

AGENUS INC
Form 424B5
May 11, 2018
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As Filed Pursuant to 424(b)(5)
Registration No. 333-221008

PROSPECTUS SUPPLEMENT

(To Prospectus Dated October 18, 2017)

Agenus Inc.

Up to 20,000,000 Shares

Common Stock

This prospectus supplement relates to the issuance and sale of up to 20,000,000 shares of our common stock from time to time through our sales agent, B. Riley FBR, Inc., or BRFBR. We have entered into an At Market Issuance Sales Agreement (Sales Agreement) with BRFBR relating to shares of our common stock, \$0.01 par value per share offered by this prospectus supplement and the accompanying prospectus. In accordance with the terms of the Sales Agreement, we may offer and sell shares up to 20,000,000 shares of our common stock from time to time through BRFBR.

Our common stock trades on the Nasdaq Capital Market (Nasdaq) under the symbol AGEN. The last reported sale price of our Common Stock on Nasdaq on May 9, 2018 was \$3.41 per share.

Sales of shares of our common stock under this prospectus supplement and the accompanying prospectus, if any, may be made by any method deemed to be an at the market offering as defined in Rule 415(a)(4) under the Securities Act of 1933, as amended (the Securities Act). The sales agent has agreed to use its commercially reasonable efforts consistent with its normal trading and sales practices to sell the shares, on mutually agreed terms between the sales agent and us. There is no arrangement for funds to be received in any escrow, trust or similar arrangement.

The compensation payable to the sales agent for sales of common stock sold pursuant to the Sales Agreement will be an aggregate of 3.0% of the gross proceeds of the sales price of common stock sold.

In connection with the sale of common stock on our behalf, BRFBR will be deemed to be an underwriter within the meaning of the Securities Act, and the compensation of BRFBR will be deemed to be underwriting commissions or discounts.

You should read carefully and consider the Risk Factors referenced on page S-6 of this prospectus supplement and the risk factors described in other documents incorporated by reference herein.

Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities or determined if this prospectus supplement is truthful or complete. Any representation to the contrary is a criminal offense.

B. RILEY FBR

The date of this Prospectus Supplement is May 11, 2018.

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ABOUT THIS PROSPECTUS SUPPLEMENT

This document contains two parts. The first part is the prospectus supplement, including the documents incorporated by reference, which describes the specific terms of this offering. The second part, the accompanying prospectus, including the documents incorporated by reference, provides more general information. Generally, when we refer to this prospectus, we are referring to both parts of this document combined. Before you invest, you should carefully read this prospectus supplement, the accompanying prospectus and all information incorporated by reference herein and therein, as well as the additional information described under the sections titled **Where You Can Find More Information** and **Incorporation of Certain Information by Reference**. These documents contain information you should consider when making your investment decision. This prospectus supplement may add, update or change information contained in the accompanying prospectus. To the extent that any statement that we make in this prospectus supplement is inconsistent with statements made in the accompanying prospectus, the statements made in this prospectus supplement or any documents incorporated by reference will be deemed to modify or supersede those made in the accompanying prospectus and such documents incorporated by reference.

If information in this prospectus supplement is inconsistent with the accompanying prospectus, you should rely on this prospectus supplement. This prospectus supplement, the accompanying prospectus and the documents incorporated into each by reference include important information about us, the shares being offered and other information you should know before investing in our common stock.

You should rely on this prospectus supplement, the accompanying prospectus and the information incorporated or deemed to be incorporated by reference in this prospectus supplement and the accompanying prospectus. We have not, and BRFB has not, authorized anyone to provide you with information that is in addition to or different from that contained or incorporated by reference in this prospectus supplement and the accompanying prospectus. We are not, and BRFB is not, offering to sell these securities in any jurisdiction where the offer or sale is not permitted. You should not assume that the information contained or incorporated by reference in this prospectus supplement or the accompanying prospectus is accurate as of any date other than as of the date of this prospectus supplement or the accompanying prospectus, as the case may be, or in the case of the documents incorporated by reference, the date of such documents regardless of the time of delivery of this prospectus supplement and the accompanying prospectus or any sale of our common stock. Our business, financial condition, liquidity, results of operations and prospects may have changed since those dates.

All references in this prospectus supplement or the accompanying prospectus to **Agenus**, **Antigenics**, the **Company**, **we**, **us** or **our** mean **Agenus Inc.** and its subsidiaries, unless we state otherwise or the context otherwise requires.

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CAUTIONARY NOTE ABOUT FORWARD-LOOKING STATEMENTS

This prospectus supplement, the accompanying prospectus, the documents incorporated by reference herein and therein and other written and oral statements we make from time to time contain certain forward-looking statements within the meaning of Section 27A of the Securities Act and Section 21E of the Securities Exchange Act of 1934, as amended (the Exchange Act). 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, future and other words and terms of similar meaning and expression in connection with any discussion of future operating or financial performance. You can also identify forward-looking statements by the fact that they do not relate strictly to historical or current facts. Such forward-looking statements are based on current expectations and involve inherent risks and uncertainties, including factors that could delay, divert or change any of them, and could cause actual outcomes to differ materially from current expectations. These statements are likely to 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, uncertainty regarding our future operating results and our profitability, anticipated sources of funds as well as our plans, objectives, expectations, and intentions. will,

We have included more detailed descