that can enhance or amplify an antigen-specific immune response.

We possess a suite of antibody discovery platforms that are designed to drive the discovery of future CPM antibody candidates. We are planning to employ a variety of techniques to identify and optimize mono-specific and multi-specific antibody candidates, internally.

We have presented clinical data on our lead antibody, AGEN1884, at the American Society of Clinical Oncology (“ASCO”) in June 2017. We have also presented pre-clinical and clinical data on our anti-CTLA-4 (AGEN1884), anti-PD-1 (AGEN2034), and ASV programs as well as our partnered programs GITR (INCAGN1876) and OX-40 (INCAGN1949) at the American Association for Cancer Research (“AACR”) April 2017, Society for Immunotherapy of Cancer (“SITC”) November 2017, the International Cancer Immunotherapy Conference (“CIMT”) May 2017, and CRI-CIMT-EATI-AACR International Cancer Immunotherapy Conference in September 2017. The presentations covered pre-clinical pharmacology for our antibodies and demonstrated that AGEN1884 binds to CTLA-4 expressed on T cells and potently blocks engagement of CD80 and CD86, leading to enhanced T cell responsiveness. We also reported:

• data that AGEN1884 augmented vaccine response in primates;

- clinical data on safety and preliminary efficacy of AGEN1884, which supported our expectations of the safety profile;
- that we observed a complete response in a compassionate use setting in a patient with advanced angiosarcoma, who was unresponsive to prior therapies;
- data demonstrating that AGEN2034, a novel human IgG4 anti-PD-1 antagonist antibody, inhibits PD-1 binding to PD-L1 and PD-L2, resulting in enhanced T cell responsiveness in vitro as well as in a non-human primate model;
- that AGEN2034 combined effectively with AGEN1884, a human IgG1 anti-CTLA-4 antibody, anti-TIGIT or anti-LAG-3 to further enhance T cell responsiveness;
- that AIM can predict immunogenic peptides containing neoantigens and tumor specific phosphopeptides to develop immunogenic vaccines; and
- clinical data demonstrating ASV can effectively deliver antigens and induce an anti-tumor immunogenic response and pre-clinical data revealing that these vaccinogenic responses are optimized with combination checkpoint modulating antibodies.

We are developing our anti-CTLA-4 antibody in combination with Keytruda in a definable cohort of patients in 1L NSCLC with high PD-L1 (>50%) expression, where Keytruda is approved for use as a monotherapy with <50% response rates. We also plan to develop our proprietary anti-CTLA-4 antibody in combination with our proprietary anti-PD-1 antibody in second line cervical cancer. Chemoradiation therapy is the current standard of care for earlier lines of treatment. In distant metastatic patients, platinum-based chemotherapy, with or without bevacizumab, is the current standard of care. However, there are no established therapies for second

line cervical cancer and the five-year survival rate of recurrent/metastatic cervical cancer is 16.8%. Cervical cancer is a malignancy that is driven by the persistent infection by certain types of human papilloma virus (“HPV”). Anti PD-1/PD-L1 have shown to be active in virally induced disease and, specifically, HPV induced squamous cell cancer of the head and neck. In these tumors, anti PD-1 blockade might induce objective responses as well as prolongation of survival.

In addition to pursuing validated targets, our discovery pipeline includes next generation molecules to known targets (such as TIGIT and CD137) as well as potential first-in-class bispecific antibodies designed to go beyond T cell targeting to address tumor escape mechanisms within the tumor micro-environment. The most advanced of our discovery pipeline is our next generation CTLA-4, an IgG1 anti-CTLA-4 antagonist. This molecule is advancing through Investigational New Drug (“IND”) enablement.

Partnered CPM Programs

In January 2015, we entered into a broad, global alliance with Incyte Corporation (“Incyte”) to discover, develop and commercialize novel immuno-therapeutics using our antibody platforms. The collaboration was initially focused on four CPM programs targeting GITR, OX40, TIM-3 and LAG-3, and in November 2015, we expanded the alliance by adding three novel undisclosed CPM targets. Pursuant to the terms of the original agreement, Incyte made non-creditable, non-refundable upfront payments to us totaling \$25.0 million. Targets under the collaboration were designated as either profit-share programs, where the parties shared all costs and profits equally, or royalty-bearing programs, where Incyte funded all costs, and we were eligible to receive milestones and royalties. Under the original collaboration agreement, programs targeting GITR, OX40 and two of the undisclosed targets were designated as profit-share programs, while the other targets were royalty-bearing programs. For each profit-share product, we were eligible to receive up to \$20.0 million in future contingent development milestones. For each royalty-bearing product, we were eligible to receive (i) up to \$155.0 million in future contingent development, regulatory, and commercialization milestones and (ii) tiered royalties on global net sales at rates generally ranging from 6%-12%. Concurrent with the execution of the original collaboration agreement, we and Incyte also entered into a stock purchase agreement pursuant to which Incyte purchased approximately 7.76 million shares of our common stock for an aggregate purchase price of \$35.0 million, or approximately \$4.51 per share. In February 2017, we and Incyte amended the terms of the original collaboration agreement to, among other things, convert the GITR and OX40 programs from profit-share to royalty-bearing programs with royalties on global net sales at a flat 15% rate for each. In addition, the profit-share programs relating to two undisclosed targets were removed from the collaboration, with one reverting to Incyte and one to Agenus (the latter being our TIGIT antibody program), each with royalties on global net sales at a flat 15% rate. The remaining three royalty-bearing programs in the collaboration targeting TIM-3, LAG-3 and one undisclosed target remain unchanged, and there are no more profit-share programs under the collaboration. Pursuant to the amended agreement, we received accelerated milestone payments of \$20.0 million from Incyte related to the clinical development of INCAGN1876 (anti-GITR agonist) and INCAGN1949 (anti-OX40 agonist). Across all programs in the collaboration, we are now eligible to receive up to a total of \$510.0 million in future potential development, regulatory and commercial milestones. Concurrent with the execution of the amendment agreement, we and Incyte entered into a separate stock purchase agreement whereby Incyte purchased an additional 10 million shares of Agenus common stock at \$6.00 per share. Immediately following the transaction, Incyte owned approximately 18.1% of our outstanding common stock.

At the April 2016 AACR conference, we presented data for two antibody candidates under the Incyte collaboration: INCAGN1949 (anti-OX40 agonist) and INCAGN1876 (anti-GITR agonist). The presentations covered pre-clinical pharmacology for each antibody, including optimized features. At AACR 2017, we presented additional pre-clinical data for both INCAGN1949 and INCAGN1876 which further characterized these antibody candidates. In June 2016, we announced that the first patient was dosed in a Phase 1/2 clinical trial of INCAGN1876. The Phase 1/2 trial is exploring the safety, tolerability, and efficacy of INCAGN1876 combined with immune therapies, ipilimumab and nivolumab, in advanced or metastatic malignancies such as advanced or metastatic endometrial cancer, gastric cancer (including stomach, esophageal, and gastroesophageal junction), and squamous cell carcinoma of the head and neck.

In addition, in November 2016 we announced the commencement of a Phase 1/2 clinical trial of INCAGN1949. The Phase 1/2 trial is exploring the safety, tolerability, and efficacy of INCAGN1949 combined with immune therapies, ipilimumab and nivolumab, in advanced or metastatic malignancies such as advanced or metastatic urothelial carcinoma or RCC.

In addition, in April 2014, we entered into a collaboration and license agreement with Merck to discover and optimize fully-human antibodies against two undisclosed CPM targets. Merck selected a lead product candidate against one of the undisclosed Merck targets to advance into preclinical studies, leading to a \$2.0 million milestone payment that we received in May 2016. In November 2017, we announced we had received a \$4.0 million payment for the advancement of the undisclosed antibody under our license and research collaboration agreement with Merck. Under the terms of the agreement, Merck is responsible for all future product development expenses for the selected antibody candidate. We are eligible to receive up to an additional \$99.0 million in milestone payments, in addition to royalties on any worldwide product sales.

In 2017, we also formalized a research collaboration with UCB Biopharma SPRL (“UCB”). The collaboration leverages the antibody engineering capabilities of UCB and Agenus in novel bispecific antibody discovery. In addition, we have a collaboration agreement with Recepta Biopharma SA on the development of our antibodies targeting CTLA-4 and PD-1, which gives Recepta certain rights to South American countries. We expect to continue exploring additional future collaborations.

Vaccine Platforms

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Our current vaccine platforms for the treatment of cancer, and potentially other indications, include our HSP based Prophage vaccine candidates, and our fully synthetic, neoantigen vaccine candidates, ASV and PSV.

HSPs are a group of proteins present at high levels in most mammalian cells. Their expression is increased when cells are exposed to elevated temperatures or other stresses. A potential role for HSPs in regulating immune responses was revealed when it was first discovered that HSP complexes purified from cancer cells produced immunity to cancer, whereas HSP complexes purified from normal tissue did not. This discovery led to the understanding that HSPs bind to and carry a broad sampling of the protein environment within cells, including mutant proteins that might arise from genetic mutations within cancer cells. It was further shown that immunization with HSP complexes purified from tumors generate both CD4 and CD8 positive T-cell immune responses. These activated T-cells target the cancer cells of the tumor, from which the HSP complexes were derived, for destruction. Thus, HSP complexes isolated from cancer cells may be particularly helpful in mediating successful immunization. Since HSPs are expressed in all tumor cells, the approach of immunizing with the HSP complexes isolated from a particular tumor may be broadly applicable to a variety of cancer types. We believe that we pioneered the use of gp96, an HSP, purified from a patient's own tumor tissue, as a way to make I-O vaccine candidates.

Prophage Vaccine Candidates

Our Prophage cancer vaccine candidate, HSPPC-96, is an autologous cancer vaccine therapy derived from cancer tissues that are surgically removed from an individual patient. As a result, a Prophage vaccine contains a broad sampling of potentially antigenic mutant proteins from each patient's own tumor. Prophage vaccines are designed to program the body's immune system to target only the specific cells that express those mutant antigens, thereby reducing the risk that the body's immune response against the tumor after vaccination will also affect healthy tissue. Enhancing immune response using personalized vaccines, particularly in combination with CPMs, could be useful in cancers where a low number of mutant proteins leads to weakened immunogenicity.

To date, more than 1,000 patients have been treated with Prophage vaccines in clinical trials internationally, covering a broad range of cancer types, and no serious immune-mediated side effects have been observed. The results of these trials have been published and/or presented at scientific conferences. These results indicate observable clinical and/or immunological activity across many types of cancer.

In January 2017, we announced a clinical trial collaboration with the National Cancer Institute ("NCI"). The double-blind, randomized controlled Phase 2 trial is evaluating the effect of Prophage in combination with pembrolizumab (Keytruda®) in patients with ndGBM. The trial is being conducted by the Brain Tumor Trials Collaborative ("BTTC"), a consortium of top academic centers led by Dr. Mark Gilbert, Chief of the Neuro-Oncology Branch at the NCI Center for Cancer Research. The trial consists of two-arms with one arm receiving pembrolizumab as a monotherapy and a second arm receiving both Prophage and pembrolizumab in combination. Forty-five patients are being randomly assigned to each arm. Under this collaboration, we are supplying Prophage, Merck is supplying pembrolizumab (Keytruda®) and NCI and BTTC member sites are recruiting patients and conducting the trial.

Neoantigen Vaccine Platforms

Mutation-based neo-epitopes, which will form the basis for the immunogens used in ASV, appear to be almost always particular to a given patient. Therefore, ASV is a largely individualized vaccine product candidate. With a small amount of a patient's tumor as a sample, our ASV program is designed to utilize next generation sequencing technologies coupled with complex bioinformatics algorithms to identify mutations in a tumor's DNA and RNA. Once these mutations have been identified, we plan to manufacture synthetic peptides encoding these neoepitopes, load these peptides on to our recombinant HSP70 and deliver a fully synthetic polyvalent vaccine to the patient. This program is based on the hypothesis that the HSP70 platform would shuttle the mutated peptides to sites where they could be recognized by the immune system and elicit a cytotoxic and helper T cell response in patients with cancer.

Biochemically based neoantigens, such as those arising from dysregulated phosphorylation of various proteins in malignant cells, can serve as a tumor fingerprint across a broad histology of tumors. Through the acquisition of PhosImmune, we have a portfolio of proprietary phosphorylated antigenic targets that can be used for therapeutic development. PSV is a vaccine candidate designed to induce immunity against this novel class of tumor specific neoepitopes. PSV is intended to induce cellular immunity to abnormal phosphopeptide neoepitopes that are characteristic of these various forms of cancer. Phosphopeptides (or phosphopeptide analogues) can be synthesized and complexed with HSP70, using an approach analogous to that used in the generation of our previous HerpV vaccine candidate. HerpV demonstrated good cellular and humoral responses to synthetic peptide immunogens complexed with HSP70 in a placebo-controlled Phase 2 study. We believe that similar responses could be obtained to phosphopeptide or phosphopeptide analogues bound to HSP70 when used as vaccines. Phosphorylation-based neoepitopes can apparently be found on specific types of cancer in many patients. Studies to optimize the immunogens to be used in PSV are ongoing.

QS-21 Stimulon Adjuvant

QS-21 Stimulon is an adjuvant, which is a substance added to a vaccine or other immunotherapy that is intended to enhance an immune response to the target antigens. QS-21 Stimulon is a natural product, a triterpene glycoside, or saponin, purified from the bark of the Chilean soapbark tree, *Quillaja saponaria*. QS-21 Stimulon has the ability to stimulate an antibody-mediated immune response and has also been shown to activate cellular immunity. It has become a key component in the development of investigational

preventive vaccine formulations across a wide variety of diseases. These studies have been carried out by academic institutions and pharmaceutical companies in the United States and internationally. A number of these studies have shown QS-21 Stimulon to be significantly more effective in stimulating immune responses than aluminum hydroxide or aluminum phosphate, the adjuvants most commonly used in approved vaccines in the United States today.

Partnered QS-21 Stimulon Programs

In July 2006, we entered into a license agreement and a supply agreement with GSK for the use of QS-21 Stimulon (the “GSK License Agreement” and the “GSK Supply Agreement,” respectively). In January 2009, we entered into an Amended and Restated Manufacturing Technology Transfer and Supply Agreement (the “Amended GSK Supply Agreement”) under which GSK has the right to manufacture all of its requirements of commercial grade QS-21 Stimulon. GSK is obligated to supply us, or our affiliates, licensees, or customers, certain quantities of commercial grade QS-21 Stimulon for a stated period of time. In March 2012, we entered into a First Right to Negotiate and Amendment Agreement amending the GSK License Agreement and the Amended GSK Supply Agreement to clarify and include additional rights for the use of QS-21 Stimulon (the “GSK First Right to Negotiate Agreement”). In addition, we granted GSK the first right to negotiate for the purchase of Agenus or certain of our assets, which expired in March 2017. As consideration for entering into the GSK First Right to Negotiate Agreement, GSK paid us an upfront, non-refundable payment of \$9.0 million, \$2.5 million of which is creditable toward future royalty payments. We refer to the GSK License Agreement, the Amended GSK Supply Agreement and the GSK First Right to Negotiate Agreement collectively as the GSK Agreements. In 2017, we received a final milestone payment of \$1.0 million from GSK and are no longer entitled to any additional milestone payments under the GSK Agreements. Under the terms of the Agreement, we are generally entitled to receive 2% royalties on net sales of prophylactic vaccines for a period of 10 years after the first commercial sale of a resulting GSK product, with some exceptions; however, we have already sold this potential royalty stream in its entirety as discussed in more detail below. The GSK License and Amended GSK Supply Agreements may be terminated by either party upon a material breach if the breach is not cured within the time specified in the respective agreement. The termination or expiration of the GSK License Agreement does not relieve either party from any obligation which accrued prior to the termination or expiration. Among other provisions, the license rights granted to GSK survive expiration of the GSK License Agreement. The license rights and payment obligations of GSK under the Amended GSK Supply Agreement survive termination or expiration, except that GSK's license rights and future royalty obligations do not survive if we terminate due to GSK's material breach unless we elect otherwise.

QS-21 Stimulon is a key component included in certain of GSK's proprietary adjuvant systems, and we believe that a number of GSK's vaccine candidates currently in development are formulated using adjuvant systems containing QS-21 Stimulon, including its shingles vaccine that was approved by the FDA and Health Canada in 2017. We do not incur clinical development costs for products partnered with GSK.

In September 2015, we monetized a portion of the royalties associated with the GSK License Agreement to an investor group led by Oberland Capital Management for up to \$115.0 million in the form of a non-dilutive royalty transaction. Under the terms of a Note Purchase Agreement with the investor group (the “Note Purchase Agreement”) we received \$100.0 million at closing for which the investors had the right to receive 100% of our worldwide royalties under the GSK License Agreement on sales of GSK's shingles (HZ/su) and malaria (RTS,S) prophylactic vaccine products that contain our QS-21 Stimulon adjuvant to pay down principle and interest. In November 2017 and pursuant to the Note Purchase Agreement, we received an additional \$15.0 million in cash from the investors based on the approval of HZ/su by the FDA. Pursuant to the terms of this transaction, we retained the right to receive all royalties from GSK after all principal, interest and other obligations were satisfied under the Note Purchase Agreement. The Note Purchase Agreement also allowed us to buy back the loan and extinguish the notes early under pre-specified terms.

In January 2018, we sold 100% of all royalties we were entitled to receive from GSK to Healthcare Royalty Partners III, L.P. and certain of its affiliates (collectively, “HCR”) and used the proceeds to extinguish the debt under the Note

Purchase Agreement. HCR paid approximately \$190.0 million at closing for the royalty rights, of which approximately \$161.9 was used to extinguish the prior notes, yielding us approximately \$28.0 million in net proceeds. We are also entitled to receive up to \$40.35 million in milestone payments from HCR based on sales of GSK's vaccines as follows: (i) \$15.1 million upon reaching \$2.0 billion last-twelve-months net sales any time prior to 2024 and (ii) \$25.25 million upon reaching \$2.75 billion last-twelve-months net sales any time prior to 2026. We would owe a reverse milestone payment of approximately \$25.9 million to HCR in 2021 if neither of the following sales milestones are achieved: (i) 2019 sales of the GSK vaccines exceed \$1.0 billion or (ii) 2020 sales of the GSK vaccines exceed \$1.75 billion (the "Rebate Payment"). As part of the transaction, we provided a guaranty for the potential Rebate Payment and secured the obligation with substantially all of our assets pursuant to a security agreement, subject to certain customary exceptions and excluding all assets necessary for AgenTus Therapeutics, Inc.

Manufacturing

Manufacturing CPM Antibodies

In December 2015, Agenus acquired an antibody manufacturing pilot plant in Berkeley, CA from XOMA Corporation ("XOMA"), which we refer to as "Agenus West." A team of former XOMA employees with valuable chemistry, manufacturing and controls experience joined us and continue to operate the facility. In addition, in February 2017, we amended our collaboration with

Incyte, transferring manufacturing responsibilities for all antibodies under the collaboration to them. This includes antibodies targeting GITR, OX40, TIM-3, LAG-3 and one undisclosed target. We have transferred the manufacturing know how to Incyte to support these endeavors and allow Agenus to focus on the manufacture of antibodies for some of our own CPM programs and those of existing and potential third-party collaborators. Since the acquisition of Agenus West, we have made significant improvements in the plant, and added additional headcount increasing both scale and capacity. The plant is currently providing antibody production for our lead programs. We aim to utilize our Agenus West pilot plant capabilities to accelerate antibody delivery, improve quality and increase product yield while providing us with greater manufacturing flexibility, all at reduced costs.

Manufacturing Cancer Vaccines

We manufacture our cancer vaccine candidates in our Lexington, MA facility.

Each Prophage vaccine is manufactured using a patient's own tumor. After the patient undergoes surgery to remove cancerous tumor tissue, the tumor is shipped frozen in a specially designed kit provided by us to our Lexington, Massachusetts facility. Each Prophage vaccine is produced to a specific standard, in a process taking approximately ten hours, after which it undergoes extensive quality testing for approximately two weeks. The turnaround time from the date of surgery to delivery of vaccine is approximately three to four weeks, which generally fits well with the patient's recovery time from surgery. Once we release the vaccine, it is shipped frozen overnight to the hospital pharmacy or clinician. Prophage vaccines are given as a simple intradermal injection.

ASV and PSV vaccine candidates are manufactured using HSP70 loaded with synthetic peptide synthesized by approved manufacturers. The sequence of the peptides is determined by sequencing and analysis of patient and tumor DNA and RNA and run through complex algorithms by our bioinformatics group who have specialized knowledge of the attributes required to elicit immune responsiveness. This process takes several weeks, after which the manufactured vaccine undergoes extensive quality testing, including sterility testing, which takes a further two weeks.

We have established, within a single facility, well-defined, cost efficient vaccine manufacturing under GMPs, including bioanalytical, quality control and quality assurance, logistics, distribution and supply chain management. After manufacturing, Prophage and ASV vaccine candidates are tested and released by our analytical and quality systems staff. The quality control organization performs a series of release assays designed to ensure that the product meets all applicable specifications. Our quality assurance staff also reviews manufacturing and quality control records prior to batch release in an effort to assure conformance with current GMP ("cGMP") as mandated by the FDA and foreign regulatory agencies.

Our manufacturing staff is trained and routinely evaluated for conformance to rigorous manufacturing procedures and quality standards. This oversight is intended to ensure compliance with FDA and foreign regulations and to provide consistent vaccine output. Our quality control and quality assurance staff is similarly trained and evaluated as part of our effort to ensure consistency in the testing and release of the product, as well as consistency in materials, equipment and facilities.

QS-21 Stimulon

Except in the case of GSK, we have retained worldwide manufacturing rights for QS-21 Stimulon, and we have the right to subcontract manufacturing for QS-21 Stimulon. In addition, under the terms of our agreement with GSK, upon request by us, GSK is committed to supply certain quantities of commercial grade QS-21 Stimulon to us and our licensees for a fixed period of time.

Intellectual Property Portfolio

We seek to protect our technologies through a combination of patents, trade secrets and know-how, and we currently own, co-own or have exclusive rights to approximately 35 issued United States patents and approximately 120 issued foreign patents. We also own, co-own or have exclusive rights to approximately 30 pending United States patent applications and approximately 125 pending foreign patent applications. We may not have rights in all territories where we may pursue regulatory approval for our product candidates.

Through various acquisitions, we own, co-own, or have exclusive rights to a number of patents and patent applications directed to various methods and compositions, including methods for identifying therapeutic antibodies and product candidates arising out of such entities' technology platforms. In particular, we own patents and patent applications relating to our Retrocyte Display technology platform, a high throughput antibody expression platform for the identification of fully-human and humanized monoclonal antibodies. This patent family is projected to expire between 2029 and 2031. We own, co-own, or have exclusive rights to patents and patent applications directed to various methods and compositions, including a patent directed to methods for identifying phosphorylated proteins using mass spectrometry. This patent is projected to expire in 2023. We also own patents and patent applications relating to the SECANT[®] platform, a platform used for the generation of novel monoclonal antibodies. This patent family is projected to expire between 2028 and 2029. In addition, as we advance our research and development efforts with our institutional and corporate collaborators, we are seeking patent protection for certain newly identified therapeutic antibodies and product candidates. We can provide no assurance that any of our patents, including the patents that we acquired or in-licensed, will have commercial value, or that any of our existing or future patent applications, including the patent applications that were acquired or in-licensed, will result in the issuance of valid and enforceable patents. Our issued patents covering Prophage vaccine and methods of use thereof, alone or in combination with other agents, expired or will expire at various dates between 2015 and 2024. In particular, our issued U.S. patents

covering Prophage composition of matter expired in 2015. In addition, our issued patents covering QS-21 Stimulon composition of matter expired in 2008. We continue to explore means of extending the life cycle of our patent portfolio.

Various patents and patent applications have been exclusively licensed to us by the following entities:

University of Virginia

In connection with our acquisition of PhosImmune in December 2015, we obtained exclusive rights to a portfolio of patent applications and one issued patent relating to PTTs under a patent license agreement with the University of Virginia (“UVA”). The UVA license gives us exclusive rights to develop and commercialize the PTT technology and an exclusive option to license any further PTT technology arising from ongoing research at UVA until December 2018. Under the license agreement, we will pay low to mid-single digit running royalties on net sales of PTT products, and a modest flat percentage of sublicensing income. In addition, we may be obligated to make milestone payments of up to \$2.7 million for each indication of a licensed PTT product to complete clinical trials and achieve certain sales thresholds. The term of the UVA license agreement ends when the last of the licensed patents expires or becomes no longer valid. The UVA license agreement may be terminated as follows: (i) by UVA in connection with our bankruptcy or cessation of business relating to the licensed technology, (ii) by UVA if we commit a material, uncured breach or (iii) by us for our convenience on 180 days written notice.

Ludwig Institute for Cancer Research

On December 5, 2014, our wholly-owned subsidiary, Agenus Switzerland Inc. (formerly known as 4-Antibody AG)(“4-AB”), entered into a license agreement with the Ludwig Institute for Cancer Research Ltd. (“Ludwig”), which replaced and superseded a prior agreement entered into between the parties in May 2011. Pursuant to the terms of the license agreement, Ludwig granted 4-AB an exclusive, worldwide license under certain intellectual property rights of Ludwig and Memorial Sloan Kettering Cancer Center arising from the prior agreement to further develop and commercialize GITR, OX40 and TIM-3 antibodies. On January 25, 2016, we and 4-AB entered into a second license agreement with Ludwig, on substantially similar terms, to develop CTLA-4 and PD-1 antibodies. Pursuant to the December 2014 license agreement, 4-AB made an upfront payment of \$1.0 million to Ludwig. The December 2014 license agreement also obligates 4-AB to make potential milestone payments of up to \$20.0 million for events prior to regulatory approval of licensed GITR, OX40 and TIM-3 products, and potential milestone payments in excess of \$80.0 million if such licensed products are approved in multiple jurisdictions, in more than one indication, and certain sales milestones are achieved. Under the January 2016 license agreement, we are obligated to make potential milestone payments of up to \$12.0 million for events prior to regulatory approval of CTLA-4 and PD-1 licensed products, and potential milestone payments of up to \$32.0 million if certain sales milestones are achieved. Under each of these license agreements, we and/or 4-AB will also be obligated to pay low to mid-single digit royalties on all net sales of licensed products during the royalty period, and to pay Ludwig a percentage of any sublicensing income, ranging from a low to mid-double digit percentage depending on various factors. During the year ended December 31, 2017, we paid a percentage of sublicensing income totaling \$2.0 million to Ludwig under the license agreements. The license agreements may each be terminated as follows: (i) by either party if the other party commits a material, uncured breach; (ii) by either party if the other party initiates bankruptcy, liquidation or similar proceedings; or (iii) by 4-AB or us (as applicable) for convenience upon 90 days’ prior written notice. The license agreements also contain customary representations and warranties, mutual indemnification, confidentiality and arbitration provisions.

University of Connecticut Health Center

In May 2001, we entered into a license agreement with the University of Connecticut Health Center (“UConn”) which was amended in March 2003 and June 2009. Through the license agreement, we obtained an exclusive, worldwide license to patent rights resulting from inventions discovered under a research agreement that was effective from February 1998 until December 2006. The term of the license agreement ends when the last of the licensed patents

expires in 2028 or becomes no longer valid. UConn may terminate the agreement: (1) if, after 30 days written notice for breach, we continue to fail to make any payments due under the license agreement, or (2) we cease to carry on our business related to the patent rights or if we initiate or conduct actions in order to declare bankruptcy. We may terminate the agreement upon 90 days written notice. We are required to make royalty payments on any obligations created prior to the effective date of termination of the license agreement. Upon expiration or termination of the license agreement due to breach, we have the right to continue to manufacture and sell products covered under the license agreement which are considered to be works in progress for a period of six months. The license agreement contains aggregate milestone payments of approximately \$1.2 million for each product we develop covered by the licensed patent rights. These milestone payments are contingent upon regulatory filings, regulatory approvals, and commercial sales of products. We have also agreed to pay UConn a royalty on the net sales of products covered by the license agreement as well as annual license maintenance fees beginning in May 2006. Royalties otherwise due on the net sales of products covered by the license agreement may be credited against the annual license maintenance fee obligations. Under the March 2003 amendment, we agreed to pay UConn an upfront payment and to make future payments for each patent or patent application with respect to which we exercised our option under the research agreement. As of December 31, 2017, we had paid approximately \$900,000 to UConn under the license agreement. The license agreement gives us complete discretion over the commercialization of products covered by the licensed patent rights but also requires us to use commercially reasonable diligent efforts to introduce commercial products within and outside the United States. If we fail to meet these diligence requirements, UConn may be able to terminate the license agreement.

Regulatory Compliance

Governmental authorities in the United States and other countries extensively regulate the pre-clinical and clinical testing, manufacturing, labeling, storage, record keeping, advertising, promotion, export, marketing and distribution, among other things, of our investigational product candidates. In the United States, the FDA under the Federal Food, Drug, and Cosmetic Act, the Public Health Service Act and other federal statutes and regulations, subject pharmaceutical products to rigorous review.

In order to obtain approval of a new product from the FDA, we must, among other requirements, submit proof of safety and efficacy as well as detailed information on the manufacture and composition of the product. In most cases, this proof entails extensive pre-clinical, clinical, and laboratory tests. Before approving a new drug or marketing application, the FDA may also conduct pre-licensing inspections of the company, its contract research organizations and/or its clinical trial sites to ensure that clinical, safety, quality control, and other regulated activities are compliant with Good Clinical Practices (“GCP”), or Good Laboratory Practices (“GLP”), for specific non-clinical toxicology studies. The FDA may also require confirmatory trials, post-marketing testing, and extra surveillance to monitor the effects of approved products, or place conditions on any approvals that could restrict the commercial applications of these products. Once approved, the labeling, advertising, promotion, marketing, and distribution of a drug or biologic product must be in compliance with FDA regulatory requirements.

In Phase 1 clinical trials, the sponsor tests the product in a small number of patients or healthy volunteers, primarily for safety at one or more doses. Phase 1 trials in cancer are often conducted with patients who have end-stage or metastatic cancer. In Phase 2, in addition to safety, the sponsor evaluates the efficacy of the product in a patient population somewhat larger than Phase 1 trials. Phase 3 trials typically involve additional testing for safety and clinical efficacy in an expanded population at geographically dispersed test sites. The FDA may order the temporary or permanent discontinuation of a clinical trial at any time.

The sponsor must submit to the FDA the results of pre-clinical and clinical testing, together with, among other things, detailed information on the manufacture and composition of the product, in the form of a new drug application (“NDA”), or in the case of biologics, a BLA. In a process that can take a year or more, the FDA reviews this application and, when and if it decides that adequate data are available to show that the new compound is both safe and effective for a particular indication and that other applicable requirements have been met, approves the drug or biologic for marketing.

Whether or not we have obtained FDA approval, we must generally obtain approval of a product by comparable regulatory authorities of international jurisdictions prior to the commencement of marketing the product in those jurisdictions. We are also subject to cGMP, GCP, and GLP compliance obligations and are subject to inspection by international regulatory authorities. International requirements may in some circumstances be more rigorous than U.S. requirements and may require additional investment in manufacturing process development, non-clinical studies, clinical studies, and record-keeping that are not required for U.S. regulatory compliance or approval. The time required to obtain this approval may be longer or shorter than that required for FDA approval and can also require significant resources in time, money and labor.

Under the laws of the United States, the countries of the European Union and other nations, we and the institutions where we sponsor research are subject to obligations to ensure the protection of personal information of human subjects participating in our clinical trials. We have instituted procedures that we believe will enable us to comply with these requirements and the contractual requirements of our data sources. The laws and regulations in this area are evolving, and further regulation, if adopted, could affect the timing and the cost of future clinical development activities.

We are also subject to regulation under the Occupational Safety and Health Act, the Toxic Substances Control Act, the Resource Conservation and Recovery Act, and other current and potential future federal, state, or local regulations.

Our research and development activities involve the controlled use of hazardous materials, chemicals, biological materials, various radioactive compounds, and for some experiments we use recombinant DNA. We believe that our procedures comply with the standards prescribed by local, state, and federal regulations; however, the risk of injury or accidental contamination cannot be completely eliminated. We conduct our activities in compliance with the National Institutes of Health Guidelines for Recombinant DNA Research.

Additionally, the U.S. Foreign Corrupt Practices Act (“FCPA”), prohibits U.S. corporations and their representatives from offering, promising, authorizing or making payments to any foreign government official, government staff member, political party or political candidate in an attempt to obtain or retain business abroad. The scope of the FCPA includes interactions with certain healthcare professionals in many countries. Other countries have enacted similar anti-corruption laws and/or regulations.

Competition

Competition in the pharmaceutical and biotechnology industries is intense. Many pharmaceutical or biotechnology companies have products on the market and are actively engaged in the research and development of products for the treatment of cancer.

Many competitors have substantially greater financial, manufacturing, marketing, sales, distribution, and technical resources, and more experience in research and development, clinical trials, and regulatory matters, than we do. Competing companies developing or acquiring rights to more efficacious therapeutic products for the same diseases we are targeting, or which offer significantly lower costs of treatment, could render our products noncompetitive or obsolete. See Part I-Item 1A. “Risk Factors-Risks

Related to our Business-Our competitors may have superior products, manufacturing capability, selling and marketing expertise and/or financial and other resources.”

Academic institutions, governmental agencies, and other public and private research institutions conduct significant amounts of research in biotechnology, medicinal chemistry and pharmacology. These entities have become increasingly active in seeking patent protection and licensing revenues for their research results. They also compete with us in recruiting and retaining skilled scientific talent.

The CPM landscape is crowded with several competitors developing assets against a number of targets. Development plans are spread out across various indications and lines of therapy, either alone or in combination with other assets. Competitors range from small cap to large cap companies, with assets in pre-clinical or clinical stages of development. Therefore, the landscape is dynamic and constantly evolving. We and our partners have CPM antibody programs currently in clinical stage development targeting PD-1, CTLA-4, GITR and OX40. We are aware of many companies that have antibody-based products on the market or in clinical development that are directed to the same biological targets as these programs, including, without limitation, the following: (1) Bristol-Myers Squibb (“BMS”) markets ipilimumab, an anti-CTLA-4 antibody, and nivolumab, an anti-PD-1 antibody, and is developing agonists to GITR and OX-40, (2) Merck has an approved anti-PD-1 antibody in the United States, as well as an anti-CTLA-4 antagonist and an anti-GITR agonist in clinical development, (3) Ono Pharmaceuticals has an approved anti-PD-1 antibody in Japan, (4) AstraZeneca has an approved anti PD-L1 antibody, as well as anti-CTLA-4, PD-1, GITR and OX40 targeting antibodies in development, (5) Pfizer has an approved anti-PD-L1 (with Merck KgaA) as well as anti-PD-1, and anti-OX40 antibodies in clinical development, (6) Novartis has anti-PD-1, anti-PD-L1 and anti-GITR antibodies in clinical trials, and (7) Roche/Genentech has an approved anti-PD-L1 antibody. We are also aware of other competitors with PD-1/PD-L1 antibodies in clinical development, including AbbVie, Arcus Biosciences, Boehringer Ingelheim, Tesaro, Beigene, Regeneron, Eli Lilly, Jiangsu HengRui Medicine, Shanghai Junshi, MacroGenics, CytomX, Janssen, CBT Pharmaceuticals, Checkpoint Therapeutics, CStone Pharmaceuticals, Livzon MabPharm Inc and Suzhou Alphamab. We are also aware of competitors with pre-clinical antibodies against these targets. In addition, we are also aware of competitors with clinical stage antibodies against targets in our earlier stage programs such as TIM-3, LAG-3, CD137, TIGIT and other undisclosed targets. These include, but are not limited to, BMS, Pfizer, Novartis, Merck, Roche, Tesaro, Eli Lilly, OncoMed, Boehringer Ingelheim, Astellas and Regeneron. Additionally, we are also aware of competitors with assets against these targets that are in pre-clinical development. There is no guarantee that our antibody product candidates will be able to compete with our competitors’ antibody products and product candidates.

We are planning to advance combinations of our anti-CTLA-4 antibody, AGEN1884, with Keytruda in 1L NSCLC, where Keytruda is currently approved. Specifically, we will be evaluating efficacy within a subset of patients who express $\geq 50\%$ PD-L1 protein biomarker levels. We are aware of competitors who have approved immunotherapy products in this indication, including having a label specifically directed at this patient subset, such as Merck (anti-PD-1 monotherapy). We are also aware of industry sponsored clinical trials, including exploratory studies, that are directed at this specific patient population. These include, but are not restricted to, Incyte/Merck (anti-IDO + anti-PD-1), Merck (anti-PD-1 + anti-CTLA-4), Regeneron (anti-PD-1), Roche (anti-PD-L1), Merck KgaA / Pfizer (anti-PD-L1), and Tesaro (anti-PARP + anti-PD-1). We are also aware of competitors who have approved immunotherapies or have published positive Phase 3 data for immunotherapies in 1L NSCLC. These include, and are not restricted to, Merck, BMS, and Roche. Additional competitors have ongoing clinical trials for immunotherapies in the 1L NSCLC setting, across PD-L1 expression levels.

We are planning to develop our anti PD-1 antibody in second line cervical cancer. We are aware of industry sponsored clinical trials, including exploratory studies that are underway in cervical cancer. Our competitors include, but are not restricted to, Regeneron (anti-PD-1), Merck (anti-PD-1), Ono Pharmaceuticals and BMS (anti-PD-1 alone or in combination with anti-CTLA-4 or anti-LAG-3 or anti-IDO), Advaxis (HPV targeting vaccine alone or in combination with AstraZeneca’s anti-PD-L1 antibody or BMS’ anti-PD-1 antibody) and Lion Biotechnologies (autologous TILs). We are also aware that Advaxis has submitted a conditional Marketing Authorization Application (“MAA”) for its HPV

targeting vaccine in the European Union. Additionally, we are also aware of other early stage clinical trials testing alternate CPM targets in cervical cancer patients. These include, but are not restricted to, OX40 +/- CD137 agonists (Pfizer) and anti-PD-1 + anti-ICOS (GSK/Merck).

We have autologous vaccine programs in development including our Prophage vaccine in clinical development for GBM and our neo-antigen based AutoSynVax vaccine in clinical development. We are aware of many companies pursuing personalized cancer vaccines in pre-clinical or clinical development, including, without limitation, the following: Aduro Biotech, Neon Therapeutics, Gritstone Oncology, Advaxis/Amgen, BioNTech, Moderna/Merck, Genocea Biosciences, Argos Therapeutics, EpiVax Inc. Nouscom, Immatix, EpiVax Inc., Achilles Therapeutics and BrightPath Biotherapeutics.

Several companies have products that utilize similar technologies and/or patient-specific medicine techniques that compete with our HSP based vaccines. Schering Corporation, a subsidiary of Merck, markets temozolomide for treatment of patients with ndGBM and refractory astrocytoma. Other companies are developing vaccines for the treatment of patients with ndGBM, including but not limited to Northwest Biotherapeutics (DC-Vax), Mimivax Inc. (SurVaxM) and Annias Immunotherapeutics (CMV Vaccine). Other companies may begin development programs as well.

To the extent we develop our vaccines in other indications or in combination with other product candidates, such as available standard of care agents (Avastin[®]), or with CPMs, they could face additional competition in those indications or in those combinations. In addition, and prior to regulatory approval, if ever, our vaccines and our other product candidates may compete for access to patients with other products in clinical development, with products approved for use in the indications we are studying, or with off-label use of products in the indications we are studying. We anticipate that we will face increased competition in the future as new companies enter markets we seek to address and scientific developments surrounding immunotherapy and other traditional cancer and infectious disease therapies continue to accelerate.

We are aware of compounds that claim to be comparable to QS-21 Stimulon that are being used in clinical trials. Several other vaccine adjuvants are in development and could compete with QS-21 Stimulon for inclusion in vaccines in development. These adjuvants include, but are not limited to, (1) oligonucleotides, under development by Pfizer, Idera, and Dynavax, (2) MF59, under development by Novartis, (3) IC31, under development by Intercell (now part of Valneva), and (4) MPL, under development by GSK. In the past, we have provided QS-21 Stimulon to other entities under materials transfer arrangements. In at least one instance, it is possible that this material was used unlawfully to develop synthetic formulations and/or derivatives of QS-21. In addition, companies such as Adjuvance Technologies, Inc., CSL Limited, and Novavax, Inc., as well as academic institutions and manufacturers of saponin extracts, are developing saponin adjuvants, including derivatives and synthetic formulations. These sources may be competitive to our ability to execute future partnering and licensing arrangements involving QS-21 Stimulon. The existence of products developed by these and other competitors, or other products of which we are not aware, or which other companies may develop in the future, may adversely affect the marketability of products developed or sold using QS-21 Stimulon.

We are also aware of a third party that manufactures pre-clinical material purporting to be comparable to QS-21 Stimulon. The claims being made by this third party may create marketplace confusion and have an adverse effect on the goodwill generated by us and our partners with respect to QS-21 Stimulon. Any diminution of this goodwill may have an adverse effect on our ability to commercialize future products, if any, incorporating this technology, either alone or with a third party.

Employees

As of February 28, 2018, we had 255 employees, of whom 81 were PhDs and one was an MD. None of our employees are subject to a collective bargaining agreement. We believe that we have good relations with our employees.

Corporate History

Antigenics L.L.C. was formed as a Delaware limited liability company in 1994 and was converted to Antigenics Inc., a Delaware corporation, in February 2000 in conjunction with our initial public offering of common stock. On January 6, 2011, we changed our name from Antigenics Inc. to Agenus Inc.

Availability of Periodic SEC Reports

Our Internet website address is www.agenusbio.com. We make available free of charge through our website our annual reports on Form 10-K, quarterly reports on Form 10-Q, current reports on Form 8-K, and amendments to those reports filed or furnished pursuant to Section 13(a) or 15(d) of the Securities Exchange Act of 1934, as amended (“Exchange Act”), as soon as reasonably practicable after we electronically file such material with, or furnish such material to, the Securities and Exchange Commission (the “SEC”). The contents of our website are not part of, or incorporated into, this document. In addition, we regularly use our website to post information regarding our business, product development programs and governance, and we encourage investors to use our website, particularly the information in the sections entitled “Financial” and “News,” as sources of information about us.

The public may read and copy any materials filed by Agenus with the SEC at the SEC's Public Reference Room at 100 F Street, NE, Room 1580, Washington, DC 20549. The public may obtain information on the operation of the Public Reference Room by calling the SEC at 1-800-SEC-0330. The SEC maintains an Internet site that contains reports, proxy and information statements, and other information regarding issuers that file electronically with the SEC at www.sec.gov.

The contents of the websites referred to above are not incorporated into this filing. Further, our references to the URLs for these websites are intended to be inactive textual references only.

Item 1A. Risk Factors

Our future operating results could differ materially from the results described in this Annual Report on Form 10-K due to the risks and uncertainties described below. You should consider carefully the following information about risks below in evaluating our business. If any of the following risks actually occur, our business, financial conditions, results of operations and future growth prospects would likely be materially and adversely affected. In these circumstances, the market price of our common stock would likely decline.

We cannot assure investors that our assumptions and expectations will prove to be correct. Important factors could cause our

actual results to differ materially from those indicated or implied by forward-looking statements. See “Note Regarding Forward-Looking Statements” in this Annual Report on Form 10-K. Factors that could cause or contribute to such differences include those factors discussed below.

Risks Related to our Business

If we incur operating losses for longer than we expect, or we are not able to raise additional capital, we may be unable to continue our operations, or we may become insolvent.

Our net losses for the years ended December 31, 2017, 2016, and 2015, were \$120.7 million, \$127.0 million, and \$87.9 million, respectively. We expect to incur additional losses over the next several years as we continue to research and develop our technologies and pursue partnering opportunities, regulatory strategies, commercialization, and related activities. Furthermore, our ability to generate cash from operations is dependent on the success of our licensees and collaboration partners, as well as the likelihood and timing of new strategic licensing and partnering relationships and/or successful development and commercialization of product candidates, including through our antibody programs and platforms, our vaccine programs, and our saponin-based vaccine adjuvants.

On December 31, 2017, we had \$60.2 million in cash and cash equivalents and short-term investments. We believe that, based on our current plans and activities, including additional funding we anticipate from multiple sources between now and the end of the second quarter of 2018, including out-licensing and/or partnering opportunities, our working capital resources at December 31, 2017, along with the net proceeds of approximately \$28.0 million received from HCR in January 2018 in connection with our royalty transaction, will be sufficient to satisfy our liquidity requirements through the first quarter of 2019. We also continue to monitor the likelihood of success of our key initiatives and can discontinue funding of such activities if they do not prove to be successful, restrict capital expenditures and/or reduce the scale of our operations if necessary.

To date, we have financed our operations primarily through the sale of equity and debt securities. In order to finance future operations, we will be required to raise additional funds in the capital markets, through arrangements with collaboration partners or from other sources. Additional financing may not be available on favorable terms, or at all. If we are unable to raise additional funds when we need them or if we incur operating losses for longer than we expect, we may not be able to continue some or all of our operations, or we may become insolvent. We also may be forced to license or sell technologies to others under agreements that are on unfavorable terms or allocate to third parties substantial portions of the potential value of these technologies.

There are a number of factors that will influence our future capital requirements, including, without limitation, the following:

- the number and characteristics of the product candidates we and our partners pursue;
 - our and our partners’ ability to successfully develop, manufacture, and commercialize product candidates;
- the scope, progress, results and costs of researching and developing our future product candidates and conducting pre-clinical and clinical trials;
- the timing of, and the costs involved in, obtaining regulatory approvals for our and our licensees’ product candidates;
- the cost of manufacturing;
- our ability to establish and maintain strategic partnerships, licensing or other arrangements and the financial terms of such arrangements;
- the costs involved in preparing, filing, prosecuting, maintaining, defending and enforcing our intellectual property rights;
- the costs associated with any successful commercial operations; and
- the timing, receipt and amount of sales of, or royalties on, our future products and those of our partners, if any.

General economic conditions in the United States economy and abroad may have a material adverse effect on our liquidity and financial condition, particularly if our ability to raise additional funds is impaired. The ability of potential patients and/or health care payers to pay for our future products, if any, could also be adversely impacted, thereby limiting our potential revenue. In addition, any negative impacts from any deterioration in the credit markets on our collaboration partners could limit potential revenue from our product candidates.

Our obligations to HCR and the holders of our 2015 Subordinated Notes could materially and adversely affect our liquidity.

In January 2018, we and our wholly-owned subsidiary, Antigenics LLC (“Antigenics”), entered into a Royalty Purchase Agreement (“RPA”) HCR, pursuant to which HCR purchased 100% of Antigenics’ worldwide rights to receive royalties from GSK on sales of GSK’s vaccines containing our QS-21 Stimulon adjuvant. As consideration for the purchase of the royalty rights, HCR paid \$190.0 million at closing, less certain transaction expenses. Of the closing proceeds, approximately \$161.9 million was used to redeem

Antigenics' \$115.0 million principal amount of notes issued pursuant to the Note Purchase Agreement with Oberland Capital SA Zermatt LLC, and we retained approximately \$28.0 million of net proceeds. Antigenics is also entitled to receive up to \$40.35 million in milestone payments based on sales of GSK's vaccines as follows: (i) \$15.1 million upon reaching \$2.0 billion last-twelve-months net sales any time prior to 2024 and (ii) \$25.25 million upon reaching \$2.75 billion last-twelve-months net sales any time prior to 2026. Antigenics will owe approximately \$25.9 million to HCR in 2021 if neither of the following sales milestones are achieved: (i) 2019 sales of GSK's vaccines exceed \$1.0 billion or (ii) 2020 sales of GSK's vaccines exceed \$1.75 billion (the "Rebate Payment"). As part of the transaction, we provided a guaranty for the potential Rebate Payment and secured the obligation with substantially all of our assets pursuant to a security agreement. If GSK's sales do not achieve either of the relevant milestones and we are obligated to make the Rebate Payment, our liquidity could be materially and adversely affected.

In February 2015, we exchanged senior subordinated promissory notes that we issued in 2013 for new senior subordinated promissory notes in the aggregate principal amount of \$5.0 million with annual interest at 8%, and we issued an additional \$9.0 million principal amount of such notes (the "2015 Subordinated Notes"). The 2015 Subordinated Notes were originally due February 2018, and in March 2017, we amended the 2015 Subordinated Notes to extend the maturity date to February 2020. The 2015 Subordinated Notes include default provisions that allow for the acceleration of the principal payment of the 2015 Subordinated Notes in the event we become involved in certain bankruptcy proceedings, become insolvent, fail to make a payment of principal or (after a grace period) interest on the 2015 Subordinated Notes, default on other indebtedness with an aggregate principal balance of \$13.5 million or more if such default has the effect of accelerating the maturity of such indebtedness, or become subject to a legal judgment or similar order for the payment of money in an amount greater than \$13.5 million if such amount will not be covered by third-party insurance. If we default on the 2015 Subordinated Notes and the repayment of such indebtedness is accelerated, our liquidity could be materially and adversely affected.

If we do not have sufficient cash on hand to pay the Rebate Payment when due, or to otherwise service our 2015 Subordinated Notes, we may be required, among other things, to:

- seek additional financing in the debt or equity markets;
- refinance or restructure all or a portion of our indebtedness;
- sell, out-license, or otherwise dispose of assets; and/or
- reduce or delay planned expenditures on research and development and/or commercialization activities.

Such measures might not be sufficient to enable us to make principal and interest payments. In addition, any such financing, refinancing, or sale of assets might not be available on favorable terms, if at all.

We are dependent upon our collaboration with Incyte to further develop, manufacture and commercialize antibodies against certain targets. If we or Incyte fail to perform as expected, the potential for us to generate future revenues under the collaboration could be significantly reduced, the development and/or commercialization of these antibodies may be terminated or substantially delayed, and our business could be adversely affected.

In February 2017, we amended the terms of our collaboration agreement with Incyte to, among other things, convert the GITR and OX40 programs from profit-share programs, where we and Incyte shared all costs and profits on a 50:50 basis, to royalty-bearing programs, where Incyte funds 100% of the costs and we are eligible for potential milestones and royalties. In addition, the profit-share programs relating to TIGIT and one undisclosed target were removed from the collaboration, with TIGIT reverting to Agenus and the undisclosed target reverting to Incyte, each with a potential 15% royalty to the other party on any global net sales. The remaining three royalty-bearing programs in the collaboration targeting TIM-3, LAG-3 and one undisclosed target remain unchanged, and there are no more profit-share programs under the collaboration. For each program in the collaboration, we serve as the lead for pre-clinical development activities through the filing of an IND, and Incyte has exclusive rights and all decision-making authority for manufacturing, clinical development and commercialization. Accordingly, the timely and successful completion by Incyte of clinical development and commercialization activities will significantly affect the timing and amount of any royalties or milestones we may receive under the collaboration agreement. In addition,

in March 2017 we transferred manufacturing responsibilities to Incyte for antibodies under that collaboration. Any delays or weaknesses in the ability of Incyte to successfully manufacture could have an adverse impact on those programs. Incyte's activities will be influenced by, among other things, the efforts and allocation of resources by Incyte, which we cannot control. If Incyte does not perform in the manner we expect or fulfill its responsibilities in a timely manner, or at all, the clinical development, manufacturing, regulatory approval, and commercialization efforts related to antibodies under the collaboration could be delayed or terminated. There can be no assurance that any of the development, regulatory or sales milestones will be achieved, or that we will receive any future milestone or royalty payments under the collaboration agreement.

In addition, our collaboration with Incyte may be unsuccessful due to other factors, including, without limitation, the following:

- Incyte may terminate the agreement or any individual program for convenience upon 12 months' notice;
- Incyte has control over the development of assets in the collaboration;
- Incyte may change the focus of its development and commercialization efforts or prioritize other programs more highly and, accordingly, reduce the efforts and resources allocated to our collaboration;
- Incyte may choose not to develop and commercialize antibody products, if any, in all relevant markets or for one or more indications, if at all; and
- If Incyte is acquired during the term of our collaboration, the acquirer may have competing programs or different strategic priorities that could cause it to reduce its commitment to our collaboration.

If Incyte terminates our collaboration agreement, we may need to raise additional capital and may need to identify and come to agreement with another collaboration partner to advance certain of our antibody programs. Even if we are able to find another partner, this effort could cause delays in our timelines and/or additional expenses, which could adversely affect our business prospects and the future of our antibody product candidates under the collaboration.

Our antibody programs are in early stage development, and there is no guarantee that we or our partners will be successful in advancing antibody product candidates into and through clinical development.

Our antibody programs are currently in early stage development, and many of our antibody programs are pre-clinical. Even if our pre-clinical studies or our and/or our partners' clinical trials produce positive results, they may not necessarily be predictive of the results of future clinical trials. Many companies in the pharmaceutical, biopharmaceutical and biotechnology industries have suffered significant setbacks in clinical trials after achieving positive results in pre-clinical development or earlier clinical trials, and we cannot be certain that we will not face similar setbacks. These setbacks have been caused by, among other things, pre-clinical findings made while clinical trials were underway or safety or efficacy observations made in clinical trials, including adverse events. Moreover, pre-clinical and clinical data are often susceptible to varying interpretations and analyses, and many companies that believed their product candidates performed satisfactorily in pre-clinical studies and clinical trials nonetheless failed to obtain regulatory approval. If we and our partners fail to produce positive results in clinical trials of antibodies, our business and financial prospects would be materially adversely affected.

Although we are targeting to file our first BLA as early as the second half of 2019 and becoming a commercial organization in 2020, there is no guarantee that we will be able to do so on that timeline or at all. Our stated timelines are aggressive and subject to various factors outside of our control, including patient accrual rates for our clinical trials. If our trials are unable to accrue patients at the rate we expect, we are unlikely to hit our anticipated timelines and our business and financial prospects could be materially adversely affected.

Similarly, although we are striving to file numerous INDs to advance novel antibodies and cell therapy candidates into the clinic in the next 12-18 months, there is no guarantee that we will be able to do so on that timeline, if at all. Our stated timelines are aggressive and subject to various risks, including resource constraints. If we are unable to advance novel candidates into the clinic as planned due to resource constraints or otherwise, our business and partnering prospects could be materially adversely affected.

We have undergone significant growth across multiple locations over the past few years, and are focusing on further enhancing core areas and capabilities as we move toward commercialization. In addition, we have consolidated certain sites while expanding others to focus on our core priorities and future needs. We may encounter difficulties in managing these growth and/or consolidation efforts, either of which could disrupt our operations.

Since our acquisition of Agenus Switzerland Inc., formerly known as 4-Antibody AG ("4-AB") in January 2014, we have more than tripled our headcount, in part through various acquisitions and the expansion of our research and development activities both nationally and internationally. While we have restructured our organization over the past

two years, we expect to continue increasing our headcount in certain core areas as we continue to build our development, manufacturing and commercialization capabilities and integrate our acquired technology platforms. To manage these organizational changes, we must continue to implement and improve our managerial, operational and financial systems and continue to recruit, train and retain qualified personnel. If our management is unable to effectively manage our growth, our expenses may increase more than expected, our ability to generate revenue could be reduced, and we may not be able to implement our business strategy.

As part of our efforts to optimize efficiency across our organization, we closed our Jena office in 2016 and consolidated these operations in the United Kingdom and Switzerland. In 2017, we completed a reduction in force in our Lexington, MA facility, which included certain members of our management, in line with our prioritization efforts, and we closed our office in Basel, Switzerland and transferred our research and development assets and capabilities there to the United Kingdom. If these transition efforts prove to

be unsuccessful, or if we identify management or operational gaps in connection with our changes, it could cause delays in discovery timelines and increased costs for certain of our internal and partnered programs, which also could have an adverse effect on our business, financial condition and results of operations.

Our synthetic Heat Shock Protein (“HSP”) peptide-based platform is in early stage development, and there is no guarantee that a product candidate will progress from this platform.

In June 2014, we reported positive results from a Phase 2 trial with HerpVTM, a vaccine candidate for genital herpes from our synthetic HSP peptide-based platform. While the HerpV Phase 2 trial met its formal endpoints, subjects were not followed long enough to determine whether the magnitude of the effect on viral load would be sufficient to significantly reduce the incidence, severity, or duration of herpetic lesions or reduce the risk of viral transmission. Although we have not advanced this program into a Phase 3 trial, we initiated our ASV synthetic cancer vaccine program based on our prior findings with this platform. We initiated our first clinical trial for our first AutoSynVax product candidate in 2017; however, there is no guarantee that results of this trial or any potential future clinical trials will be positive. Although we are targeting to initiate a combination trial with ASV and one or more of our antibodies in 2018, there is no guarantee that we will be able to do so on that timeline or at all. Furthermore, it is possible that research and discoveries by others will render any product candidate from this platform as obsolete or noncompetitive.

We may not be able to advance clinical development or commercialize our cancer vaccine candidates or realize any benefits from these programs.

The probability of future clinical development efforts leading to marketing approval and commercialization of Prophage vaccines is highly uncertain. Prophage vaccines have been in clinical development for over 16 years, including multiple Phase 1 and 2 trials in eight different tumor types as well as randomized Phase 3 trials in metastatic melanoma and adjuvant renal cell carcinoma. To date, none of our clinical trials with Prophage vaccines have resulted in a marketing approval, except in Russia where commercialization of the approved product was unsuccessful. All of our currently planned trials involving Prophage are intended to be sponsored by third parties, and there is no guarantee that they will occur at all. In addition, while we believe Prophage vaccines may provide clinical benefit to some patients as a monotherapy and in combination with other therapies, there is no guarantee that, if completed, subsequent Prophage trials would yield useful translational and/or efficacy data.

Our current clinical trial plans with Prophage vaccines entails one government sponsored IND in which we provide support and product supply. For third-party sponsored trials, we lack the ability to control trial design, timelines, tumor tissue procurement and data availability. For example, in January 2017, we announced a clinical trial collaboration with the National Cancer Institute (“NCI”), whereby the NCI is conducting a double-blind, randomized controlled Phase 2 trial to evaluate the effect of Prophage vaccine in conjunction with Merck’s pembrolizumab on the overall survival rate of patients with newly diagnosed glioblastoma (“ndGBM”). In addition, the Phase 2 trial of Prophage vaccine in combination with bevacizumab in patients with surgically resectable recurrent glioma that was being conducted under the sponsorship of the Alliance for Clinical Trials in Oncology, a cooperative group of the NCI, has been closed. In addition, our other cancer vaccine programs (ASV and PSV) are in Phase 1 and pre-clinical development, respectively, and there is no guarantee that they will successfully advance in and through the clinic. ASV also utilizes QS-21 Stimulon, and any inability or delay in securing adequate supplies of the adjuvant could have an adverse impact on the program or otherwise delay timelines. Current and future studies may eventually be terminated due to, among other things, slow enrollment, lack of probability that they will yield useful translational and/or efficacy data, lengthy timelines, or the unlikelihood that results will support timely or successful regulatory filings.

Changes in our manufacturing strategies, manufacturing problems, or increased demand may cause delays, unanticipated costs, or loss of revenue streams within or across our programs.

Our antibody programs will require substantial manufacturing development and investment to progress. We are currently progressing a portfolio of antibody programs that are at different stages of development. If these efforts are delayed or do not produce the desired outcomes, this will cause delays in development timelines and increased costs, which may cause us to limit the size and scope of our efforts and studies. In December 2015, we secured our own antibody manufacturing capabilities with the purchase of a manufacturing pilot plant from XOMA Corporation (“XOMA”), and we expect this facility to supply us with antibody drug substance requirements through clinical proof-of-concept studies. We will also need to develop or secure later phase and/or commercial manufacturing capabilities for larger, registrational studies or any commercial supply requirements. For the programs for which we will produce our own drug substance, we will continue to rely on third parties for fill-finish services and other parts of the manufacturing process. These services include the storage and maintenance of our drug substance during all stages of the manufacturing process. While we maintain insurance to cover certain potential losses, there is no guarantee that our insurance coverage will be adequate. Furthermore, we currently rely on contract manufacturing organizations (“CMOs”) and contract research organizations (“CROs”) to support some of our existing antibody programs. Our dependence on external CMOs for the manufacture of certain antibodies results in intrinsic risks to our performance, timelines, and costs of our accelerated development plans, and which

could divert resources away from our antibody programs and/or lead to delays in the development of our product candidates. In the event that our antibody programs require progressively larger production capabilities, our options for qualified CMOs may become more limited.

The long-term success of the antibody pilot plant manufacturing facility and capabilities that we acquired from XOMA will depend, in part, on our ability to realize the anticipated synergies, business opportunities and growth prospects from combining our manufacturing facilities in Lexington, MA with the antibody pilot plant manufacturing facility in Berkeley, CA. We may never realize these anticipated synergies, business opportunities and growth prospects. Assumptions underlying estimates of expected cost savings as a result of the acquisition of the antibody pilot plant manufacturing facility may be inaccurate. If any of these factors limit our ability to successfully manufacture antibodies to support our current and future clinical trials, the expectations of future results of operations, including certain cost savings and synergies expected to result from the acquisition of XOMA's antibody pilot plant manufacturing facility, might not be met.

We currently manufacture our Prophage vaccines in our Lexington, MA facility. Manufacturing of the Prophage vaccines is complex, and various factors could cause delays or an inability to supply the vaccine. Deviations in the processes controlling manufacture or deficiencies in size or quality of source material could result in production failures. Specific vulnerabilities in the process may exist in tumor types in which quality or quantity of tissue is limited. In addition, regulatory bodies may require us to make our manufacturing facility a single product facility. In such an instance, we would no longer have the ability to manufacture Prophage vaccines in addition to other product candidates in our current facility.

We have given our corporate QS-21 Stimulon licensee, GSK, manufacturing rights for QS-21 Stimulon for use in their product programs. We have retained the right to manufacture QS-21 for ourselves and third parties, although no other such programs are anticipated to bring us substantial revenues in the near future, if ever. Although we have the right to secure certain quantities of QS-21 from GSK and we have some internal supply in-house, we currently do not have an alternative long term supply partner for this adjuvant.

Our ability to efficiently manufacture our product candidates is contingent, in part, upon our own, and our CMOs', ability to ramp up production in a timely manner without the benefit of years of experience and familiarity with the processes, which we may not be able to adequately transfer. We currently rely upon and expect to continue to rely upon third parties, potentially including our collaborators or licensees, to produce materials required to support our product candidates, pre-clinical studies, clinical trials, and any future commercial efforts. A number of factors could cause production interruptions at either our manufacturing facility or the facilities of our CMOs or suppliers, including equipment malfunctions, labor or employment retention problems, natural disasters, power outages, terrorist activities, or disruptions in the operations of our suppliers. Alternatively, there is the possibility we may have excess manufacturing capacity if product candidates do not progress as planned.

As mentioned above, reliance on third-party manufacturers entails risks to which we would not be subject if we manufactured all of our product candidates ourselves, including reliance on the third party for regulatory compliance, the possibility of breach of the manufacturing agreement by the third party because of factors beyond our control, and the possibility of termination or non-renewal of the agreement by the third party, based on its own business priorities, at a time that is costly or inconvenient for us.

Biopharmaceutical manufacturing is also subject to extensive government regulation. Components of a finished therapeutic product approved for commercial sale or used in late-stage clinical trials must be manufactured in accordance with current good manufacturing practices. These regulations govern manufacturing processes and procedures (including record-keeping) and the implementation and operation of quality systems to control and assure the quality of investigational products and products approved for sale. Our facilities and quality systems and the facilities and quality systems of some or all of our third-party contractors must pass a pre-approval inspection for compliance with the applicable regulations as a condition of regulatory approval of a product candidate. In addition,

facilities are subject to on-going inspections and routine audits, and minor changes in manufacturing processes may require additional regulatory approvals and audits, either of which could cause us to incur significant additional costs, set-backs or delays and eventual loss of revenue.

Risks associated with doing business internationally could negatively affect our business.

We currently have research and development operations in the United Kingdom, and we expect to pursue pathways to develop and commercialize our product candidates in both U.S. and non-U.S. jurisdictions. Various risks associated with foreign operations may impact our success. Possible risks of foreign operations include fluctuations in the value of foreign and domestic currencies requirements to comply with various jurisdictional requirements such as data privacy regulations, disruptions in the import, export, and transportation of patient tumors and our products or product candidates, the product and service needs of foreign customers, difficulties in building and managing foreign relationships, the performance of our licensees or collaborators, geopolitical instability, unexpected regulatory, economic, or political changes in foreign and domestic markets, including without limitation any resulting

from the United Kingdom's withdrawal from the European Union or our current political regime, and limitations on the flexibility of our operations and costs imposed by local labor laws.

Our competitors may have superior products, manufacturing capability, selling and marketing expertise and/or financial and other resources.

Our product candidates and the product candidates in development by our collaboration partners may fail because of competition from major pharmaceutical companies and specialized biotechnology companies that market products, or that are engaged in the development of product candidates and for the treatment cancer. Many of our competitors, including large pharmaceutical companies, have greater financial and human resources and more experience than we do. Our competitors may:

- develop safer or more effective therapeutic drugs or therapeutic vaccines and other products;
- establish superior intellectual property positions;
- discover technologies that may result in medical insights or breakthroughs, which render our drugs or vaccines obsolete, possibly before they generate any revenue, if ever;
- adversely affect our ability to recruit patients for our clinical trials;
- solidify partnerships or strategic acquisitions that may increase the competitive landscape;
 - develop or commercialize their product candidates sooner than we commercialize our own, if ever; or
- implement more effective approaches to sales and marketing and capture some of our potential market share.

There is no guarantee that our product candidates will be able to compete with potential future products being developed by our competitors.

The CPM landscape is crowded with several competitors developing assets against a number of targets. Development plans are spread out across various indications and lines of therapy, either alone or in combination with other assets. Competitors range from small cap to large cap companies, with assets in pre-clinical or clinical stages of development. Therefore, the landscape is dynamic and constantly evolving. We and our partners have CPM antibody programs currently in clinical stage development targeting PD-1, CTLA-4, GITR and OX40. We are aware of many companies that have antibody-based products on the market or in clinical development that are directed to the same biological targets as these programs, including, without limitation, the following: (1) BMS markets ipilimumab, an anti-CTLA-4 antibody, and nivolumab, an anti-PD-1 antibody, and is developing agonists to GITR and OX-40, (2) Merck has an approved anti-PD-1 antibody in the United States, as well as an anti-CTLA-4 antagonist and an anti-GITR agonist in clinical development, (3) Ono Pharmaceuticals has an approved anti-PD-1 antibody in Japan, (4) AstraZeneca has an approved anti PD-L1 antibody, as well as anti-CTLA-4, PD-1, GITR and OX40 targeting antibodies in development, (5) Pfizer has an approved anti-PD-L1 (with Merck KgaA) as well as anti-PD-1, and anti-OX40 antibodies in clinical development, (6) Novartis has anti-PD-1, anti-PD-L1 and anti-GITR antibodies in clinical trials, and (7) Roche/Genentech has an approved anti-PD-L1 antibody. We are also aware of other competitors with PD-1/PD-L1 antibodies in clinical development, including AbbVie, Arcus Biosciences, Boehringer Ingelheim, Tesaro, Beigene, Regeneron, Eli Lilly, Jiangsu HengRui Medicine, Shanghai Junshi, MacroGenics, CytomX, Janssen, CBT Pharmaceuticals, Checkpoint Therapeutics, CStone Pharmaceuticals, Livzon MabPharm Inc and Suzhou Alphamab. We are also aware of competitors with pre-clinical antibodies against these targets. In addition, we are aware of competitors with clinical stage antibodies against targets in our earlier stage programs such as TIM-3, LAG-3, CD137, TIGIT and other undisclosed targets. These include, but are not limited to, BMS, Pfizer, Novartis, Merck, Roche, Tesaro, Eli Lilly, OncoMed, Boehringer Ingelheim, Astellas and Regeneron. Additionally, we are aware of competitors with assets against these targets that are in pre-clinical development. There is no guarantee that our antibody product candidates will be able to compete with our competitors' antibody products and product candidates.

We are planning to advance combinations of our anti-CTLA-4 antibody, AGEN1884, with Keytruda in 1L NSCLC, where Keytruda is currently approved. Specifically, we will be evaluating efficacy within a subset of patients who express $\geq 50\%$ PD-L1 protein biomarker levels. We are aware of competitors who have approved immunotherapy

products in this indication, including having a label specifically directed at this patient subset, such as Merck (anti-PD-1 monotherapy). We are also aware of industry sponsored clinical trials, including exploratory studies, that are directed at this specific patient population. These include, but are not limited to, Incyte/Merck (anti-IDO + anti-PD-1), Merck (anti-PD-1 + anti-CTLA-4), Regeneron (anti-PD-1), Roche (anti-PD-L1), Merck KgaA / Pfizer (anti-PD-L1), and Tesaro (anti-PARP + anti-PD-1). We are also aware of competitors who have approved immunotherapies or have published positive Phase 3 data for immunotherapies in 1L NSCLC. These include, and are not limited to, Merck, BMS, and Roche. Additional competitors have ongoing clinical trials for immunotherapies in the 1L NSCLC setting, across PD-L1 expression levels.

We are also conducting activities in second line cervical cancer. We are aware of industry sponsored clinical trials, including exploratory studies, that are underway in cervical cancer. Our competitors include, but are not limited to, Regeneron (anti-PD-1), Merck (anti-PD-1), Ono Pharmaceuticals and BMS (anti-PD-1 alone or in combination with anti-CTLA-4 or anti-LAG-3 or anti-IDO),

Advaxis (HPV targeting vaccine alone or in combination with AstraZeneca's anti-PD-L1 antibody or BMS' anti-PD-1 antibody) and Lion Biotechnologies (autologous TILs). We are also aware that Advaxis has submitted a conditional Marketing Authorization Application (MAA) for its HPV targeting vaccine in the European Union. Additionally, we are aware of other early stage clinical trials testing alternate CPM targets in cervical cancer patients. These include, but are not limited to, OX40 +/- CD137 agonists (Pfizer) and anti-PD-1 + anti-ICOS (GSK/Merck).

We have autologous vaccine programs in development including our Prophage vaccine in clinical development for GBM and our neo-antigen based AutoSynVax vaccine in clinical development. We are aware of many companies pursuing personalized cancer vaccines in pre-clinical or clinical development, including, without limitation, the following: Aduro Biotech, Neon Therapeutics, Gritstone Oncology, Advaxis/Amgen, BioNTech, Moderna/Merck, Genocea Biosciences, Argos Therapeutics, EpiVax Inc. Nouscom, Immatics, EpiVax Inc., Achilles Therapeutics and BrightPath Biotherapeutics.

Several companies have products that utilize similar technologies and/or patient-specific medicine techniques that compete with our HSP based vaccines. Schering Corporation, a subsidiary of Merck, markets temozolomide for treatment of patients with ndGBM and refractory astrocytoma. Other companies are developing vaccines for the treatment of patients with ndGBM, including, but not limited to, Northwest Biotherapeutics (DC-Vax), Mimivax Inc. (SurVaxM) and Annias Immunotherapeutics (CMV Vaccine). Other companies may begin development programs as well.

To the extent we develop our vaccines in other indications or in combination with other product candidates, such as available standard of care agents (Avastin®), or with CPMs, they could face additional competition in those indications or in those combinations. In addition, and prior to regulatory approval, if ever, our vaccines and our other product candidates may compete for access to patients with other products in clinical development, with products approved for use in the indications we are studying, or with off-label use of products in the indications we are studying. We anticipate that we will face increased competition in the future as new companies enter markets we seek to address and scientific developments surrounding immunotherapy and other traditional cancer and infectious disease therapies continue to accelerate.

We are aware of compounds that claim to be comparable to QS-21 Stimulon that are being used in clinical trials. Several other vaccine adjuvants are in development and could compete with QS-21 Stimulon for inclusion in vaccines in development. These adjuvants include, but are not limited to, (1) oligonucleotides, under development by Pfizer, Idera, and Dynavax, (2) MF59, under development by Novartis, (3) IC31, under development by Intercell (now part of Valneva), and (4) MPL, under development by GSK. In the past, we have provided QS-21 Stimulon to other entities under materials transfer arrangements. In at least one instance, it is possible that this material was used unlawfully to develop synthetic formulations and/or derivatives of QS-21. In addition, companies such as Adjuvance Technologies, Inc., CSL Limited, and Novavax, Inc., as well as academic institutions and manufacturers of saponin extracts, are developing saponin adjuvants, including derivatives and synthetic formulations. These sources may be competitive to our ability to execute future partnering and licensing arrangements involving QS-21 Stimulon. The existence of products developed by these and other competitors, or other products of which we are not aware or which other companies may develop in the future, may adversely affect the marketability of products developed or sold using QS-21 Stimulon.

We are also aware of a third party that manufactures pre-clinical material purporting to be comparable to QS-21 Stimulon. The claims being made by this third party may create marketplace confusion and have an adverse effect on the goodwill generated by us and our partners with respect to QS-21 Stimulon. Any diminution of this goodwill may have an adverse effect on our ability to commercialize future products, if any, incorporating this technology, either alone or with a third party.

Failure to realize the anticipated benefits of our strategic acquisitions and licensing transactions could adversely affect our business, operations and financial condition.

An important part of our business strategy has been to identify and advance a pipeline of product candidates by acquiring and in-licensing product candidates, technologies and businesses that we believe are a strategic fit with our existing business. Since we acquired 4-AB), in February 2014, we have completed numerous additional strategic acquisitions and licensing transactions. The ultimate success of these strategic transactions entails numerous operational and financial risks, including:

- higher than expected development and integration costs;
- difficulty in combining the technologies, operations and personnel of acquired businesses with our technologies, operations and personnel;
- exposure to unknown liabilities;
- difficulty or inability to form a unified corporate culture across multiple office sites both nationally and internationally;
- inability to retain key employees of acquired businesses;
- disruption of our business and diversion of our management's time and attention; and
- difficulty or inability to secure financing to fund development activities for such acquired or in-licensed product candidates, technologies or businesses.

We have limited resources to integrate acquired and in-licensed product candidates, technologies and businesses into our current infrastructure, and we may fail to realize the anticipated benefits of our strategic transactions. Any such failure could have an adverse effect on our business, operations and financial condition.

Failure to enter into and/or maintain significant licensing, distribution and/or collaboration agreements on favorable terms to us may hinder or cause us to cease our efforts to develop and commercialize our product candidates, increase our development timelines, and/or increase our need to rely on partnering or financing mechanisms, such as sales of debt or equity securities, to fund our operations and continue our current and anticipated programs.

As previously noted, our ability to advance our antibody programs depends in part on collaboration agreements such as our collaboration with Incyte. See “Risk Factors—Risks Related to Our Business—We are dependent upon our collaboration with Incyte to further develop, manufacture and commercialize antibodies against certain targets. If we or Incyte fail to perform as expected, the potential for us to generate future revenues under the collaboration could be significantly reduced, the development and/or commercialization of these antibodies may be terminated or substantially delayed, and our business could be adversely affected.” In addition, from time to time we engage in efforts to enter into licensing, distribution and/or collaboration agreements with one or more pharmaceutical or biotechnology companies to assist us with development and/or commercialization of our other product candidates. If we are successful in entering into such agreements, we may not be able to negotiate agreements with economic terms similar to those negotiated by other companies. We may not, for example, obtain significant upfront payments, substantial royalty rates or milestones. If we fail to enter into any such agreements, our efforts to develop and/or commercialize our product candidates may be undermined. In addition, if we do not raise funds through any such agreements, we will need to rely on other financing mechanisms, such as sales of debt or equity securities, to fund our operations. Such financing mechanisms, if available, may not be sufficient or timely enough to advance our programs forward in a meaningful way in the short-term.

Because we rely on collaborators and licensees for the development and commercialization of certain of our product candidate programs, these programs may not prove successful, and/or we may not receive significant payments from such parties.

Part of our strategy is to develop and commercialize many of our product candidates by continuing or entering into arrangements with academic, government, or corporate collaborators and licensees. Our success depends on our ability to negotiate such agreements on favorable terms and on the success of the other parties in performing research, pre-clinical and clinical testing, completing regulatory applications, and commercializing product candidates. Our research, development, and commercialization efforts with respect to antibody candidates from our technology platforms are, in part, contingent upon the participation of institutional and corporate collaborators. For example, in February 2015, we began a broad collaboration with Incyte to pursue the discovery and development of antibodies. See “Risk Factors-Risks Related to our Business—We are dependent upon our collaboration with Incyte to further develop, manufacture and commercialize antibodies against certain targets. If we or Incyte fail to perform as expected, the potential for us to generate future revenues under the collaboration could be significantly reduced, the development and/or commercialization of these antibodies may be terminated or substantially delayed, and our business could be adversely affected.” Furthermore, we have a collaboration arrangement with Recepta for CTLA-4 and PD-1, giving Recepta rights to certain South American countries and requiring us to agree upon development plans for these candidates. Disagreements or the failure of either party to perform satisfactorily could have an adverse impact on these programs.

The Brain Tumor Trials Collaborative is sponsoring a Phase 2 clinical trial of our Prophage vaccine candidate in combination with Merck’s pembrolizumab in patients with glioma. When our licensees or third-party collaborators sponsor clinical trials using our product candidates, we cannot control the timing of enrollment, data readout, or quality of such trials or related activities.

Development activities for our collaboration programs may fail to produce marketable products due to unsuccessful results or abandonment of these programs, failure to enter into future collaborations or license agreements, or the inability to manufacture product supply requirements for our collaborators and licensees. Several of our agreements also require us to transfer important rights and regulatory compliance responsibilities to our collaborators and licensees. As a result of these collaboration agreements, we will not control the nature, timing, or cost of bringing these product candidates to market. Our collaborators and licensees could choose not to, or be unable to, devote resources to these arrangements or adhere to required timelines, or, under certain circumstances, may terminate these arrangements early. They may cease pursuing product candidates or elect to collaborate with different companies. In addition, these collaborators and licensees, outside of their arrangements with us, may develop technologies or products that are competitive with those that we are developing. From time to time, we may also become involved in disputes with our collaborators or licensees. Such disputes could result in the incurrence of significant expense, or the termination of collaborations. We may be unable to fulfill all of our obligations to our collaborators, which may result in the termination of collaborations. As a result of these factors, our strategic collaborations may not yield revenue. Furthermore, we may not be able to enter into new collaborations on favorable terms or at all. Failure to generate significant revenue from collaborations could increase our need to fund our operations through sales of debt or equity securities and would negatively affect our business prospects.

Our internal computer systems, or those of our third-party CROs, CMOs, licensees, collaborators or other contractors or consultants, may fail or suffer security breaches, which could result in a material disruption in our business and operations or could subject us to sanctions and penalties that could have a material adverse effect on our reputation or financial condition.

Despite the implementation of security measures, our internal computer systems and those of our current and future CROs, CMOs, licensees, collaborators and other contractors and consultants are vulnerable to damage from computer viruses, unauthorized access, natural disasters, terrorism, war and telecommunication and electrical failures. While we are not aware of any such material system failure, accident or security breach to date, if such an event were to occur and cause interruptions in our operations, it could result in a material disruption of our development programs and our business operations. For example, the loss of clinical trial data from completed, on-going or future clinical trials could result in delays in our regulatory approval efforts and significant costs to recover or reproduce the data. Likewise, we rely on third parties to manufacture certain of our drug candidates and conduct clinical trials, and similar events relating to their computer systems could also have a material adverse effect on our business. To the extent that any disruption or security breach were to result in a loss of, or damage to, our data or applications, or inappropriate disclosure of confidential or proprietary information, we could incur liabilities and the further development and commercialization of our product candidates could be delayed.

We use and store customer, vendor, employee and business partner and, in certain instances patient, personally identifiable information in the ordinary course of our business. We are subject to various domestic and international privacy and security regulations, including but not limited to the Health Insurance Portability and Accountability Act of 1996 (“HIPAA”), which mandates, among other things, the adoption of uniform standards for the electronic exchange of information in common healthcare transactions, as well as standards relating to the privacy and security of individually identifiable health information, which require the adoption of administrative, physical and technical safeguards to protect such information. In addition, many states have enacted comparable laws addressing the privacy and security of health information, some of which are more stringent than HIPAA. Failure to comply with these standards, or a computer security breach or cyber-attack that affects our systems or results in the unauthorized release of proprietary or personally identifiable information, could subject us to criminal penalties and civil sanctions, and our reputation could be materially damaged, and our operations could be impaired. We may also be exposed to a risk of loss or litigation and potential liability, which could have a material adverse effect on our business, results of operations and financial condition.

We are highly reliant on certain members of our management team. In addition, we have limited internal resources and if we fail to recruit and/or retain the services of key employees and external consultants as needed, we may not be able to achieve our strategic and operational objectives.

Garo H. Armen, Ph.D., the Chairman of our Board of Directors and our Chief Executive Officer who co-founded the Company in 1994, is integral to building our company and developing our technology. If Dr. Armen is unable or unwilling to continue his relationship with Agenus, our business may be adversely impacted. We have an employment agreement with Dr. Armen, and he plays an important role in our day-to-day activities. We do not carry a key employee insurance policy for Dr. Armen or any other employee.

Our future growth success depends to a significant extent on the skills, experience and efforts of our executive officers and key members of our clinical and scientific staff. We face intense competition for qualified individuals from other pharmaceutical, biopharmaceutical and biotechnology companies, as well as academic and other research institutions. We may be unable to retain our current personnel or attract or assimilate other highly qualified management and clinical personnel in the future on acceptable terms. The loss of any or all of these individuals could harm our business and could impair our ability to support our collaboration partners or our growth generally. If our management is unable to effectively manage our growth, our expenses may increase more than expected, our ability to generate revenue could be reduced and we may not be able to implement our business strategy. Moreover, in connection with our 2017 restructuring activities, certain positions on our management team were eliminated and Dr. Robert Stein

retired from his role as President of R&D to become a senior R&D advisor to the Company. Any key capability gaps identified following this restructuring could have a material adverse effect on our business, financial condition and results of operations.

We intend to advance our cell therapy business through our new subsidiary, AgenTus Therapeutics, eventually with separate funding. Moving intellectual property assets into AgenTus Therapeutics in foreign jurisdictions could have adverse tax consequences, and there is no guarantee that we will be able to attract external funding. Moreover, even if the business is funded, there is no guarantee that it will be successful.

We are currently in the process of pursuing external funding and partnership opportunities to advance AgenTus Therapeutics, but Agenus is currently funding such operations. There is no guarantee that external funding will be available. If funding is available, there is no guarantee that it will be on attractive or acceptable terms, or that it will be adequate to advance the business to an inflection point for additional funding. Similarly, there is no guarantee that partnership opportunities will be available on attractive terms, if at all. If external funding is not available, we may be forced to either retire these programs or use internal resources to advance them. In addition, our cell therapy assets are pre-clinical. Even if adequate funding and partnership opportunities are available, there is no guarantee that we will be successful in advancing one or more product candidates into and through clinical development. In addition,

most of the efforts being made on behalf of AgenTus Therapeutics are being led by a separate AgenTus chief executive officer, utilizing Agenus' management team and internal resources. The current structure could distract management and divert Agenus resources from Agenus' own core pipeline and programs.

The cell therapy assets necessary to enable AgenTus Therapeutics are currently owned or controlled by Agenus in the United States and Switzerland. In connection with capitalizing AgenTus Therapeutics, these assets will be transferred or licensed to new legal entities within the United States and Europe. Transferring these assets or licensing them on an exclusive basis would require that taxes be paid based on the fair market value of the assets. While we expect to have adequate net operating losses to offset any tax liabilities, there is no guarantee that this will be the case in all relevant jurisdictions. Moreover, we have previously disclosed our interest in potentially issuing a tax-free dividend to Agenus' stockholders in the form of stock of AgenTus Therapeutics. There is no guarantee that any such dividend will be tax-free or that it will be issued at all, or the timing thereof. If we issue a dividend in the form of stock, there could be adverse tax consequences for certain of our stockholders.

Calamities, power shortages or power interruptions could disrupt our business and materially adversely affect our operations.

If a natural disaster, power outage or other event occurred that prevented us from using all or a significant portion of our facilities, that damaged critical infrastructure (such as our manufacturing facility) or that otherwise disrupted operations, it may be difficult or, in certain cases, impossible for us to continue certain activities, such as for example our manufacturing capabilities, for a substantial period of time. We own an antibody pilot plant manufacturing facility and lease additional office space in Berkeley, CA. This location is in an area of seismic activity near active earthquake faults. Any earthquake, terrorist attack, fire, power shortage or other calamity affecting our facilities or those of third parties upon whom we depend may disrupt our business and could have a material adverse effect on our business, results of operations, financial condition and prospects. The disaster recovery and business continuity plans we have in place currently are limited and are unlikely to prove adequate in the event of a serious disaster or similar event. We may incur substantial expenses and delays as a result of the limited nature of our disaster recovery and business continuity plans, which could have a material adverse effect on our business.

Comprehensive tax reform legislation could adversely affect our business and financial condition.

On December 22, 2017, President Trump signed into law the "Tax Cuts and Jobs Act" (the "TCJA") that significantly reforms the Internal Revenue Code of 1986, as amended (the "Code"). The TCJA, among other things, includes changes to U.S. federal tax rates, imposes significant additional limitations on the deductibility of interest and net operating loss carryforwards, allows for the expensing of capital expenditures, and puts into effect the migration from a "worldwide" system of taxation to a territorial system. Our net deferred tax assets and liabilities will be revalued at the newly enacted U.S. corporate rate. We do not expect to recognize any tax expense in the year of enactment as our net deferred tax assets have a full valuation allowance recorded. We continue to examine the impact this tax reform legislation may have on our business. The impact of this tax reform is uncertain and could be adverse.

Risks Related to Regulation of the Biopharmaceutical Industry

The drug development and approval process is uncertain, time-consuming, and expensive.

Drug development, including non-clinical testing and clinical development, and the process of obtaining regulatory approvals for new therapeutic products, is lengthy, expensive, and uncertain. For example, as of December 31, 2017, we had spent more than 20 years and \$684.6 million on our research and development programs. The development and regulatory approval processes also can vary substantially based on the therapeutic area, type, complexity, and novelty of the product. We must provide regulatory authorities with manufacturing, product characterization, and pre-clinical and clinical data demonstrating that our product candidates are safe and effective before they can be approved for commercial sale. It may take us many years to complete our testing, and failure can occur at any stage.

Results of pre-clinical studies do not necessarily predict clinical results, and promising results in early clinical studies might not be confirmed in later studies. Any pre-clinical or clinical test may fail to produce results satisfactory to regulatory authorities for many reasons, including but not limited to emerging manufacturing or control issues, limitations of pre-clinical assessments, difficulties to enroll a sufficient number of patients, changing therapeutic landscape or failure to prospectively identify the benefit/risk profile of the new product. Pre-clinical and clinical data can be interpreted in different ways, which could delay, limit, or prevent regulatory approval. Negative or inconclusive results from a pre-clinical study or clinical trial, adverse medical events during a clinical trial, or safety issues emerging with products of the same class of drug could require additional studies or cause a program to be terminated, even if other studies or trials relating to the program are successful. We or the FDA, other regulatory agencies, or an institutional review board may suspend or terminate human clinical trials at any time on various grounds.

The timing and success of a clinical trial is dependent on obtaining and maintaining sufficient cash resources, successful production of clinical trial material, enrolling sufficient patients in a timely manner, avoiding or mitigating serious or significant adverse patient reactions, and demonstrating efficacy of the product candidate in order to support a favorable risk versus benefit profile, among other considerations. The timing and success of our clinical trials, in particular, are also dependent on clinical sites and

regulatory authorities accepting each trial's protocol, statistical analysis plan, product characterization tests, and final clinical results. In addition, regulatory authorities may request additional information or data that is not readily available. Delays in our ability to respond to such requests would delay, and failure to adequately address concerns would prevent, our commercialization efforts. We have encountered in the past, and may encounter in the future, delays in initiating trial sites and enrolling patients into our clinical trials. Future enrollment delays will postpone the dates by which we expect to complete the impacted trials and the potential receipt of regulatory approval. There is no guarantee we will successfully initiate and/or complete our clinical trials.

Delays or difficulties in obtaining regulatory approvals or clearances for our product candidates may:

- adversely affect the marketing of any products we or our licensees or collaborators develop;
- impose significant additional costs on us or our licensees or collaborators;
- diminish any competitive advantages that we or our licensees or collaborators may attain;
- limit our ability to receive royalties and generate revenue and profits; and
- adversely affect our business prospects and ability to obtain financing.

Delays or failures in our receiving regulatory approval for our product candidates in a timely manner may result in us having to incur additional development expense and subject us to having to secure additional financing. As a result, we may not be able to commercialize them in the time frame anticipated, and our business will suffer.

Even if we or our partners receive marketing approval for our product candidates, such product approvals could be subject to restrictions or withdrawals. Regulatory requirements are subject to change. Further, even if we or our partners receive marketing approval, we may not receive sufficient coverage and adequate reimbursement for our products.

Regulatory authorities generally approve products for particular indications. If an approval is for a limited indication, this limitation reduces the size of the potential market for that product. Product approvals, once granted, are subject to continual review and periodic inspections by regulatory authorities. Our operations and practices are subject to regulation and scrutiny by the United States government, as well as governments of any other countries in which we do business or conduct activities. Later discovery of previously unknown problems or safety issues, and/or failure to comply with domestic or foreign laws, knowingly or unknowingly, can result in various adverse consequences, including, among other things, possible delay in approval or refusal to approve a product, warning letters, fines, injunctions, civil penalties, recalls or seizures of products, total or partial suspension of production, refusal of the government to renew marketing applications, complete withdrawal of a marketing application, corrective action requirements, and/or criminal prosecution, withdrawal of an approved product from the market, and/or exclusion from government health care programs. Such regulatory enforcement could have a direct and negative impact on the product for which approval is granted and could have a negative impact on the approval of any pending applications for marketing approval of new drugs or supplements to approved applications.

Because we operate in a highly regulated industry, regulatory authorities could take enforcement action against us in connection with our licensees' or collaborators', and/or our business and marketing activities for various reasons. For example, the Foreign Corrupt Practices Act prohibits U.S. companies and their representatives from offering, promising, authorizing, or making payments to foreign governmental officials for the purpose of obtaining or retaining business abroad.

From time to time, new legislation is passed into law that could significantly change the statutory provisions governing the approval, manufacturing, and marketing of products regulated by the FDA and other foreign health authorities. Additionally, regulations and guidance are often revised or reinterpreted by health agencies in ways that may significantly affect our business and our products. It is impossible to predict whether further legislative changes will be enacted, or whether regulations, guidance, or interpretations will change, and what the impact of such changes, if any, may be. For example, the Patient Protection and Affordable Care Act and the Health Care and Education Affordability Reconciliation Act of 2010 (collectively the "ACA"), enacted in March 2010, substantially changed the

way healthcare is financed by both governmental and private insurers, and significantly impacted the pharmaceutical industry. With regard to pharmaceutical products, among other things, ACA is expected to expand, increase, and change the methodology regarding industry rebates for drugs covered under Medicaid programs; impose an annual, nondeductible fee on any entity that manufactures or imports specific branded prescription drugs and biologic agents, apportioned among those entities according to market share in certain government healthcare programs; expand eligibility criteria for Medicaid programs by, among other things, allowing states to offer Medicaid coverage to certain individuals with income at or below 133% of the federal poverty level; expand the entities eligible for discounts under the Public Health Service pharmaceutical pricing program; create a new Patient Centered Outcomes Research Institute to oversee, identify priorities in, and conduct comparative clinical effectiveness research, along with funding for such research; and make changes to the coverage requirements under the Medicare D program. Significant legislative changes to the ACA also appear possible in the 115th U.S. Congress under the Trump Administration.

We expect both government and private health plans to continue to require healthcare providers, including healthcare providers

that may one day purchase our products, to contain costs and demonstrate the value of the therapies they provide. Even if our product candidates are approved, the commercial success of our products will depend substantially on the extent to which they are covered by third-party payors, including government health authorities and private health insurers. In the United States, no uniform policy of coverage and reimbursement for products exists among third-party payors, and coverage and reimbursement for products can differ significantly from payor to payor. If coverage and reimbursement are not available, or reimbursement is available only to limited levels, we or our collaborators may not be able to successfully commercialize our product candidates.

New data from our research and development activities, and/or resource considerations could modify our strategy and result in the need to adjust our projections of timelines and costs of programs.

Because we are focused on novel technologies, our research and development activities, including our nonclinical studies and clinical trials, involve the ongoing discovery of new facts and the generation of new data, based on which we determine next steps for a relevant program. These developments can occur with varying frequency and constitute the basis on which our business is conducted. We make determinations on an ongoing basis as to which of these facts or data will influence timelines and costs of programs. We may not always be able to make such judgments accurately, which may increase the costs we incur attempting to commercialize our product candidates. We monitor the likelihood of success of our initiatives and we may need to discontinue funding of such activities if they do not prove to be commercially feasible, due to our limited resources.

We may need to successfully address a number of technological challenges in order to complete development of our product candidates. Moreover, these product candidates may not be effective in treating any disease or may prove to have undesirable or unintended side effects, toxicities, or other characteristics that may preclude our obtaining regulatory approvals or prevent or limit commercial use.

Risks Related to Intellectual Property Rights

If we are unable to obtain and enforce patent protection for our product candidates and related technology, our business could be materially harmed.

We rely upon a combination of patents, trade secret protection and confidentiality agreements to protect the intellectual property related to our product candidates and technology. Any disclosure to or misappropriation by third parties of our confidential proprietary information could enable competitors to duplicate or surpass our technological achievements, eroding our competitive position in the market. Our patent applications may not result in issued patents, and, even if issued, the patents may be challenged and invalidated. Moreover, our patents and patent applications may not be sufficiently broad to prevent others from practicing our technologies or developing competing products. We also face the risk that others may independently develop similar or alternative technologies or may design around our proprietary property.

Issued patents may be challenged, narrowed, invalidated or circumvented. In addition, court decisions may introduce uncertainty in the enforceability or scope of patents owned by biotechnology companies. The legal systems of certain countries do not favor the aggressive enforcement of patents, and the laws of foreign countries may not allow us to protect our inventions with patents to the same extent as the laws of the United States. Because patent applications in the United States and many foreign jurisdictions are typically not published until 18 months after filing, or in some cases not at all, and because publications of discoveries in scientific literature lag behind actual discoveries, we cannot be certain that we were the first to make the inventions claimed in our issued patents or pending patent applications, or that we were the first to file for protection of the inventions set forth in our patents or patent applications. As a result, we may not be able to obtain or maintain protection for certain inventions. Therefore, the enforceability and scope of our patents in the United States and in foreign countries cannot be predicted with certainty and, as a result, any patents that we own, or license may not provide sufficient protection against competitors. We may not be able to obtain or maintain patent protection from our pending patent applications, from those we may file in the future, or from those

we may license from third parties. Moreover, even if we are able to obtain patent protection, such patent protection may be of insufficient scope to achieve our business objectives.

Patent terms may be inadequate to protect our competitive position on our product candidates for an adequate amount of time. Patents have a limited lifespan. In the United States, the natural expiration of a patent is generally 20 years after its effective filing date. Various extensions may be available; however, the life of a patent, and the protection it affords, is limited. Without patent protection for our product candidates, we may be open to competition from biosimilar or generic versions of our product candidates. Furthermore, the product development timeline for biotechnology products is lengthy and it is possible that our issued patents covering our product candidates in the United States and other jurisdictions may expire prior to commercial launch. For example, if we encounter delays in our development efforts, including our clinical trials, the period of time during which we could market our product candidates under patent protection could be reduced.

Our strategy depends on our ability to identify and seek patent protection for our discoveries. This process is expensive and time consuming, and we and our current or future licensors or licensees may not be able to file and prosecute all necessary or desirable

patent applications at a reasonable cost or in a timely manner or in all jurisdictions where protection may be commercially advantageous. It is also possible that we or our current licensors or licensees, or any future licensors or licensees, may not identify patentable aspects of inventions made in the course of development and commercialization activities in time to obtain patent protection on them. Therefore, these and any of our patents and applications may not be prosecuted and enforced in a manner consistent with the best interests of our business. Defects of form in the preparation or filing of our patents or patent applications may exist, or may arise in the future, for example with respect to proper priority claims, inventorship, etc. If we or our current licensors or licensees, or any future licensors or licensees, fail to establish, maintain or protect such patents and other intellectual property rights, such rights may be reduced or eliminated. If our current licensors or licensees, or any future licensors or licensees, are not fully cooperative or disagree with us as to the prosecution, maintenance or enforcement of any patent rights, such patent rights could be compromised. If there are material defects in the form or preparation of our patents or patent applications, such patents or applications may be invalid and unenforceable. Despite our efforts to protect our proprietary rights, unauthorized parties may be able to obtain and use information that we regard as proprietary. The issuance of a patent does not ensure that it is valid or enforceable, so even if we obtain patents, they may not be valid or enforceable against third parties. In addition, the issuance of a patent does not give us the right to practice the patented invention. Third parties may have blocking patents that could prevent us from marketing our own patented product and practicing our own patented technology. Any of these outcomes could impair our ability to prevent competition from third parties, which may have an adverse impact on our business.

The patent landscape in the field of therapeutic antibody development, manufacture and commercialization is crowded. For example, we are aware of third party patents directed to methods for identifying and producing therapeutic antibodies. We are also aware of third party patents directed to antibodies to numerous targets for which we also seek to identify, develop, and commercialize antibodies. For example, some patents claim antibodies based on competitive binding with existing antibodies, some claim antibodies based on specifying sequence or other structural information, and some claim various methods of discovery, production, or use of such antibodies.

These or other third-party patents could impact our freedom to operate in relation to our technology platforms, as well as in relation to development and commercialization of antibodies identified by us as therapeutic candidates. As we discover and develop our candidate antibodies, we will continue to conduct analyses of these third-party patents to determine whether we believe we might infringe them, and if so, whether they would be likely to be deemed valid and enforceable if challenged. If we determine that a license for a given patent or family of patents is necessary or desirable, there can be no guarantee that a license would be available on favorable terms, or at all. Inability to obtain a license on favorable terms, should such a license be determined to be necessary or desirable, could, without limitation, result in increased costs to design around the third-party patents, delay product launch, or result in cancellation of the affected program or cessation of use of the affected technology.

Third parties may also seek to market biosimilar versions of any approved products. Alternatively, third parties may seek approval to market their own products similar to or otherwise competitive with our products. In these circumstances, we may need to defend and/or assert our patents, including by filing lawsuits alleging patent infringement. In any of these types of proceedings, a court or agency with jurisdiction may find our patents invalid and/or unenforceable. Even if we have valid and enforceable patents, these patents still may not provide protection against competing products or processes sufficient to achieve our business objectives.

We own, co-own or have exclusive rights to approximately 35 issued United States patents and approximately 120 issued foreign patents. We also own, co-own or have exclusive rights to approximately 30 pending United States patent applications and approximately 125 pending foreign patent applications. However, our patents may not protect us against our competitors. Our patent positions, and those of other biopharmaceutical, pharmaceutical and biotechnology companies, are generally uncertain and involve complex legal, scientific, and factual questions. The standards which the United States Patent and Trademark Office (“USPTO”) uses to grant patents, and the standards which courts use to interpret patents, are not always applied predictably or uniformly and can change, particularly as new technologies develop. Consequently, the level of protection, if any, that will be provided by our patents if we

attempt to enforce them and they are challenged, is uncertain. In addition, the type and extent of patent claims that will be issued to us in the future is uncertain. Any patents that are issued may not contain claims that permit us to stop competitors from using similar technology.

Through our acquisitions of 4-AB, PhosImmune and certain assets of Celexion, we own, co-own, or have exclusive rights to a number of patents and patent applications directed to various methods and compositions, including methods for identifying therapeutic antibodies and product candidates arising out of such entities' technology platforms. In particular, we own patents and patent applications relating to our Retrocyte Display™ technology platform, a high throughput antibody expression platform for the identification of fully-human and humanized monoclonal antibodies. This patent family is projected to expire between 2029 and 2031. Through our acquisition of PhosImmune, we own, co-own, or have exclusive rights to patents and patent applications directed to various methods and compositions, including a patent directed to methods for identifying phosphorylated proteins using mass spectrometry. This patent is projected to expire in 2023. We also own patents and patent applications relating to the SECANT® platform, a platform used for the generation of novel monoclonal antibodies. This patent family is projected to expire between 2028 and 2029. In addition, as we advance our research and development efforts with our institutional and corporate collaborators, we are

seeking patent protection for newly identified therapeutic antibodies and product candidates. We can provide no assurance that any of our patents, including the patents that we acquired or in-licensed in connection with our acquisitions of 4-AB, PhosImmune and certain assets of Celexion, will have commercial value, or that any of our existing or future patent applications, including the patent applications that we acquired or in-licensed in connection with our acquisitions of 4-AB, PhosImmune and certain assets of Celexion, will result in the issuance of valid and enforceable patents.

Our issued patents covering Prophage vaccine and methods of use thereof, alone or in combination with other agents, expired or will expire at various dates between 2015 and 2024. In particular, our issued U.S. patents covering Prophage composition of matter expired in 2015. In addition, our issued patents covering QS-21 Stimulon composition of matter expired in 2008. We continue to explore means of extending the life cycle of our patent portfolio.

The patent position of biopharmaceutical, pharmaceutical or biotechnology companies, including ours, is generally uncertain and involves complex legal and factual considerations. The standards which the USPTO and its foreign counterparts use to grant patents are not always applied predictably or uniformly and can change. There is also no uniform, worldwide policy regarding the subject matter and scope of claims granted or allowable in biopharmaceutical, pharmaceutical or biotechnology patents. The laws of some foreign countries do not protect proprietary information to the same extent as the laws of the United States, and many companies have encountered significant problems and costs in protecting their proprietary information in these foreign countries. Outside the United States, patent protection must be sought in individual jurisdictions, further adding to the cost and uncertainty of obtaining adequate patent protection outside of the United States. Accordingly, we cannot predict whether additional patents protecting our technology will issue in the United States or in foreign jurisdictions, or whether any patents that do issue will have claims of adequate scope to provide competitive advantage. Moreover, we cannot predict whether third parties will be able to successfully obtain claims or the breadth of such claims. The allowance of broader claims may increase the incidence and cost of patent interference proceedings, opposition proceedings, post-grant review, inter partes review, and/or reexamination proceedings, the risk of infringement litigation, and the vulnerability of the claims to challenge. On the other hand, the allowance of narrower claims does not eliminate the potential for adversarial proceedings and may fail to provide a competitive advantage. Our issued patents may not contain claims sufficiently broad to protect us against third parties with similar technologies or products or provide us with any competitive advantage.

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

Third parties may infringe or misappropriate our intellectual property, including our existing patents, patents that may issue to us in the future, or the patents of our licensors or licensees to which we have a license. As a result, we may be required to file infringement claims to stop third-party infringement or unauthorized use. Further, we may not be able to prevent, alone or with our licensors or licensees, misappropriation of our intellectual property rights, particularly in countries where the laws may not protect those rights as fully as in the United States.

If we or one of our licensors or licensees were to initiate legal proceedings against a third party to enforce a patent covering our product candidates, the defendant could counterclaim that the patent covering our product candidates is invalid and/or unenforceable. In patent litigation in the United States, defendant counterclaims alleging invalidity and/or unenforceability are commonplace, and there are numerous grounds upon which a third party can assert invalidity or unenforceability of a patent.

In addition, within and outside of the United States, there has been a substantial amount of litigation and administrative proceedings, including interference and reexamination proceedings before the USPTO or oppositions and other comparable proceedings in various foreign jurisdictions, regarding patent and other intellectual property rights in the biopharmaceutical industry. Recently, the AIA introduced new procedures, including inter partes review

and post grant review. These procedures may be used by competitors to challenge the scope and/or validity of our patents, including those that patents perceived by our competitors as blocking entry into the market for their products, and the outcome of such challenges.

Even after they have been issued, our patents and any patents which we license may be challenged, narrowed, invalidated or circumvented. If our patents are invalidated or otherwise limited or will expire prior to the commercialization of our product candidates, other companies may be better able to develop products that compete with ours, which could adversely affect our competitive business position, business prospects and financial condition.

The following are non-exclusive examples of litigation and other adversarial proceedings or disputes that we could become a party to involving our patents or patents licensed to us:

- we or our collaborators may initiate litigation or other proceedings against third parties to enforce our patent rights;
- third parties may initiate litigation or other proceedings seeking to invalidate patents owned by or licensed to us or to obtain a declaratory judgment that their product or technology does not infringe our patents or patents licensed to us;

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- third parties may initiate opposition proceedings, post-grant review, inter partes review, or reexamination proceedings challenging the validity or scope of our patent rights, requiring us or our collaborators and/or licensors or licensees to participate in such proceedings to defend the validity and scope of our patents;
- there may be a challenge or dispute regarding inventorship or ownership of patents currently identified as being owned by or licensed to us;
- the USPTO may initiate an interference or derivation proceeding between patents or patent applications owned by or licensed to us and those of our competitors, requiring us or our collaborators and/or licensors or licensees to participate in an interference or derivation proceeding to determine the priority of invention, which could jeopardize our patent rights; or
 - third parties may seek approval to market biosimilar versions of our future approved products prior to expiration of relevant patents owned by or licensed to us, requiring us to defend our patents, including by filing lawsuits alleging patent infringement.

These lawsuits and proceedings would be costly and could affect our results of operations and divert the attention of our managerial and scientific personnel. There is a risk that a court or administrative body could decide that our patents are invalid or not infringed by a third party's activities, or that the scope of certain issued claims must be further limited. An adverse outcome in a litigation or proceeding involving our own patents could limit our ability to assert our patents against these or other competitors, affect our ability to receive royalties or other licensing consideration from our licensees, and may curtail or preclude our ability to exclude third parties from making, using and selling similar or competitive products. An adverse outcome may also put our pending patent applications at risk of not issuing, or issuing with limited and potentially inadequate scope to cover our product candidates. The outcome following legal assertions of invalidity and unenforceability is unpredictable. With respect to the validity question, for example, we cannot be certain that there is no invalidating prior art, of which we and the patent examiner were unaware during prosecution. Additionally, it is also possible that prior art of which we are aware, but which we do not believe affects the validity or enforceability of a claim, may, nonetheless, ultimately be found by a court of law or an administrative panel to affect the validity or enforceability of a claim, for example, if a priority claim is found to be improper. If a defendant were to prevail on a legal assertion of invalidity and/or unenforceability, we could lose at least part, and perhaps all, of the patent protection on our relevant product candidates. Such a loss of patent protection could have a material adverse impact on our business.

Furthermore, because of the substantial amount of discovery required in connection with intellectual property litigation or administrative proceedings, there is a risk that some of our confidential information could be compromised by disclosure. In addition, during the course of litigation or administrative proceedings, there could be public announcements of the results of hearings, motions or other interim proceedings or developments or public access to related documents. If investors perceive these results to be negative, the market price for our common stock could be significantly harmed. Any of these occurrences could adversely affect our competitive business position, business prospects, and financial condition.

Intellectual property rights do not necessarily address all potential threats to our competitive advantage. The degree of future protection for our proprietary rights is uncertain because legal means afford only limited protection and may not adequately protect our rights or permit us to gain or keep our competitive advantage. For example:

- others may be able to develop a platform that is similar to, or better than, ours in a way that is not covered by the claims of our patents;
- others may be able to make compounds that are similar to our product candidates but that are not covered by the claims of our patents;
- we might not have been the first to make the inventions covered by patents or pending patent applications;
- we might not have been the first to file patent applications for these inventions;
- any patents that we obtain may not provide us with any competitive advantages or may ultimately be found invalid or unenforceable; or
- we may not develop additional proprietary technologies that are patentable.

Our commercial success depends significantly on our ability to operate without infringing the patents and other proprietary rights of third parties.

Our success will depend in part on our ability to operate without infringing the proprietary rights of third parties. Other entities may have or obtain patents or proprietary rights that could limit our ability to make, use, sell, offer for sale or import our future approved products or impair our competitive position. In particular the patent landscape around the discovery, development, manufacture and commercial use of our pre-clinical CPM antibody programs and therapeutic antibodies is crowded.

Third parties may have or obtain valid and enforceable patents or proprietary rights that could block us from developing product candidates using our technology. Our failure to obtain a license to any technology that we require may materially harm our business, financial condition and results of operations. Moreover, our failure to maintain a license to any technology that we require may also materially harm our business, financial condition, and results of operations. Furthermore, we would be exposed to a threat of litigation.

In the biopharmaceutical industry, significant litigation and other proceedings regarding patents, patent applications, trademarks and other intellectual property rights have become commonplace. The types of situations in which we may become a party to such litigation or proceedings include:

- we or our collaborators may initiate litigation or other proceedings against third parties seeking to invalidate the patents held by those third parties or to obtain a judgment that our products or processes do not infringe those third parties' patents;
- if our competitors file patent applications that claim technology also claimed by us or our licensors or licensees, we or our licensors or licensees may be required to participate in interference, derivation or other proceedings to determine the priority of invention, which could jeopardize our patent rights and potentially provide a third party with a dominant patent position;
- if third parties initiate litigation claiming that our processes or products infringe their patent or other intellectual property rights, we and our collaborators will need to defend against such proceedings; and
- if a license to necessary technology is terminated, the licensor may initiate litigation claiming that our processes or products infringe or misappropriate their patent or other intellectual property rights and/or that we breached our obligations under the license agreement, and we and our collaborators would need to defend against such proceedings.

These lawsuits would be costly and could affect our results of operations and divert the attention of our management and scientific personnel. There is a risk that a court would decide that we or our collaborators are infringing the third party's patents and would order us or our collaborators to stop the activities covered by the patents. In that event, we or our collaborators may not have a viable alternative to the technology protected by the patent and may need to halt work on the affected product candidate or cease commercialization of an approved product. In addition, there is a risk that a court will order us or our collaborators to pay the other party damages. An adverse outcome in any litigation or other proceeding could subject us to significant liabilities to third parties and require us to cease using the technology that is at issue or to license the technology from third parties. We may not be able to obtain any required licenses on commercially acceptable terms or at all. Any of these outcomes could have a material adverse effect on our business.

The biopharmaceutical industry has produced a significant number of patents, and it may not always be clear to industry participants, including us, which patents cover various types of products or methods of use. The coverage of patents is subject to interpretation by the courts, and the interpretation is not always uniform or predictable. If we are sued for patent infringement, we would need to demonstrate that our products or methods either do not infringe the patent claims of the relevant patent or that the patent claims are invalid, and we may not be able to do this. Proving invalidity is difficult. For example, in the United States, proving invalidity requires a showing of clear and convincing evidence to overcome the presumption of validity enjoyed by issued patents. Even if we are successful in these proceedings, we may incur substantial costs and divert management's time and attention in pursuing these proceedings, which could have a material adverse effect on us. If we are unable to avoid infringing the patent rights of others, we may be required to seek a license, defend an infringement action or challenge the validity of the patents in court. Patent litigation is costly and time consuming. We may not have sufficient resources to bring these actions to a successful conclusion. In addition, if we do not obtain a license, develop or obtain non-infringing technology, fail to defend an infringement action successfully or have infringed patents declared invalid, we may incur substantial monetary damages, encounter significant delays in bringing our product candidates to market and be precluded from manufacturing or selling our product candidates.

The cost of any patent litigation or other proceeding, even if resolved in our favor, could be substantial. Some of our competitors may be able to sustain the cost of such litigation and proceedings more effectively than we can because of

their substantially greater resources. Uncertainties resulting from the initiation and continuation of patent litigation or other proceedings could have a material adverse effect on our ability to compete in the marketplace. Patent litigation and other proceedings may also absorb significant management time.

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

We are currently party to various intellectual property license agreements. These license agreements impose, and we expect that future license agreements may impose, various diligence, milestone payment, royalty, insurance and other obligations on us. These licenses typically include an obligation to pay an upfront payment, yearly maintenance payments and royalties on sales. If we fail to comply with our obligations under the licenses, the licensors may have the right to terminate their respective license agreements, in

which event we might not be able to market any product that is covered by the agreements. Termination of the 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, which could adversely affect our competitive business position and harm our business.

If we are unable to protect the confidentiality of our proprietary information, the value of our technology and products could be adversely affected.

In addition to patent protection, we also rely on other proprietary rights, including protection of trade secrets, and other proprietary information. To maintain the confidentiality of trade secrets and proprietary information, we enter into confidentiality agreements with our employees, consultants, collaborators and others upon the commencement of their relationships with us. These agreements require that all confidential information developed by the individual or made known to the individual by us during the course of the individual's relationship with us be kept confidential and not disclosed to third parties. Our agreements with employees and our personnel policies also provide that any inventions conceived by the individual in the course of rendering services to us shall be our exclusive property. However, we may not obtain these agreements in all circumstances, and individuals with whom we have these agreements may not comply with their terms. Thus, despite such agreement, such inventions may become assigned to third parties. In the event of unauthorized use or disclosure of our trade secrets or proprietary information, these agreements, even if obtained, may not provide meaningful protection, particularly for our trade secrets or other confidential information. To the extent that our employees, consultants or contractors use technology or know-how owned by third parties in their work for us, disputes may arise between us and those third parties as to the rights in related inventions. To the extent that an individual who is not obligated to assign rights in intellectual property to us is rightfully an inventor of intellectual property, we may need to obtain an assignment or a license to that intellectual property from that individual, or a third party or from that individual's assignee. Such assignment or license may not be available on commercially reasonable terms or at all.

Adequate remedies may not exist in the event of unauthorized use or disclosure of our proprietary information. The disclosure of our trade secrets would impair our competitive position and may materially harm our business, financial condition and results of operations. Costly and time-consuming litigation could be necessary to enforce and determine the scope of our proprietary rights, and failure to maintain trade secret protection could adversely affect our competitive business position. In addition, others may independently discover or develop our trade secrets and proprietary information, and the existence of our own trade secrets affords no protection against such independent discovery.

As is common in the biopharmaceutical industry, we employ individuals who were previously or concurrently employed at research institutions and/or other biopharmaceutical, biotechnology or pharmaceutical companies, including our competitors or potential competitors. We may be subject to claims that these employees, or we, have inadvertently or otherwise used or disclosed trade secrets or other proprietary information of their former employers, or that patents and applications we have filed to protect inventions of these employees, even those related to one or more of our product candidates, are rightfully owned by their former or concurrent employer. Litigation may be necessary to defend against these claims. Even if we are successful in defending against these claims, litigation could result in substantial costs and be a distraction to management.

Obtaining and maintaining our patent protection depends on compliance with various procedural, documentary, fee payment and other requirements imposed by governmental patent agencies, and our patent protection could be reduced or eliminated for non-compliance with these requirements.

Periodic maintenance fees, renewal fees, annuity fees and various other governmental fees on patents and/or applications will be due to the USPTO and various foreign patent offices at various points over the lifetime of our patents and/or applications. We have systems in place to remind us to pay these fees, and we rely on our outside counsel or service providers to pay these fees when due. Additionally, the USPTO and various foreign patent offices

require compliance with a number of procedural, documentary, fee payment and other similar provisions during the patent application process. We employ reputable law firms and other professionals to help us comply, and in many cases, an inadvertent lapse can be cured by payment of a late fee or by other means in accordance with rules applicable to the particular jurisdiction. However, there are situations in which noncompliance can result in abandonment or lapse of the patent or patent application, resulting in partial or complete loss of patent rights in the relevant jurisdiction. If such an event were to occur, it could have a material adverse effect on our business. In addition, we are responsible for the payment of patent fees for patent rights that we have licensed from other parties.

If any licensor of these patents does not itself elect to make these payments, and we fail to do so, we may be liable to the licensor for any costs and consequences of any resulting loss of patent rights.

Changes in U.S. patent law could diminish the value of patents in general, thereby impairing our ability to protect our product candidates.

Obtaining and enforcing patents in the biopharmaceutical industry involves both technological and legal complexity, and

therefore, is costly, time-consuming and inherently uncertain. In addition, the United States has recently enacted and is currently implementing wide-ranging patent reform legislation. Further, recent U.S. Supreme Court rulings have either narrowed the scope of patent protection available in certain circumstances or weakened the rights of patent owners in certain situations. In addition to increasing uncertainty with regard to our ability to obtain patents in the future, this combination of events has created uncertainty with respect to the value of patents, once obtained.

For our U.S. patent applications containing a claim not entitled to priority before March 16, 2013, there is a greater level of uncertainty in the patent law. In September 2011, the Leahy-Smith America Invents Act, or the American Invents Act (“AIA”), was signed into law. The AIA includes a number of significant changes to U.S. patent law, including provisions that affect the way patent applications will be prosecuted and may also affect patent litigation. The USPTO is currently developing regulations and procedures to govern administration of the AIA, and many of the substantive changes to patent law associated with the AIA. It is not clear what other, if any, impact the AIA will have on the operation of our business. Moreover, the AIA 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 have a material adverse effect on our business and financial condition.

An important change introduced by the AIA is that, as of March 16, 2013, the United States transitioned to a “first-inventor-to-file” system for deciding which party should be granted a patent when two or more patent applications are filed by different parties claiming the same invention. A third party that files a patent application in the USPTO after that date but before us could therefore be awarded a patent covering an invention of ours even if we had made the invention before it was made by the third party. This will require us to be cognizant going forward of the time from invention to filing of a patent application. Furthermore, our ability to obtain and maintain valid and enforceable patents depends on whether the differences between our technology and the prior art allow our technology to be patentable over the prior art. Since patent applications in the United States and most other countries are confidential for a period of time after filing, we cannot be certain that we were the first to either (i) file any patent application related to our product candidates or (ii) invent any of the inventions claimed in our patents or patent applications.

Among some of the other changes introduced by the AIA are changes that limit where a patentee may file a patent infringement suit and providing opportunities for third parties to challenge any issued patent in the USPTO. This applies to all of our U.S. patents, even those issued before March 16, 2013. Because of a lower evidentiary standard in USPTO proceedings compared to the evidentiary standard in United States federal court necessary to invalidate a patent claim, a third party could potentially provide evidence in a USPTO proceeding sufficient for the USPTO to hold a claim invalid even though the same evidence would be insufficient to invalidate the claim if first presented in a district court action. Accordingly, a third party may attempt to use the USPTO procedures to invalidate our patent claims that would not have been invalidated if first challenged by the third party as a defendant in a district court action.

We may be subject to claims that our employees, consultants or independent contractors have wrongfully used or disclosed confidential information of third parties.

We may have received confidential and proprietary information from third parties. In addition, we employ individuals who were previously employed at other biopharmaceutical, biotechnology or pharmaceutical companies. We may be subject to claims that we or our employees, consultants or independent contractors have inadvertently or otherwise improperly used or disclosed confidential information of these third parties or our employees’ former employers. Further, we may be subject to ownership disputes in the future arising, for example, from conflicting obligations of consultants or others who are involved in developing our product candidates. We may also be subject to claims that former employees, consultants, independent contractors, collaborators or other third parties have an ownership interest in our patents or other intellectual property. Litigation may be necessary to defend against these and other claims challenging our right to and use of confidential and proprietary information. If we fail in defending any such claims, in addition to paying monetary damages, we may lose our rights therein. Such an outcome could have a material adverse effect on our business. Even if we are successful in defending against these claims, litigation could result in

substantial cost and be a distraction to our management and employees.

We may not be able to protect our intellectual property rights throughout the world.

Filing, prosecuting and defending patents on our product candidates in all countries throughout the world would be prohibitively expensive. The requirements for patentability may differ in certain countries, particularly developing countries. For example, China has a heightened requirement for patentability, and specifically requires a detailed description of medical uses of a claimed drug. In addition, the laws of some foreign countries do not protect intellectual property rights to the same extent as laws in the United States. Consequently, we may not be able to prevent third parties from practicing our inventions in all countries outside the United States. Competitors may use our technologies in jurisdictions where we have not obtained patent protection to develop their own products and, further, may export otherwise infringing products to territories where we have patent protection, but enforcement on infringing activities is inadequate. These products may compete with our product candidates, and our patents or other intellectual property rights may not be effective or sufficient to prevent them from competing.

Many companies have encountered significant problems in protecting and defending intellectual property rights in foreign jurisdictions. The legal systems of certain countries, particularly certain developing countries, do not favor the enforcement of patents and other intellectual property protection, particularly those relating to biopharmaceuticals, which could make it difficult for us to stop the infringement of our patents or marketing of competing products in violation of our proprietary rights generally. Proceedings to enforce our patent rights in foreign jurisdictions could result in substantial costs and divert our efforts and attention from other aspects of our business, could put our patents at risk of being invalidated or interpreted narrowly and our patent applications at risk of not issuing, and could provoke third parties to assert claims against us. We may not prevail in any lawsuits that we initiate and the damages or other remedies awarded, if any, may not be commercially meaningful. In addition, certain countries in Europe and certain developing countries, including India and China, have compulsory licensing laws under which a patent owner may be compelled to grant licenses to third parties. In those countries, we may have limited remedies if our patents are infringed or if we are compelled to grant a license to our patents to a third party, which could materially diminish the value of those patents. This could limit our potential revenue opportunities. Accordingly, our efforts to enforce our intellectual property rights around the world may be inadequate to obtain a significant commercial advantage from the intellectual property that we own or license. Finally, our ability to protect and enforce our intellectual property rights may be adversely affected by unforeseen changes in foreign intellectual property laws.

Risks Related to Litigation

We may face litigation or regulatory investigations that could result in substantial damages and may divert management's time and attention from our business.

From time to time we may become a party to legal proceedings, claims and investigations that arise in the ordinary course of business such as, but not limited to, patent, employment, securities, commercial and environmental matters. While we currently believe that the ultimate outcome of any of these proceedings will not have a material adverse effect on our financial position, results of operations, or liquidity, litigation is subject to inherent uncertainty. Furthermore, litigation and regulatory investigations consume both cash and management attention.

We maintain property and general commercial insurance coverage as well as errors and omissions and directors and officers insurance policies. This insurance coverage may not be sufficient to cover us for future claims.

If we or our employees fail to comply with laws or regulations, it could adversely impact our reputation, business and stock price.

We are exposed to the risk of employee fraud or other misconduct. Misconduct by employees could include intentional and/or negligent 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 health care fraud and abuse, transparency, and/or data privacy and security laws and regulations, to report financial information or data accurately or to disclose unauthorized activities to us. In particular, sales, marketing and business arrangements in the healthcare industry are subject to extensive laws and regulations intended to prevent fraud, kickbacks, self-dealing and other abusive practices; to promote transparency; and to protect the privacy and security of patient data. These laws and regulations may restrict or prohibit a wide range of pricing, discounting, marketing and promotion, sales commission, customer incentive programs and other business arrangements.

While we have adopted a corporate compliance program, we may not be able to protect against all potential issues of noncompliance. Efforts to ensure that our business complies with all applicable healthcare laws and regulations will 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 laws and regulations.

Employee misconduct could also involve the improper use or disclosure of information obtained in the course of clinical trials, which could result in regulatory sanctions and serious harm to our reputation. In addition, during the

course of our operations, our directors, executives and employees may have access to material, nonpublic information regarding our business, our results of operations or potential transactions we are considering. We may not be able to prevent a director, executive or employee from trading in our common stock on the basis of, or while having access to, material, nonpublic information. If a director, executive or employee was to be investigated, or an action was to be brought against a director, executive or employee for insider trading, it could have a negative impact on our reputation and our stock price. Such a claim, with or without merit, could also result in substantial expenditures of time and money, and divert attention of our management team.

Product liability and other claims against us may reduce demand for our products and/or result in substantial damages.

We face an inherent risk of product liability exposure related to testing our product candidates in human clinical trials and manufacturing antibodies in our Berkeley, CA facility and may face even greater risks if we ever sell products commercially. An

individual may bring a product liability claim against us if one of our product candidates causes, or merely appears to have caused, an injury. Product liability claims may result in:

- regulatory investigations;
- injury to our reputation;
- withdrawal of clinical trial volunteers;
- costs of related litigation; and
- substantial monetary awards to plaintiffs; and
- decreased demand for any future products.

We manufacture the Prophage vaccines from a patient's cancer cells, and medical professionals must inject the vaccines into the same patient from which they were manufactured. A patient may sue us if a hospital, a shipping company, or we fail to receive the removed cancer tissue or deliver that patient's vaccine. We do not have any other insurance that covers loss of or damage to the Prophage vaccines or tumor material, and we do not know whether such insurance will be available to us at a reasonable price or at all. We have limited product liability coverage for use of our product candidates. Our product liability policy provides \$10.0 million aggregate coverage and \$10.0 million per occurrence coverage. This limited insurance coverage may be insufficient to fully cover us for future claims.

We are also subject to laws generally applicable to businesses, including but not limited to, federal, state and local wage and hour, employee classification, mandatory healthcare benefits, unlawful workplace discrimination and whistle-blowing. Any actual or alleged failure to comply with any regulation applicable to our business or any whistle-blowing claim, even if without merit, could result in costly litigation, regulatory action or otherwise harm our business, results of operations, financial condition, cash flow and future prospects.

If we do not comply with environmental laws and regulations, we may incur significant costs and potential disruption to our business.

We use or may use hazardous, infectious, and radioactive materials, and recombinant DNA in our operations, which have the potential of being harmful to human health and safety or the environment. We store these hazardous (flammable, corrosive, toxic), infectious, and radioactive materials, and various wastes resulting from their use, at our facilities pending use and ultimate disposal. We are subject to a variety of federal, state, and local laws and regulations governing use, generation, storage, handling, and disposal of these materials. We may incur significant costs complying with both current and future environmental health and safety laws and regulations. In particular, we are subject to regulation by the Occupational Safety and Health Administration, the Environmental Protection Agency, the Drug Enforcement Agency, the Department of Transportation, the Centers for Disease Control and Prevention, the National Institutes of Health, the International Air Transportation Association, and various state and local agencies. At any time, one or more of the aforementioned agencies could adopt regulations that may affect our operations. We are also subject to regulation under the Toxic Substances Control Act and the Resource Conservation Development programs.

Although we believe that our current procedures and programs for handling, storage, and disposal of these materials comply with federal, state, and local laws and regulations, we cannot eliminate the risk of accidents involving contamination from these materials. Although we have a workers' compensation liability policy, we could be held liable for resulting damages in the event of an accident or accidental release, and such damages could be substantially in excess of any available insurance coverage and could substantially disrupt our business.

Risks Related to our Common Stock

Provisions in our organizational documents could prevent or frustrate attempts by stockholders to replace our current management.

Our certificate of incorporation and bylaws contain provisions that could make it more difficult for a third party to acquire us without the consent of our Board of Directors. Our certificate of incorporation provides for a staggered board and removal of directors only for cause. Accordingly, stockholders may elect only a minority of our Board at any annual meeting, which may have the effect of delaying or preventing changes in management. In addition, under our certificate of incorporation, our Board of Directors may issue additional shares of preferred stock and determine the terms of those shares of stock without any further action by our stockholders. Our issuance of additional preferred stock could make it more difficult for a third party to acquire a majority of our outstanding voting stock and thereby effect a change in the composition of our Board of Directors. Our certificate of incorporation also provides that our stockholders may not take action by written consent. Our bylaws require advance notice of stockholder proposals and director nominations and permit only our president or a majority of the Board of Directors to call a special stockholder meeting. These

provisions may have the effect of preventing or hindering attempts by our stockholders to replace our current management. In addition, Delaware law prohibits a corporation from engaging in a business combination with any holder of 15% or more of its capital stock until the holder has held the stock for three years unless, among other possibilities, the board of directors approves the transaction. Our Board of Directors may use this provision to prevent changes in our management. Also, under applicable Delaware law, our Board of Directors may adopt additional anti-takeover measures in the future.

Our stock has historically had low trading volume, and its public trading price has been volatile.

During the period from our initial public offering on February 4, 2000 to December 31, 2017, and the year ended December 31, 2017, the closing price of our common stock has fluctuated between \$1.80 (or \$0.30 pre-reverse stock split) and \$315.78 (or \$52.63 pre-reverse stock split) per share and \$3.24 and \$5.37 per share, respectively. The average daily trading volume for the year ended December 31, 2017 was approximately 1,174,002 shares, while the average daily trading volume for the year ended December 31, 2016 was approximately 1,207,067. The market may experience significant price and volume fluctuations that are often unrelated to the operating performance of individual companies. In addition to general market volatility, many factors may have a significant adverse effect on the market price of our stock, including:

- continuing operating losses, which we expect over the next several years as we continue our development activities;
- announcements of decisions made by public officials or delays in any such announcements;
- results of our pre-clinical studies and clinical trials or delays in anticipated timing;
- delays in our regulatory filings or those of our partners;
- announcements of new collaboration agreements with strategic partners or developments by our existing collaboration partners;
- announcements of acquisitions;
- announcements of technological innovations, new commercial products, failures of products, or progress toward commercialization by our competitors or peers;
- failure to realize the anticipated benefits of acquisitions;
- developments concerning proprietary rights, including patent and litigation matters;
- publicity regarding actual or potential results with respect to product candidates under development;
 - quarterly fluctuations in our financial results, including our average monthly cash used in operating activities;
- variations in the level of expenses related to any of our product candidates or clinical development programs;
- additions or departures of key management or scientific personnel;
- conditions or trends in the biopharmaceutical, biotechnology and pharmaceutical industries generally;
- other events or factors, including those resulting from war, incidents of terrorism, natural disasters or responses to these events;
- changes in accounting principles;
- general economic and market conditions and other factors that may be unrelated to our operating performance or the operating performance of our competitors, including changes in market valuations of similar companies; and
- sales of common stock by us or our stockholders in the future, as well as the overall trading volume of our common stock.

In the past, securities class action litigation has often been brought against a company following a significant decline in the market price of its securities. This risk is especially relevant for us because many biopharmaceutical, biotechnology and pharmaceutical companies experience significant stock price volatility.

The trading market for our common stock will depend in part on the research and reports that securities or industry analysts publish about us or our business. If one or more of the analysts who covers us downgrades our stock, or publishes inaccurate or unfavorable research about our business, our stock price would likely decline. If one or more of these analysts ceases coverage of us or fails to publish reports on us regularly, demand for our stock could decrease, which could cause our stock price and trading volume to decline.

The sale of a significant number of shares could cause the market price of our stock to decline.

The sale by us or the resale by stockholders of a significant number of shares of our common stock could cause the market price of our common stock to decline. As of February 28, 2018, we had 102,556,797 shares of common stock outstanding. All of these shares are eligible for sale on Nasdaq, although certain of the shares are subject to sales volume and other limitations. We have filed registration statements to permit the sale of approximately 22,200,000 shares of common stock under our equity incentive plans, and to permit the sale of 1,500,000 shares of common stock under our 2015 Inducement Equity Plan. We have also filed registration statements to permit the sale of approximately 167,000 shares of common stock under our Employee Stock Purchase Plan, to permit the sale of 325,000 shares of common stock under our Directors' Deferred Compensation Plan, to permit the sale of approximately 31,100,319 shares of common stock pursuant to various private placement agreements and to permit the sale of up to 15,000,000 shares of our common stock pursuant to our Controlled Equity OfferingSM Sales Agreement. As of the date of filing, an aggregate of approximately 34,000,000 of these shares remained available for sale. In connection with our acquisition of 4-AB in February 2014, we are obligated to make contingent milestone payments to the former shareholders of 4-AB, payable in cash or shares of our common stock at our option, as follows (i) \$10.0 million upon our market capitalization exceeding \$750.0 million for 30 consecutive trading days prior to the earliest of (a) February 12, 2024 (b) the sale of 4-AB or (c) the sale of Agenus and (ii) \$10.0 million upon our market capitalization exceeding \$1.0 billion for 30 consecutive trading days prior to the earliest of (a) February 12, 2024, (b) the sale of 4-AB or (c) the sale of Agenus. In connection with our acquisition of PhosImmune in December 2015, we issued 1,631,521 shares of our common stock to the shareholders of PhosImmune and other third parties having a fair market value of approximately \$7.4 million at closing. In addition, we may be obligated in the future to pay certain contingent milestones payments, payable at our election in cash or shares of our common stock of up to \$35.0 million in the aggregate. We are also obligated to file registration statements covering any additional shares that may be issued to XOMA or the former shareholders of PhosImmune in the future pursuant to the terms of our agreements with XOMA and PhosImmune, respectively. The market price of our common stock may decrease based on the expectation of such sales. The market price of our common stock may decrease based on the expectation of such sales.

As of December 31, 2017, warrants to purchase approximately 4,351,450 shares of our common stock with a weighted average exercise price per share of \$9.01 were outstanding.

As of December 31, 2017, options to purchase 14,366,787 shares of our common stock with a weighted average exercise price per share of \$4.22 were outstanding. These options are subject to vesting that occurs over a period of up to four years following the date of grant. As of December 31, 2017, we had 8,164,576 vested options and 1,313,550 non-vested shares outstanding.

As of December 31, 2017, our outstanding shares of Series A-1 Convertible Preferred Stock were convertible into 333,333 shares of our common stock.

We may issue additional common stock, preferred stock, restricted stock units, or securities convertible into or exchangeable for our common stock. Furthermore, substantially all shares of common stock for which our outstanding stock options or warrants are exercisable are, once they have been purchased, eligible for immediate sale in the public market. The issuance of additional common stock, preferred stock, restricted stock units, or securities convertible into or exchangeable for our common stock or the exercise of stock options or warrants would dilute existing investors and could adversely affect the price of our securities. In addition, such securities may have rights senior to the rights of securities held by existing investors.

We do not intend to pay cash dividends on our common stock and, consequently your ability to obtain a return on your investment will depend on appreciation in the price of our common stock.

We have never declared or paid any cash dividend on our common stock and do not intend to do so in the foreseeable future. We currently anticipate that we will retain future earnings for the development, operation and expansion of our

business. Therefore, the success of an investment in shares of our common stock will depend upon any future appreciation in their value. There is no guarantee that shares of our common stock will appreciate in value or maintain their current value.

Our independent auditor's report for the fiscal year ended December 31, 2017 includes an explanatory paragraph regarding substantial doubt about our ability to continue as a going concern.

In its report accompanying our audited consolidated financial statements for the year ended December 31, 2017, our independent registered public accounting firm included an explanatory paragraph regarding concerns about our ability to continue as a going concern. The inclusion of a going concern explanatory paragraph may negatively impact the trading price of our common stock and make it more difficult, time consuming or expensive to obtain necessary financing. In the event we are unable to continue our operations, we may have to liquidate our assets and it is likely that investors will lose all or a part of their investment.

Failure to maintain effective internal controls in accordance with Section 404 of the Sarbanes-Oxley Act of 2002 and to comply with changing regulation of corporate governance and public disclosure could have a material adverse effect on our operating results and the price of our common stock.

The Sarbanes-Oxley Act of 2002 and rules adopted by the SEC and Nasdaq have resulted in significant costs to us. In particular, our efforts to comply with Section 404 of the Sarbanes-Oxley Act of 2002 and related regulations regarding the required assessment of our internal control over financial reporting, and our independent registered public accounting firm's audit of internal control over financial reporting, have required commitments of significant management time. We expect these commitments to continue.

Our internal control over financial reporting (as defined in Rules 13a-15 of the Exchange Act) is a process designed to provide reasonable assurance regarding the reliability of financial reporting and the preparation of our consolidated financial statements for external purposes in accordance with U.S. GAAP. Because of its inherent limitations, internal control over financial reporting may not prevent or detect all deficiencies or weaknesses in our financial reporting. While our management has concluded that there were no material weaknesses in our internal control over financial reporting as of December 31, 2017, our procedures are subject to the risk that our controls may become inadequate because of changes in conditions or as a result of a deterioration in compliance with such procedures. No assurance is given that our procedures and processes for detecting weaknesses in our internal control over financial reporting will be effective.

Changing laws, regulations and standards relating to corporate governance and public disclosure, are creating uncertainty for companies. Laws, regulations and standards are subject to varying interpretations in some cases due to their lack of specificity, and as a result, their application in practice may evolve over time as new guidance is provided, which could result in continuing uncertainty regarding compliance matters and higher costs caused by ongoing revisions to disclosure and governance practices. If we fail to comply with these laws, regulations and standards, our reputation may be harmed, and we might be subject to sanctions or investigation by regulatory authorities, such as the SEC. Any such action could adversely affect our operating results and the market price of our common stock.

Item 1B. Unresolved Staff Comments

None.

Item 2. Properties

We lease our manufacturing, research and development, and corporate offices in Lexington, Massachusetts occupying approximately 82,000 square feet. This lease agreement terminates in August 2023 with an option to renew for one additional ten-year period.

During December 2012 we entered into a commercial lease for approximately 5,600 square feet of office space in New York, New York for use as corporate offices that terminates in May 2020.

We also lease research and office facilities in Cambridge, United Kingdom. This lease terminates in November 2025. In 2017, we closed our offices in Basel, Switzerland, terminating the lease and consolidating these operations in the United Kingdom and United States.

In December 2015, we acquired a manufacturing facility with approximately 24,000 square feet in Berkeley, California to be used in the production and manufacture of antibody product candidates. In December 2015, we also entered into a commercial lease in Berkeley, California for approximately 10,900 square feet to be used for corporate offices which expires in December 2020. We also have a sublease in Berkeley, California for parking that expires in May 2020.

We believe substantially all of our property and equipment is in good condition and that we have sufficient capacity to meet our current operational needs. We do not anticipate experiencing significant difficulty in retaining occupancy of any of our manufacturing or office facilities and will do so through lease renewals prior to expiration or through replacing them with equivalent facilities.

Item 3. Legal Proceedings

We are not party to any material legal proceedings.

Item 4. Mine Safety Disclosures

Not applicable.

PART II

Item 5. Market for Registrant’s Common Equity, Related Stockholder Matters and Issuer Purchases of Equity Securities

Our common stock is currently listed on The Nasdaq Capital Market under the symbol “AGEN.”

The following table sets forth, for the periods indicated, the high and low sale prices per share of our common stock.

	High	Low
2016		
First Quarter	\$4.63	\$2.61
Second Quarter	4.82	2.97
Third Quarter	7.31	4.04
Fourth Quarter	7.49	3.71
2017		
First Quarter	4.54	3.69
Second Quarter	4.08	3.24
Third Quarter	5.37	3.48
Fourth Quarter	4.78	3.26

As of February 28, 2018, there were 402 holders of record and 27,634 beneficial holders of our common stock.

We have never paid cash dividends on our common stock, and we do not anticipate paying any cash dividends in the foreseeable future. We currently intend to retain future earnings, if any, for the future operation and expansion of our business. Any future payment of dividends on our common stock will be at the discretion of our Board of Directors and will depend upon, among other things, our earnings, financial condition, capital requirements, level of indebtedness, and other factors that our Board of Directors deem relevant.

Stock Performance

The following graph shows the cumulative total stockholder return on our common stock over the period spanning December 31, 2012 to December 31, 2017, as compared with that of the Nasdaq Stock Market (U.S. Companies) Index and the Nasdaq Biotechnology Index, based on an initial investment of \$100 in each on December 31, 2012. Total stockholder return is measured by dividing share price change plus dividends, if any, for each period by the share price at the beginning of the respective period and assumes reinvestment of dividends.

This stock performance graph shall not be deemed “filed” with the SEC or subject to Section 18 of the Exchange Act, nor shall it be deemed incorporated by reference in any of our filings under the Securities Act of 1933, as amended (the “Securities Act”).

COMPARISON OF CUMULATIVE TOTAL RETURN OF AGENUS INC.,

NASDAQ STOCK MARKET (U.S. COMPANIES) INDEX

AND NASDAQ BIOTECHNOLOGY INDEX

	12/31/2012	12/31/2013	12/31/2014	12/31/2015	12/31/2016	12/31/2017
Agenus Inc.	100.00	64.39	96.83	110.73	100.49	79.51
NASDAQ Stock Market (U.S. Companies) Index	100.00	138.32	156.85	165.84	178.28	228.63
NASDAQ Biotechnology Index	100.00	165.61	217.88	247.44	193.79	234.60

Item 6. Selected Financial Data

We have derived the condensed consolidated balance sheet data set forth below as of December 31, 2017 and 2016, and the condensed consolidated statement of operations data for each of the years in the three-year period ended December 31, 2017, from our audited consolidated financial statements included elsewhere in this Annual Report on Form 10-K.

You should read the selected condensed consolidated financial data in conjunction with “Management’s Discussion and Analysis of Financial Condition and Results of Operations,” our consolidated financial statements, and the notes to our consolidated financial statements included elsewhere in this Annual Report on Form 10-K.

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Changes in cash, cash equivalents, and short-term investments, total current assets, total assets, total current liabilities, long-term debt and stockholders' (deficit) equity in the periods presented below include the effects of the receipt of net proceeds from our debt offerings, equity offerings, the exercise of stock options, and employee stock purchases that totaled approximately \$81.5 million, \$3.4 million, \$220.4 million, \$57.0 million, and \$36.6 million in the years ended December 31, 2017, 2016, 2015, 2014, and 2013, respectively.

	For the Year Ended December 31,				
	2017	2016	2015	2014	2013
	(in thousands except per share data)				
Condensed Consolidated Statement of Operations Data:					
Revenue	\$42,877	\$22,573	\$24,817	\$6,977	\$3,045
Operating expenses:					
Cost of goods sold	—	—	—	-	(536)
Research and development	(116,125)	(94,971)	(70,444)	(22,349)	(13,005)
General and administrative	(33,741)	(33,126)	(28,370)	(21,250)	(14,484)
Contingent purchase price consideration fair value adjustment	3,188	(1,953)	(6,704)	(6,699)	—
Operating loss	(103,801)	(107,477)	(80,701)	(43,321)	(24,980)
Non-operating income (expense)	1,977	(2,202)	(5,968)	2,096	(2,673)
Interest expense, net	(18,868)	(17,316)	(6,599)	(1,261)	(2,420)
Loss before taxes	(120,692)	(126,995)	(93,268)	(42,486)	(30,073)
Income tax benefit (1)	—	—	5,387	—	—
Net loss	(120,692)	(126,995)	(87,881)	(42,486)	(30,073)
Dividends on Series A-1 convertible preferred stock	(206)	(204)	(203)	(204)	(3,159)
Net loss attributable to common stockholders	\$(120,898)	\$(127,199)	\$(88,084)	\$(42,690)	\$(33,232)
Net loss attributable to common stockholders per common share, basic and diluted	\$(1.23)	\$(1.46)	\$(1.13)	\$(0.71)	\$(1.12)
Weighted average number of common shares outstanding,					
basic and diluted	98,415	87,070	78,212	59,754	29,766

	As of December 31,				
	2017	2016	2015	2014	2013
	(in thousands)				
Condensed Consolidated Balance Sheet Data:					
Cash, cash equivalents and short-term investments	\$60,187	\$76,437	\$171,668	\$40,224	\$27,352
Total current assets	73,554	91,312	184,095	42,670	28,175
Total assets	138,402	156,986	242,228	74,527	34,835
Total current liabilities	56,438	40,851	28,934	9,229	10,296
Long-term debt, less current portion	142,385	130,542	114,326	4,769	5,384
Stockholders' (deficit) equity	(75,816)	(39,126)	70,728	23,018	(4,481)

(1) Given our history of incurring operating losses, no income tax benefit has been recognized in our consolidated statements of operations for the years ended December 31, 2017, 2016, 2014, and 2013 because of the loss before income taxes, and the need to recognize a valuation allowance on the portion of our deferred tax assets which will

not be offset by the reversal of deferred tax liabilities. For the year ended December 31, 2015, we recognized an income tax benefit as a result of the deferred tax liabilities recognized in connection with the PhosImmune and XOMA antibody manufacturing facility acquisitions.

Item 7. Management's Discussion and Analysis of Financial Condition and Results of Operations
Overview

We are a clinical-stage immuno-oncology ("I-O") company dedicated to becoming a leader in the discovery and development of innovative combination therapies and committed to bringing effective medicines to patients with cancer. Our business is designed to drive success in I-O through speed, innovation, and effective combination therapies. We have assembled fully integrated capabilities from novel target discovery, antibody generation, cell line development, and good manufacturing practice ("GMP") manufacturing together with a comprehensive portfolio consisting of antibody-based therapeutics, adjuvants and cancer vaccine platforms. We leverage our immune biology platforms to identify effective combination therapies for development and have developed productive partnerships to advance our innovation.

We are developing a comprehensive I-O portfolio driven by the following platforms and programs, which we intend to utilize individually and in combination:

- our antibody discovery platforms, including our Retrocyte Display™, SECAN[®] yeast display, and phage display technologies designed to drive the discovery of future CPM antibody candidates;
- our antibody candidate programs, including our CPM programs;
- our vaccine programs, including Prophage™, AutoSynVax™ and PhosPhoSyn™; and
- our saponin-based vaccine adjuvants, principally our QS-21 Stimulon[®] adjuvant, or QS-21 Stimulon.

We assess development, commercialization and partnering strategies for each of our product candidates periodically based on several factors, including pre-clinical and clinical trial results, competitive positioning and funding requirements and resources. We have formed collaborations with companies such as Incyte Corporation ("Incyte"), Merck Sharpe & Dohme and Recepta Biopharma SA ("Recepta"). Through these alliances, as well as our own internal programs, we currently have more than a dozen antibody programs in pre-clinical or early phase development, including our anti-CTLA-4 and anti-PD-1 antibody programs (both partnered with Recepta for certain South America territories) and anti-GITR and anti-OX40 antibody programs (both partnered with Incyte). In February 2017, we amended our collaboration agreement with Incyte to, among other things, convert the GITR and OX40 programs from profit-share to royalty-bearing programs. We are now eligible to receive royalties on global net sales at a flat 15% rate for each of these programs. There are no longer any profit-share programs remaining under the collaboration, and we are eligible to receive up to a total of \$510.0 million in future potential development, regulatory and commercial milestones across all programs in the collaboration. Pursuant to the amended agreement, we received accelerated milestone payments of \$20.0 million from Incyte related to the clinical development of INCAGN1876 (anti-GITR agonist) and INCAGN1949 (anti-OX40 agonist). Concurrent with the execution of the amendment, we and Incyte also entered into the Stock Purchase Agreement whereby Incyte purchased an additional 10 million shares of our common stock at \$6.00 per share, resulting in additional proceeds of \$60.0 million to us.

In addition to our antibody platforms and CPM programs, we are also advancing a series of vaccine programs to treat cancer. In January 2017, we announced a clinical trial collaboration with the National Cancer Institute ("NCI"), which is a double-blind, randomized controlled Phase 2 trial that is evaluating the effect of our autologous vaccine candidate, Prophage, in combination with pembrolizumab (Keytruda®, Merck & Co., Inc. ("Merck")) in patients with ndGBM. Under this collaboration, we are supplying Prophage, Merck is supplying pembrolizumab and the NCI and Brain Tumor Trials Collaborative member sites are recruiting patients and conducting the trial.

Our QS-21 Stimulon adjuvant is also partnered with GlaxoSmithKline ("GSK") and is a key component in multiple GSK vaccine programs. These programs are in various stages, with the most advanced being GSK's shingles program. In 2015, we monetized a portion of the future royalties we were contractually entitled to receive from GSK from sales of its shingles and malaria vaccines through a Note Purchase Agreement ("NPA") and received net proceeds of approximately \$78 million. In October 2017, GSK's shingles vaccine was approved in the United States by the FDA and granted marketing authorization in Canada by Health Canada and in November 2017, we exercised our option to issue the \$15.0 million in additional notes in accordance with the terms of the NPA. In January 2018, we entered into

a Royalty Purchase Agreement with Healthcare Royalty Partners III, L.P. and certain of its affiliates (together, “HCR”), pursuant to which HCR purchased 100% of our worldwide rights to receive royalties from GSK on GSK’s sales of vaccines containing our QS-21 Stimulon adjuvant. We used a portion of the upfront proceeds from HCR to redeem all of the notes issued pursuant to the Note Purchase Agreement, resulting in net proceeds to us of approximately \$28.0 million at closing. We do not incur clinical development costs for products partnered with GSK.

Our business activities include product research and development, intellectual property prosecution, manufacturing, regulatory and clinical affairs, corporate finance and development activities, and support of our collaborations. Our product candidates require clinical trials and approvals from regulatory agencies, as well as acceptance in the marketplace. Part of our strategy is to develop and

commercialize some of our product candidates by continuing our existing arrangements with academic and corporate collaborators and licensees and by entering into new collaborations.

In October 2017, we announced the launch of a subsidiary that is advancing our cell therapy business, AgenTus Therapeutics. The subsidiary is focused on the discovery, development, and commercialization of breakthrough “living drugs” to advance cures for cancer patients. AgenTus licenses intellectual property assets from Agenus, and has its own management and governance.

Our common stock is currently listed on The Nasdaq Capital Market under the symbol “AGEN.”

Our research and development expenses for the years ended December 31, 2017, 2016, and 2015, were \$116.1 million, \$95.0 million, and \$70.4 million, respectively. We have incurred significant losses since our inception. As of December 31, 2017, we had an accumulated deficit of \$1,026.5 million.

To date, we have financed our operations primarily through the sale of equity and debt securities. We believe that, based on our current plans and activities, including additional funding we anticipate from multiple sources between now and the end of the second quarter of 2018, including out-licensing and/or partnering opportunities, our working capital resources at December 31, 2017, along with the net proceeds of approximately \$28.0 million received from HCR in January 2018 in connection with our royalty transaction, will be sufficient to satisfy our liquidity requirements through the first quarter of 2019. We may attempt to raise additional funds by: (1) pursuing collaboration, out-licensing and/or partnering opportunities for our portfolio programs and product candidates with one or more third parties, (2) renegotiating third party agreements, (3) selling assets, (4) securing additional debt financing and/or (5) selling equity securities. Satisfying long-term liquidity needs may require the successful commercialization and/or substantial out-licensing or partnering arrangements for our antibody discovery platforms, CPM antibody programs, and HSP-based vaccines. Our long-term success will also be dependent on the successful identification, development and commercialization of potential other product candidates, each of which will require additional capital with no certainty of timing or probability of success. If we incur operating losses for longer than we expect and/or we are unable to raise additional capital, we may become insolvent and be unable to continue our operations.

Historical Results of Operations

Year Ended December 31, 2017 Compared to the Year Ended December 31, 2016

Revenue: We generated revenue of \$42.9 million and \$22.6 million during the years ended December 31, 2017 and 2016, respectively. Revenue primarily includes fees earned under our license agreements, including approximately \$14.6 million and \$16.2 million, respectively for the years ended December 31, 2017, and 2016, related to reimbursement of development costs under our Collaboration Agreement with Incyte. The increase in total revenue for the year ended December 31, 2017 is primarily attributable to the \$20.0 million in accelerated milestones, recognized as revenue during the twelve months ended December 31, 2017, related to the antibody candidates targeting G1TR and OX40 received in connection with the February 14, 2017 amendment to our License, Development and Commercialization Agreement with Incyte. During the years ended December 31, 2017 and 2016, we recorded revenue of \$3.1 million and \$3.5 million, respectively, from the amortization of deferred revenue.

Research and Development: Research and development expenses include the costs associated with our internal research and development activities, including compensation and benefits, occupancy costs, clinical manufacturing costs, contract research organization costs, costs of consultants, and related administrative costs. Research and development expense increased 22% to \$116.1 million for the year ended December 31, 2017 from \$95.0 million for the year ended December 31, 2016. Increased expenses in 2017 primarily relate to a \$17.6 million increase in third-party services and other expenses relating largely to the advancement of our CPM antibody programs, a \$3.3 million increase in payroll related costs primarily due to increased headcount, and a \$0.6 million increase in

depreciation expense, offset by a \$0.4 million decrease in expense for our foreign subsidiaries due to the closure of our facility in Basel, Switzerland, which decrease was partially offset by increased expenses attributable to Agenus UK.

General and Administrative: General and administrative expenses consist primarily of personnel costs, facility expenses, and professional fees. General and administrative expenses increased 2% to \$33.7 million for the year ended December 31, 2017 from \$33.1 million for the year ended December 31, 2016. Increased general and administrative expenses in 2017 primarily relate to a \$1.8 million increase in payroll related costs primarily due to increased headcount offset by a \$0.6 million decrease in professional fees due to the reduced use of consultants and a \$0.6 million decrease in expense for our foreign subsidiaries due to the closure of our facility in Basel, Switzerland, which decrease was partially offset by increased expenses attributable to Agenus UK.

Contingent purchase price consideration fair value adjustment: Contingent purchase price consideration fair value adjustment represents the change in the fair value of our contingent purchase price consideration during the year ended December 31, 2017, which resulted from changes in our market capitalization and share price and changes in the credit spread since the prior year end. The fair

value of our contingent purchase price consideration is based on estimates from a Monte Carlo simulation of our market capitalization and share price.

Non-operating income (expense): Non-operating income increased by \$4.2 million for the year ended December 31, 2017, from an expense of \$2.2 million for the year ended December 31, 2016, to income of \$2.0 million for the year ended December 31, 2017, primarily due to our increased foreign currency exchange gains in 2017 compared to losses in 2016.

Interest Expense, net: Interest expense, net increased to \$18.9 million for the year ended December 31, 2017 from \$17.3 million for the year ended December 31, 2016 due to the compounding of interest on our debt under our Note Purchase Agreement and the issuance of additional notes under our NPA in November 2017.

Year Ended December 31, 2016 Compared to the Year Ended December 31, 2015

Revenue: We generated revenue of \$22.6 million and \$24.8 million during the years ended December 31, 2016 and 2015, respectively. Revenue primarily includes fees earned under our license agreements, including approximately \$16.2 million and \$14.5 million, for the years ended December 31, 2016, and 2015, respectively, related to reimbursement of development costs under our Collaboration Agreement with Incyte. The decrease in revenue for the year ended December 31, 2016 is primarily attributable to decreased amortization of deferred revenue, offset by increased reimbursement of development costs under our Collaboration Agreement with Incyte. During the years ended December 31, 2016 and 2015, we recorded revenue of \$3.5 million and \$9.2 million, respectively, from the amortization of deferred revenue.

Research and development: Research and development expense increased 35% to \$95.0 million for the year ended December 31, 2016 from \$70.4 million for the year ended December 31, 2015. Increased expenses in 2016 primarily include the \$18.3 million increase in third-party services and other expenses relating largely to the advancement of our CPM antibody programs, a \$17.0 million increase in payroll related costs and share-based compensation due to increased headcount, and \$3.1 million increase in depreciation expense, offset by a \$13.2 million decrease in in-process research and development related to a 2015 asset acquisition.

General and administrative: General and administrative expenses increased 17% to \$33.1 million for the year ended December 31, 2016 from \$28.4 million for the year ended December 31, 2015. Increased general and administrative expenses in 2016 primarily relate to a \$2.0 million increase in payroll related expenses due to increased headcount and a \$1.9 million increase in share-based compensation.

Contingent purchase price consideration fair value adjustment: Contingent purchase price consideration fair value adjustment represents the change in the fair value of our contingent purchase price consideration during the year ended December 31, 2016, which resulted from changes in our market capitalization and share price and changes in the credit spread since the prior year end. The fair value of our contingent purchase price consideration is based on estimates from a Monte Carlo simulation of our market capitalization and share price.

Non-operating income (expense): Non-operating expense increased by \$3.8 million for the year ended December 31, 2016 which primarily represents our increased foreign currency exchange loss of \$1.7 million, offset by the absence of the prior year change in the fair value of our contingent royalty obligation of \$6.9 million, loss on extinguishment of our senior subordinated promissory notes issued in April 2013, and corresponding offset by the \$1.5 million gain on the purchase related to the antibody manufacturing facility acquisition from XOMA Corporation in December 2015.

Interest expense, net: Interest expense net increased to \$17.3 million for the year ended December 31, 2016 from \$6.6 million for the year ended December 31, 2015 due to the outstanding 2015 Subordinated Notes, issued in February

2015 and the Notes under our NPA, executed in September 2015.

Income tax benefit: For the year ended December 31, 2015, an income tax benefit arose from deferred tax liabilities recognized in connection with our PhosImmune and XOMA acquisitions during the year and relates to the resulting release of our existing valuation allowance on our deferred tax assets. There was no similar benefit for the year ended December 31, 2016.

Inflation

We believe that inflation has not had a material adverse effect on our business, results of operations, or financial condition to date.

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Research and Development Programs

For the year ended December 31, 2017, our research and development programs consisted largely of our CPM antibody programs as indicated in the following table (in thousands).

Research and Development Program	Product	For the Year Ended December 31,			Prior to 2015	Total
		2017	2016	2015		
Heat shock proteins for cancer	Prophage					
	and ASV	\$12,499	\$8,202	\$5,508	\$309,681	\$335,890
Antibody programs*	Various	95,656	83,919	63,290	13,422	256,287
Vaccine adjuvant	QS-21					
	Stimulon	222	77	142	13,657	14,098
Other research and development programs		7,748	2,772	1,504	66,318	78,342
Total research and development expenses		\$116,125	\$94,970	\$70,444	\$403,078	684,617

*Prior to 2014, costs were incurred by 4-AB, which we acquired in February 2014.

Research and development program costs include compensation and other direct costs plus an allocation of indirect costs, based on certain assumptions and our review of the status of each program. Our product candidates are in various stages of development and significant additional expenditures will be required if we start new clinical trials, encounter delays in our programs, apply for regulatory approvals, continue development of our technologies, expand our operations, and/or bring our product candidates to market. The total cost of any particular clinical trial is dependent on a number of factors such as trial design, length of the trial, number of clinical sites, number of patients, and trial sponsorship. The process of obtaining and maintaining regulatory approvals for new therapeutic products is lengthy, expensive, and uncertain. Because our CPM antibody programs are early stage, and because further development of HSP-based vaccines is dependent on clinical trial results, among other factors, we are unable to reliably estimate the cost of completing our research and development programs or the timing for bringing such programs to various markets or substantial partnering or out-licensing arrangements, and, therefore, when, if ever, material cash inflows are likely to commence. Active programs involving QS-21 Stimulon depend on our licensee successfully completing clinical trials, successfully manufacturing QS-21 Stimulon to meet demand, obtaining regulatory approvals and successfully commercializing product candidates containing QS-21 Stimulon.

Product Development Portfolio

Antibody Discovery Platforms and CPM Programs

Checkpoint antibodies regulate immune response against pathogens that invade the body and are achieving positive outcomes in a number of cancers that were untreatable only a few years ago. Two classes of checkpoint targets include:

1. inhibitory checkpoints that help suppress an immune response in order to prevent excessive immune reaction resulting in undesired inflammation and/or auto-immunity, and
2. stimulatory checkpoints that can enhance or amplify an antigen-specific immune response.

We possess a suite of antibody discovery platforms that are designed to drive the discovery of future CPM antibody candidates. We are planning to employ a variety of techniques to identify and optimize mono-specific and multi-specific antibody candidates, internally.

We and our partners currently have more than a dozen antibody programs in pre-clinical or early phase development, including our anti-CTLA-4 and anti-PD-1 antibody programs (both partnered with Recepta for certain South America territories) and anti-GITR and anti-OX40 antibody programs (both partnered with Incyte). For additional information regarding our antibody discovery platforms and checkpoint antibody program, please read Part I-Item 1. "Business" of this Annual Report on Form 10-K.

Prophage Vaccine Candidates

Our Prophage cancer vaccine candidate, HSPPC-96, is an autologous cancer vaccine therapy derived from cancer tissues that are surgically removed from an individual patient. As a result, a Prophage vaccine contains a broad sampling of potentially antigenic mutant proteins from each patient's own tumor. Prophage vaccines are designed to program the body's immune system to target only the specific cells that express those mutant antigens, thereby reducing the risk that the body's immune response against the tumor after

vaccination will also affect healthy tissue. Enhancing immune response using personalized vaccines, particularly in combination with CPMs, could be useful in cancers where a low number of mutant proteins leads to weakened immunogenicity.

To date, more than 1,000 patients have been treated with Prophage vaccines in clinical trials internationally, covering a broad range of cancer types, and no serious immune-mediated side effects have been observed. The results of these trials have been published and/or presented at scientific conferences. These results indicate observable clinical and/or immunological activity across many types of cancer.

In January 2017, we announced a clinical trial collaboration with the National Cancer Institute (“NCI”). The double-blind, randomized controlled Phase 2 trial is evaluating the effect of Prophage in combination with pembrolizumab (Keytruda®) in patients with ndGBM. The trial is being conducted by the Brain Tumor Trials Collaborative (“BTTC”), a consortium of top academic centers led by Dr. Mark Gilbert, Chief of the Neuro-Oncology Branch at the NCI Center for Cancer Research. The trial consists of two-arms with one arm receiving pembrolizumab as a monotherapy and a second arm receiving both Prophage and pembrolizumab in combination. Forty-five patients are being randomly assigned to each arm. Under this collaboration, we are supplying Prophage, Merck is supplying pembrolizumab (Keytruda®) and NCI and BTTC member sites are recruiting patients and conducting the trial. For additional information regarding regulatory risks and uncertainties, please read the risks identified under Part I-Item 1A. “Risk Factors” of this Annual Report on Form 10-K.

Neoantigen Vaccine Platforms

Mutation-based neo-epitopes, which will form the basis for the immunogens used in ASV, appear to be almost always particular to a given patient. Therefore, ASV is a largely individualized vaccine product candidate. With a small amount of a patient’s tumor as a sample, our ASV program is designed to utilize next generation sequencing technologies coupled with complex bioinformatics algorithms to identify mutations in a tumor’s DNA and RNA. Once these mutations have been identified, we plan to manufacture synthetic peptides encoding these neoepitopes, load these peptides on to our recombinant HSP70 and deliver a fully synthetic polyvalent vaccine to the patient. This program is based on the hypothesis that the HSP70 platform would shuttle the mutated peptides to sites where they could be recognized by the immune system and elicit a cytotoxic and helper T cell response in patients with cancer.

Biochemically based neoantigens, such as those arising from dysregulated phosphorylation of various proteins in malignant cells, can serve as a tumor fingerprint across a broad histology of tumors. Through the acquisition of PhosImmune, we have a portfolio of proprietary phosphorylated antigenic targets that can be used for therapeutic development. PSV is a vaccine candidate designed to induce immunity against this novel class of tumor specific neoepitopes. PSV is intended to induce cellular immunity to abnormal phosphopeptide neoepitopes that are characteristic of these various forms of cancer. Phosphopeptides (or phosphopeptide analogues) can be synthesized and complexed with HSP70, using an approach analogous to that used in the generation of our previous HerpV vaccine candidate. HerpV demonstrated good cellular and humoral responses to synthetic peptide immunogens complexed with HSP70 in a placebo-controlled Phase 2 study. We believe that similar responses could be obtained to phosphopeptide or phosphopeptide analogues bound to HSP70 when used as vaccines. Phosphorylation-based neoepitopes can apparently be found on specific types of cancer in many patients. Studies to optimize the immunogens to be used in PSV are ongoing. For additional information regarding our Neoantigen Vaccine Platforms, please read Part I-Item 1. “Business” of this Annual Report on Form 10-K.

QS-21 Stimulon Adjuvant

QS-21 Stimulon is an adjuvant, which is a substance added to a vaccine or other immunotherapy that is intended to enhance an immune response to the target antigens. QS-21 Stimulon is a natural product, a triterpene glycoside, or saponin, purified from the bark of the Chilean soapbark tree, *Quillaja saponaria*. QS-21 Stimulon has the ability to stimulate an antibody-mediated immune response and has also been shown to activate cellular immunity. It has

become a key component in the development of investigational preventive vaccine formulations across a wide variety of diseases. These studies have been carried out by academic institutions and pharmaceutical companies in the United States and internationally. A number of these studies have shown QS-21 Stimulon to be significantly more effective in stimulating immune responses than aluminum hydroxide or aluminum phosphate, the adjuvants most commonly used in approved vaccines in the United States today. For additional information regarding QS-21 Stimulon, please read Part I-Item 1. "Business" of this Annual Report on Form 10-K.

Liquidity and Capital Resources

We have incurred annual operating losses since inception, and we had an accumulated deficit of \$1,026.5 million as of December 31, 2017. We expect to incur significant losses over the next several years as we continue development of our technologies and product candidates, manage our regulatory processes, initiate and continue clinical trials, and prepare for potential commercialization of products. To date, we have financed our operations primarily through the sale of equity and debt securities, and

interest income earned on cash, cash equivalents, and short-term investment balances. From our inception through December 31, 2017, we have raised aggregate net proceeds of approximately \$923.9 million through the sale of common and preferred stock, the exercise of stock options and warrants, proceeds from our Employee Stock Purchase Plan, and the issuance of convertible and other notes.

In October 2017, we filed, and the Securities and Exchange Commission declared effective a Registration Statement on Form S-3 (file no. 333-221008) (the “2017 Registration Statement”), covering the offering of up to \$250 million of common stock, preferred stock, warrants, debt securities and units. The 2017 Registration Statement included a prospectus covering the offering, issuance and sale of up to 15 million shares of our common stock from time to time in “at-the-market offerings” pursuant to a Controlled Equity OfferingSM sales agreement (the “Sales Agreement”) entered into with Cantor Fitzgerald & Co. (the “Sales Agent”). As of December 31, 2017, we had 15 million shares available for sale under the Sales Agreement. In January 2018, we received net proceeds of approximately \$2.5 million from the sale of approximately 635,000 shares of our common stock in at-the-market offerings under the Sales Agreement.

In October 2017, we also exercised our right under that certain At Market Issuance Sales Agreement by and between us and MLV & Co. LLC dated as of October 10, 2014 (the “2014 ATM Program”) to terminate the 2014 ATM Program, which termination became effective upon effectiveness of the 2017 Registration Statement. We sold approximately 1.3 million shares of our common stock pursuant to the 2014 ATM Program during the year ended December 31, 2017 and received aggregate net proceeds of \$5.6 million.

As of December 31, 2017, we had debt outstanding of \$129.1 million in principal, and \$36.8 million in accrued interest. In February 2015, we issued subordinated notes in the aggregate principal amount of \$14.0 million with annual interest at 8% (the “2015 Subordinated Notes”). The 2015 Subordinated Notes are due in February 2020. In September 2015, we and our wholly-owned subsidiary Antigenics LLC (“Antigenics”) entered into the Note Purchase Agreement (“NPA”) with certain purchasers pursuant to which Antigenics issued, and we guaranteed, limited recourse notes in the aggregate principal amount of \$100.0 million, with an option to issue an additional \$15.0 million principal amount of limited recourse notes, which was exercised in November 2017. In January 2018, we entered into a Royalty Purchase Agreement with HCR whereby we received proceeds of \$190.0 million. We used \$161.9 million of these proceeds to redeem all of the notes issued pursuant to the Note Purchase Agreement. See Note 22 for additional information.

Our cash, cash equivalents, and short-term investments at December 31, 2017 were \$60.2 million, a decrease of \$16.3 million from December 31, 2016, principally as a result cash used in operations. We believe that, based on our current plans and activities, including additional funding we anticipate from multiple sources between now and the end of the second quarter of 2018, including out-licensing and/or partnering opportunities, our working capital resources at December 31, 2017, along with the net proceeds of approximately \$28.0 million received from HCR in January 2018 in connection with our royalty transaction, will be sufficient to satisfy our liquidity requirements through the first quarter of 2019. We continue to monitor the likelihood of success of our key initiatives and are prepared to discontinue funding of such activities if they do not prove to be feasible, restrict capital expenditures and/or reduce the scale of our operations.

Our future liquidity needs will be determined primarily by the success of our operations with respect to the progression of our product candidates and key development and regulatory events in the future. Potential sources of additional funding include: (1) pursuing collaboration, out-licensing and/or partnering opportunities for our portfolio programs and product candidates with one or more third parties, (2) renegotiating third party agreements, (3) selling assets, (4) securing additional debt financing and/or (5) selling equity securities. Satisfying long-term liquidity needs may require the successful commercialization and/or substantial out-licensing or partnering arrangements for our antibody discovery platforms, CPM antibody programs, and HSP-based vaccines. Our long-term success will also be dependent on the successful identification, development and commercialization of potential other product candidates, each of which will require additional capital with no certainty of timing or probability of success. If we incur operating losses for longer than we expect, and/or we are unable to raise additional capital, we may become insolvent

and be unable to continue our operations.

Our future cash requirements include, but are not limited to, supporting clinical trial and regulatory efforts and continuing our other research and development programs. Since inception, we have entered into various agreements with contract manufacturers, institutions, and clinical research organizations (collectively "third party providers") to perform pre-clinical activities and to conduct and monitor our clinical studies. Under these agreements, subject to the enrollment of patients and performance by the applicable third-party provider, we have estimated our total payments to be \$182.1 million over the term of the related activities. Through December 31, 2017, we have expensed \$130.2 million as research and development expenses and \$124.1 million has been paid under these agreements. The timing of expense recognition and future payments related to these agreements is subject to the enrollment of patients and performance by the applicable third-party provider. We have also entered into sponsored research agreements related to our product candidates that required payments of \$9.3 million, \$8.1 million of which have been paid as of December 31, 2017. We

plan to enter into additional agreements with third party providers as well as sponsored research agreements, and we anticipate significant additional expenditures will be required to initiate and advance our various programs.

Part of our strategy is to develop and commercialize some of our product candidates by continuing our existing collaboration arrangements with academic and collaboration partners and licensees and by entering into new collaborations. As a result of our collaboration agreements, we will not completely control the efforts to attempt to bring those product candidates to market. For example, our collaboration with Incyte for the development, manufacture and commercialization of CPM antibodies against certain targets is managed by a joint steering committee, which is controlled by Incyte.

Net cash used in operating activities for the years ended December 31, 2017 and 2016 was \$94.2 million and \$80.0 million, respectively. The first product containing QS-21 Stimulon was launched in the fourth quarter of 2017. We are generally entitled to royalties on sales by GSK of vaccines using QS-21 Stimulon for at least ten years after commercial launch, with some exceptions. In September 2015, we entered into the NPA and partially monetized the potential royalties we are entitled to receive from GSK. In January 2018 we entered into a Royalty Purchase Agreement with HCR, pursuant to which HCR purchased 100% of our worldwide rights to receive royalties from GSK on GSK's sales of vaccines containing our QS-21 Stimulon adjuvant. We used a portion of the upfront proceeds from HCR to redeem all of the notes issued pursuant to the NPA. Our future ability to generate cash from operations will depend on achieving regulatory approval and market acceptance of our product candidates, achieving benchmarks as defined in existing collaboration agreements, and our ability to enter into new collaborations. Please see the "Note Regarding Forward-Looking Statements" of this Annual Report on Form 10-K and the risks highlighted under Part I-Item 1A. "Risk Factors" of this Annual Report on Form 10-K.

The table below summarizes our contractual obligations as of December 31, 2017 (in thousands).

	Total	Payments by Period			
		Less than 1 Year	1-3 Years	3-5 Years	More than 5 Years
Long-term debt (1)	\$ 131,595	\$ 16,322	\$ 15,273	\$—	\$ 100,000
Operating leases (2)	14,212	3,179	4,718	3,846	2,469
Capital lease	720	288	432	—	—
Total	\$ 146,527	\$ 19,789	\$ 20,423	\$ 3,846	\$ 102,469

(1) Includes fixed interest payments. Under the terms of the NPA, interest accrues as 13.5%, compounded quarterly and may vary based on the timing of the royalty stream under our contract with GSK and therefore the table above excludes such interest which was approximately \$36.8 million as of December 31, 2017. In January 2018, we entered into a Royalty Purchase Agreement with HCR. We used a portion of the upfront proceeds from HCR to redeem all of the notes issued pursuant to the Note Purchase Agreement. See Note 22 for additional information.

(2) The leases and subleases for our properties expire at various times between 2018 and 2025.

Off-Balance Sheet Arrangements

At December 31, 2017, we had no off-balance sheet arrangements.

Critical Accounting Policies and Estimates

The SEC defines “critical accounting policies” as those that require the application of management’s most difficult, subjective, or complex judgments, often as a result of the need to make estimates about the effect of matters that are inherently uncertain and may change in subsequent periods.

The preparation of consolidated financial statements in conformity with U.S. generally accepted accounting principles requires us to make estimates and assumptions that affect the reported amounts of assets and liabilities and disclosures of contingent assets and liabilities at the date of the financial statements and the reported amounts of revenues and expenses during the reporting period. We base those estimates on historical experience and on various assumptions that are believed to be reasonable under the circumstances. Actual results could differ from those estimates.

The following listing is not intended to be a comprehensive list of all of our accounting policies. Our significant accounting policies are described in Note 2 of the notes to our consolidated financial statements contained elsewhere in this Annual Report on Form 10-K. In many cases, the accounting treatment of a particular transaction is dictated by U.S. generally accepted accounting principles, with no need for our judgment in its application. There are also areas in which our judgment in selecting an available alternative would not produce a materially different result. We have identified the following as our critical accounting policies.

Share-Based Compensation

We recognize share-based compensation expense in accordance with the fair value recognition provisions of the Financial Accounting Standards Board (“FASB”) Accounting Standards Codification (“ASC”) 718, Compensation—Stock Compensation. Compensation expense is recognized on a straight-line basis over the requisite service period of the award. Forfeitures are recognized as they occur.

Share-based awards granted to certain non-employees have been accounted for based on the fair value method of accounting in accordance with ASC 505-50, Equity- Equity-Based Payments to Non-Employees. As a result, the non-cash charge to operations for non-employee awards with vesting or other performance criteria is affected each reporting period by changes in the fair value of our common stock. Under the provisions of ASC 505-50, the change in fair value of vested awards issued to non-employees is reflected in the statement of operations each reporting period, until the options are exercised or expire.

Determining the appropriate fair value model and calculating the fair value of share-based awards requires the use of highly subjective assumptions, including the expected life of the share-based awards and stock price volatility. The assumptions used in calculating the fair value of share-based awards represent management’s best estimates, but these estimates involve inherent uncertainties and the application of management judgment. As a result, if factors change and we use different assumptions, our share-based compensation expense could be materially different in the future. For performance condition awards, we estimate the probability that the performance condition will be met. See Note 11 of the notes to our consolidated financial statements contained elsewhere in this Annual Report on Form 10-K for a further discussion on share-based compensation.

Revenue Recognition

Revenue recognized from collaborative agreements is based upon the provisions of ASC 605-25, Revenue Recognition—Multiple Element Arrangements, as amended by Accounting Standards Update 2009-13. License fees and royalties are recognized as they are earned. Non-refundable milestone payments that represent the completion of a separate earnings process are recognized as revenue when earned. Revenue for services under research and development contracts are recognized as the services are performed, or as clinical trial materials are provided.

Fair Value Measurements

In accordance with ASC 820, Fair Value Measurements and Disclosures, we measure fair value based on a hierarchy for inputs used in measuring fair value that maximizes the use of observable inputs and minimizes the use of unobservable inputs by requiring that observable inputs be used when available. The fair value hierarchy is broken down into three levels based on the source of inputs.

We measure our contingent purchase price considerations at fair value in accordance with ASC 825, Financial Instruments. The fair value contingent purchase price considerations are based on significant inputs not observable in the market, which require them to be reported as a Level 3 liability within the fair value hierarchy. The fair values of our 4-AB and PhosImmune contingent purchase price considerations are based on estimates from a Monte Carlo simulation of our market capitalization and share price, respectively. Market capitalization and share price were evolved using a geometric brownian motion, calculated daily for the life of the contingent purchase price consideration.

Business Combinations

In February 2014 and December 2015, we acquired all of the outstanding capital stock of 4-AB and PhosImmune, respectively in business combination transactions. In December 2015, we also acquired an antibody manufacturing pilot facility from XOMA Corporation which under the applicable accounting guidance is being accounted for as a

business combination. The acquisition method of accounting requires that the assets acquired and liabilities assumed be recorded as of the date of the merger or acquisition at their respective fair values with limited exceptions. Assets acquired and liabilities assumed in a business combination that arise from contingencies are recognized at fair value if fair value can reasonably be estimated. If the acquisition date fair value of an asset acquired or liability assumed that arises from a contingency cannot be determined, the asset or liability is recognized if probable and reasonably estimable; if these criteria are not met, no asset or liability is recognized. Fair value is defined as the exchange price that would be received for an asset or paid to transfer a liability (an exit price) in the principal or most advantageous market for the asset or liability in an orderly transaction between market participants on the measurement date. Accordingly, we may be required to value assets at fair value measures that do not reflect our intended use of those assets. Any excess of the purchase price (consideration transferred) over the estimated fair values of net assets acquired is recorded as goodwill. In the event the value of the net assets acquired exceeds the purchase price consideration, then a bargain purchase has occurred. The resulting bargain purchase on the transaction will be recognized as a gain in the period in which the acquisition was executed. The operating results of the acquired businesses are reflected in our consolidated financial statements after the date of the merger or acquisition. If we determine the assets acquired do not meet the definition of a business under the acquisition method of accounting, the transaction will be accounted for as

an acquisition of assets rather than a business combination and, therefore, no goodwill will be recorded. The fair values of intangible assets, including acquired in-process research and development (“IPR&D”), are determined utilizing information available near the merger or acquisition date based on expectations and assumptions that are deemed reasonable by management. The judgments made in determining estimated fair values assigned to assets acquired and liabilities assumed in a business combination, as well as asset lives, can materially affect the Company’s results of operations.

Acquired Intangible Assets, including IPR&D

IPR&D acquired in a business combination represents the fair value assigned to research and development assets that have not reached technological feasibility. The value assigned to acquired IPR&D is determined by estimating the costs to develop the acquired technology into commercially viable products, estimating the resulting revenue from the projects, and discounting the net cash flows to present value. The revenue and costs projections used to value acquired IPR&D are, as applicable, reduced based on the probability of success of developing a new drug. Additionally, the projections consider the relevant market sizes and growth factors, expected trends in technology, and the nature and expected timing of new product introductions by us and our competitors. The rates utilized to discount the net cash flows to their present value are commensurate with the stage of development of the projects and uncertainties in the economic estimates used in the projections. Upon the acquisition of IPR&D, we complete an assessment of whether our acquisition constitutes the purchase of a single asset or a group of assets. We consider multiple factors in this assessment, including the nature of the technology acquired, the presence or absence of separate cash flows, the development process and stage of completion, quantitative significance and our rationale for entering into the transaction.

We review amounts capitalized as acquired IPR&D for impairment at least annually, as of October 31, and whenever events or changes in circumstances indicate that the carrying value of the assets might not be recoverable. When performing our impairment assessment, we have the option to first assess qualitative factors to determine whether it is necessary to recalculate the fair value of our acquired IPR&D. If we elect this option and believe, as a result of the qualitative assessment, that it is more-likely-than-not that the fair value of our acquired IPR&D is less than its carrying amount, we calculate the fair value using the same methodology as described above. If the carrying value of our acquired IPR&D exceeds its fair value, then the intangible asset is written-down to its fair value. Alternatively, we may elect to bypass the qualitative assessment and immediately recalculate the fair value of our acquired IPR&D.

Finite-lived intangible asset are amortized over their useful life. We review finite-lived intangible assets for impairment whenever events or changes in circumstances indicate that the carrying value of the assets might not be recoverable.

Goodwill

Goodwill is tested at least annually for impairment on a reporting unit basis. We have concluded that we consist of a single operating segment and one reporting unit. We assess goodwill for impairment by performing a quantitative analysis to determine whether the fair value of our single reporting unit exceeds its carrying value. We perform our annual impairment test as of October 31 of each year and the first step of our impairment analysis compares the fair value to our net book value to determine if there is an indicator of impairment. Fair value is based on the quoted market price of our common stock to derive the market capitalization as of the date of the impairment test.

Recent Accounting Pronouncements

Refer to Note 2 to our consolidated financial statements included within Item 8 of this Annual Report on Form 10-K for a description of recent accounting pronouncements applicable to our business.

Item 7A. Quantitative and Qualitative Disclosures About Market Risk

Our primary market risk exposure is foreign currency exchange rate risk. International revenues and expenses are generally transacted by our foreign subsidiary and are denominated in local currency. Approximately 18% and 39% of our cash used in operations for the years ended December 31, 2017 and 2016, respectively, was from a foreign subsidiary. Additionally, in the normal course of business, we are exposed to fluctuations in interest rates as we seek debt financing and invest excess cash. We are also exposed to foreign currency exchange rate fluctuation risk related to our transactions denominated in foreign currencies. We do not currently employ specific strategies, such as the use of derivative instruments or hedging, to manage these exposures. Our currency exposures vary, but are primarily concentrated in the Euro, pound sterling, and Swiss Franc, in large part due to our wholly-owned subsidiaries, 4-AB, a company with operations in Switzerland, and Agenus UK Limited, a company with operations in the United Kingdom. During the year ended December 31, 2017, there has been no material change with respect to our approach toward those exposures.

We had cash, cash equivalents and short-term investments at December 31, 2017 of \$60.2 million, which are exposed to the impact of interest and foreign currency exchange rate changes, and our interest income fluctuates as interest rates change. Due to the short-term nature of our investments in money market funds, our carrying value approximates the fair value of these investments at December 31, 2017, however, we are subject to investment risk.

We invest our cash and cash equivalents in accordance with our investment policy. The primary objectives of our investment policy are to preserve principal, maintain proper liquidity to meet operating needs, and maximize yields. We review our investment policy annually and amend it as deemed necessary. Currently, the investment policy prohibits investing in any structured investment vehicles and asset-backed commercial paper. Although our investments are subject to credit risk, our investment policy specifies credit quality standards for our investments and limits the amount of credit exposure from any single issue, issuer, or type of investment. We do not invest in derivative financial instruments. Accordingly, we do not believe that there is currently any material market risk exposure with respect to derivatives or other financial instruments that would require disclosure under this item.

Item 8. Financial Statements and Supplementary Data

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Report of Independent Registered Public Accounting Firm

To the Stockholders and Board of Directors
Agenus Inc.:

Opinion on the Consolidated Financial Statements

We have audited the accompanying consolidated balance sheets of Agenus Inc. and subsidiaries (the Company) as of December 31, 2017 and 2016, the related consolidated statements of operations and comprehensive loss, stockholders' (deficit) equity, and cash flows for each of the years in the three year period ended December 31, 2017, and the related notes (collectively, the consolidated financial statements). In our opinion, the consolidated financial statements present fairly, in all material respects, the financial position of the Company as of December 31, 2017 and 2016, and the results of its operations and its cash flows for each of the years in the three year period ended December 31, 2017, in conformity with U.S. generally accepted accounting principles.

We also have audited, in accordance with the standards of the Public Company Accounting Oversight Board (United States) (PCAOB), the Company's internal control over financial reporting as of December 31, 2017, based on criteria established in Internal Control – Integrated Framework (2013) issued by the Committee of Sponsoring Organizations of the Treadway Commission, and our report dated March 16, 2018 expressed an unqualified opinion on the effectiveness of the Company's internal control over financial reporting.

Going Concern

The accompanying consolidated financial statements have been prepared assuming that the Company will continue as a going concern. As discussed in Note 1 to the consolidated financial statements, the Company has suffered recurring losses from operations and has a net capital deficiency that raise substantial doubt about its ability to continue as a going concern. Management's plans in regard to these matters are also described in Note 1. The consolidated financial statements do not include any adjustments that might result from the outcome of this uncertainty.

Basis for Opinion

These consolidated financial statements are the responsibility of the Company's management. Our responsibility is to express an opinion on these consolidated financial statements based on our audits. We are a public accounting firm registered with the PCAOB and are required to be independent with respect to the Company in accordance with the U.S. federal securities laws and the applicable rules and regulations of the Securities and Exchange Commission and the PCAOB.

We conducted our audits in accordance with the standards of the PCAOB. Those standards require that we plan and perform the audit to obtain reasonable assurance about whether the consolidated financial statements are free of material misstatement, whether due to error or fraud. Our audits included performing procedures to assess the risks of material misstatement of the consolidated financial statements, whether due to error or fraud, and performing procedures that respond to those risks. Such procedures included examining, on a test basis, evidence regarding the amounts and disclosures in the consolidated financial statements. Our audits also included evaluating the accounting principles used and significant estimates made by management, as well as evaluating the overall presentation of the consolidated financial statements. We believe that our audits provide a reasonable basis for our opinion.

/s/ KPMG LLP

We have served as the Company's auditor since 1997.

Boston, Massachusetts

March 16, 2018

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AGENUS INC. AND SUBSIDIARIES

CONSOLIDATED BALANCE SHEETS

	December 31, 2017	December 31, 2016
ASSETS		
Cash and cash equivalents	\$60,186,617	\$71,448,016
Short-term investments	-	4,988,751
Inventories	79,491	88,200
Accounts Receivable	1,134,493	11,352,022
Prepaid expenses	11,070,960	2,596,675
Other current assets	1,081,993	838,538
Total current assets	73,553,554	91,312,202
Property, plant and equipment, net of accumulated amortization and depreciation of \$34,029,085 and \$31,243,967 at December 31, 2017 and 2016, respectively	26,178,622	25,633,985
Goodwill	23,048,804	22,392,411
Acquired intangible assets, net of accumulated amortization of \$5,461,834 and \$3,193,092 at December 31, 2017 and 2016, respectively	14,406,650	16,364,726
Other long-term assets	1,214,394	1,282,662
Total assets	\$138,402,024	\$156,985,986
LIABILITIES AND STOCKHOLDERS' DEFICIT		
Current portion, long-term debt	\$20,639,735	\$146,061
Current portion, deferred revenue	4,484,882	2,610,719
Accounts payable	8,086,992	5,428,452
Accrued liabilities	21,569,449	27,874,703
Other current liabilities	1,657,063	4,791,265
Total current liabilities	56,438,121	40,851,200
Long-term debt	142,385,024	130,542,424
Deferred revenue	7,748,284	12,344,782
Contingent purchase price consideration	4,373,000	7,561,000
Other long-term liabilities	3,273,387	4,812,846
Commitments and contingencies (Notes 15 and 18)		
STOCKHOLDERS' DEFICIT		
Preferred stock, par value \$0.01 per share; 5,000,000 shares authorized: Series A-1 convertible preferred stock; 31,620 shares designated, issued, and outstanding at December 31, 2017 and 2016; liquidation value of \$32,625,220, and \$32,419,678 at December 31, 2017, and 2016, respectively	316	316
Common stock, par value \$0.01 per share; 240,000,000 shares authorized; 101,706,117 shares and 87,794,933 shares issued at December 31, 2017 and 2016, respectively	1,017,061	877,949
Additional paid-in capital	951,811,958	866,854,348
Accumulated other comprehensive loss	(2,169,354)	(1,529,559)

Accumulated deficit	(1,026,475,773)	(905,329,320)
Total stockholders' deficit	(75,815,792)	(39,126,266)
Total liabilities and stockholders' deficit	\$ 138,402,024	\$ 156,985,986

See accompanying notes to consolidated financial statements.

AGENUS INC. AND SUBSIDIARIES

CONSOLIDATED STATEMENTS OF OPERATIONS AND COMPREHENSIVE LOSS

For the Years Ended December 31, 2017, 2016, and 2015

	2017	2016	2015
Revenue:			
Grant revenue	\$ 168,051	\$ 32,404	\$ 24,118
Service revenue	—	147,456	—
Research and development	42,709,035	22,393,443	24,792,907
Total revenues	42,877,086	22,573,303	24,817,025
Operating expenses:			
Research and development	(116,125,299)	(94,971,379)	(70,444,259)
General and administrative	(33,741,183)	(33,125,690)	(28,370,001)
Contingent purchase price consideration fair value adjustment	3,188,000	(1,953,000)	(6,703,700)
Operating loss	(103,801,396)	(107,476,766)	(80,700,936)
Other income (expense):			
Non-operating income (expense)	1,977,398	(2,202,336)	(5,968,170)
Interest expense, net	(18,868,494)	(17,316,073)	(6,599,083)
Loss before taxes	(120,692,492)	(126,995,175)	(93,268,188)
Income tax benefit	—	—	5,387,067
Net loss	(120,692,492)	(126,995,175)	(87,881,121)
Dividends on Series A-1 convertible preferred stock	(205,541)	(204,246)	(202,960)
Net loss attributable to common stockholders	\$(120,898,033)	\$(127,199,421)	\$(88,084,081)
Per common share data:			
Basic and diluted net loss attributable to common stockholders	\$(1.23)	\$(1.46)	\$(1.13)
Weighted average number of common shares outstanding:			
Basic and diluted	98,415,414	87,070,189	78,212,094
Other comprehensive (loss) income:			
Foreign currency translation (loss) gain	\$(615,213)	\$ 677,536	\$ 164,150
Unrealized loss on investments	—	—	(1,690)
Pension liability	(24,582)	(153,952)	(245,183)
Other comprehensive (loss) income	(639,795)	523,584	(82,723)
Comprehensive loss	\$(121,537,828)	\$(126,675,837)	\$(88,166,804)

See accompanying notes to consolidated financial statements.

AGENUS INC. AND SUBSIDIARIES

CONSOLIDATED STATEMENTS OF STOCKHOLDERS' (DEFICIT) EQUITY

For the Years Ended December 31, 2017, 2016, and 2015

	Series A-1 Convertible Preferred Stock		Common Stock			Treasury Stock			Accumulated Comprehensive Deficit	Total
	Number of Shares	Par Value	Number of Shares	Par Value	Additional Paid-In Capital	Number of Shares	Other	Accumulated		
Balance at December 31, 2014	31,620	\$316	62,720,065	\$627,201	\$715,667,633	-	\$-	\$(1,970,420)	\$(691,306,343)	\$23,018,387
Net loss									(87,881,121)	(87,881,121)
Other comprehensive loss	—	—	—	—	—	—	—	(82,723)	—	(82,723)
Shares sold in underwritten public offering	—	—	12,650,000	126,500	74,543,480	—	—	—	—	74,669,980
Share-based compensation	—	—	—	—	8,098,650	—	—	—	—	8,098,650
Reclassification of liability classified option grants	—	—	—	—	(495,742)	—	—	—	—	(495,742)
Vesting of nonvested shares	—	—	35,332	353	(353)	—	—	—	—	—
Issuance of stock for acquisition of SECANT yeast display technology	—	—	574,140	5,741	2,994,259	—	—	—	—	3,000,000
Shares sold under Stock Purchase	—	—	7,763,968	77,640	34,922,361	—	—	—	—	35,000,001

Agreement										
Issuance of shares related to milestone										
achievement	—	—	80,493	805	343,736	—	—	—	—	344,541
Issuance of warrants										
	—	—	—	—	3,038,438					3,038,438
Issuance of stock in connection with										
XOMA antibody manufacturing										
facility acquisition										
	—	—	109,211	1,092	498,908	—	—	—	—	500,000
Issuance of stock in connection with										
PhosImmune acquisition										
	—	—	1,631,521	16,315	7,383,685	—	—	—	—	7,400,000
Issuance of stock for settlement of										
contingent royalty obligation										
	—	—	300,000	3,000	2,139,000	—	—	—	—	2,142,000
Exercise of stock options and employee share purchases										
	—	—	525,967	5,260	1,969,879	—	—	—	—	1,975,139
Balance at December 31, 2015										
	31,620	\$316	86,390,697	\$863,907	\$851,103,934	—	\$—	\$(2,053,143)	\$(779,187,464)	\$70,727,550

See accompanying notes to consolidated financial statements.

AGENUS INC. AND SUBSIDIARIES

CONSOLIDATED STATEMENTS OF STOCKHOLDERS' (DEFICIT) EQUITY

(Continued)

For the Years Ended December 31, 2017, 2016, and 2015

	Series A-1 Convertible Preferred Stock		Common Stock		Treasury Stock		Accumulated		Total
	Number of Shares	Par Value	Number of Shares	Par Value	Number of Shares	Amount	Other Comprehensive Loss	Accumulated Deficit	
Additional Paid-In Capital	—	—	—	—	—	—	—	—	(126,995,175)
Comprehensive Income	—	—	—	—	—	—	523,584	—	523,584
Share-based Compensation	—	—	—	—	13,323,616	—	—	—	13,323,616
Classification Change Liability	—	—	—	—	(318,677)	—	—	—	(318,677)
Restricted Stock	—	—	570,037	5,701	(5,701)	(185,117)	(768,236)	—	(768,236)
Shares sold at Market	—	—	496,520	4,965	2,162,105	—	—	—	2,167,070
Price of New shares	—	—	23,110	231	161,332	—	—	—	161,563
Conversion of Preferred shares	—	—	(188,184)	(1,882)	(1,632,554)	188,184	781,117	853,319	-
Price of New shares	—	—	157,513	1,575	885,223	—	—	—	886,798
Exercise of Options	—	—	345,240	3,452	1,175,070	(3,067)	(12,881)	—	1,165,644

ce at ber 31,	31,620	\$316	87,794,933	\$877,949	\$866,854,348	—	\$—	\$(1,529,559)	\$(905,329,320)	\$(39,126,
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See accompanying notes to consolidated financial statements.

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AGENUS INC. AND SUBSIDIARIES

CONSOLIDATED STATEMENTS OF STOCKHOLDERS' (DEFICIT) EQUITY

(Continued)

For the Years Ended December 31, 2017, 2016, and 2015

	Series A-1 Convertible Preferred Stock	Common Stock		Additional Paid-In Capital	Treasury Stock Number of Shares	Amount	Accumulated Other Comprehensive Income (Loss)	Accumulated Deficit	Total
	Number of Shares	Number of Shares	Par Value	Capital	Number of Shares	Amount	Income (Loss)	Deficit	Total
								(120,692,492)	(120,692,492)
Comprehensive	—	—	—	—	—	—	(639,795)	—	(639,795)
of	—	—	—	1,210,916	—	—	—	(1,292,230)	(81,313)
ng	—	—	—	10,000,000	—	—	—	—	60,000
old	—	—	—	10,924,122	—	—	—	—	10,924,122
ock	—	—	—	2,015,974	—	—	—	—	2,015,974
ent	—	1,097,243	10,972	(10,972)	(155,523)	(527,223)	—	—	(527,223)
used	—	1,315,288	13,153	5,546,553	—	—	—	—	5,559,706
ation	—	—	—	731,498	—	—	—	—	731,498
ification	—	(155,523)	(1,555)	(1,363,937)	155,523	527,223	—	838,269	—
ty	—	373,351	3,734	1,482,203	—	—	—	—	1,485,937
ed	—	—	—	—	—	—	—	—	—
trants	—	—	—	—	—	—	—	—	—
of	—	—	—	—	—	—	—	—	—
ed	—	—	—	—	—	—	—	—	—
old at	—	—	—	—	—	—	—	—	—
et	—	—	—	—	—	—	—	—	—
ment to	—	—	—	—	—	—	—	—	—
trants	—	—	—	—	—	—	—	—	—
ent of	—	—	—	—	—	—	—	—	—
shares	—	—	—	—	—	—	—	—	—

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T yeast										
gy	—	—	999,317	9,993	3,537,582	—	—	—	—	3,547,882
of										
tions										
loyee										
urchases	—	—	281,508	2,815	983,671	—	—	—	—	986,486
at										
er 31,	31,620	\$316	101,706,117	\$1,017,061	\$951,811,958	—	\$—	\$(2,169,354)	\$(1,026,475,773)	\$(75,811,958)

See accompanying notes to consolidated financial statements.

AGENUS INC. AND SUBSIDIARIES

CONSOLIDATED STATEMENTS OF CASH FLOWS

For the Years Ended December 31, 2017, 2016, and 2015

	2017	2016	2015
Cash flows from operating activities:			
Net loss	\$(120,692,492)	\$(126,995,175)	\$(87,881,121)
Adjustments to reconcile net loss to net cash used in operating activities:			
Depreciation and amortization	6,006,299	4,947,787	1,957,591
Share-based compensation	12,428,655	13,188,364	7,438,308
Non-cash interest expense	18,242,299	16,530,437	5,626,918
Loss on disposal of assets	23,558	14,733	—
Change in fair value of contingent obligations	(3,188,000)	1,953,000	13,567,000
Gain on issuance of milestone payment in stock	(566,488)	—	—
In-process research and development purchase	—	—	12,245,231
Loss on extinguishment of debt	—	—	154,117
Bargain purchase	—	—	(1,522,377)
Deferred tax benefit	—	—	(5,387,067)
Changes in operating assets and liabilities:			
Accounts receivable	10,217,529	(1,549,798)	(9,331,622)
Inventories	8,709	-	7,500
Prepaid expenses	(8,453,082)	(650,824)	(703,424)
Accounts payable	1,647,430	419,708	2,668,064
Deferred revenue	(2,722,020)	(3,939,619)	15,957,820
Accrued liabilities and other current liabilities	(4,848,372)	18,275,940	9,565,639
Other operating assets and liabilities	(2,329,168)	(2,155,364)	(11,538,019)
Net cash used in operating activities	(94,225,143)	(79,960,811)	(47,175,441)
Cash flows from investing activities:			
Cash paid for acquisitions	—	—	(7,182,069)
Proceeds from sale of plant and equipment	120,000	—	—
Purchases of plant and equipment	(3,120,357)	(12,519,738)	(3,591,335)
Purchases of available-for-sale securities	(14,936,047)	(54,884,101)	(34,993,100)
Proceeds from sale of available-for-sale securities	20,000,000	85,000,000	14,534,486
Net cash provided by (used in) investing activities	2,063,596	17,596,161	(31,232,018)
Cash flows from financing activities:			
Net proceeds from sale of equity	65,559,706	2,167,070	109,669,980
Proceeds from employee stock purchases and option exercises	986,486	1,183,598	1,975,139
Purchase of treasury shares to satisfy tax withholdings	(527,223)	(667,050)	—
Proceeds from issuance of long-term debt	15,000,000	—	109,000,000
Debt issuance costs	(150,000)	—	(1,774,323)
Payments of debt	—	—	(1,111,111)
Payment of contingent purchase price consideration	—	—	(8,180,000)
Payment under a purchase agreement for in-process research and development	—	(5,000,000)	—
Payment of contingent royalty obligation	—	—	(20,000,000)
Payment of capital lease obligation	(330,744)	(144,658)	—

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Net cash provided by (used in) financing activities	80,538,225	(2,461,040)	189,579,685
Effect of exchange rate changes on cash	361,923	(429,167)	(183,873)
Net (decrease) increase in cash and cash equivalents	(11,261,399)	(65,254,857)	110,988,354
Cash and cash equivalents, beginning of period	71,448,016	136,702,873	25,714,519
Cash and cash equivalents, end of period	\$60,186,617	\$71,448,016	\$136,702,873
Supplemental cash flow information:			
Cash paid for interest	\$1,120,000	\$1,120,000	\$1,053,447
Supplemental disclosures - non-cash activities:			
Purchases of plant and equipment in accounts payable and			
accrued liabilities	\$968,400	\$695,466	\$105,245

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Issuance of common stock, \$0.01 par value, issued in connection with the settlement of milestone obligation	1,485,937	886,798	—
Issuance of common stock, \$0.01 par value, issued to directors as compensation	—	161,332	—
Issuance of common stock, \$0.01 par value, in connection with the acquisition of the SECANT yeast display technology	3,547,575	—	3,000,000
Issuance of common stock, \$0.01 par value, in connection with acquisition of PhosImmune	—	—	—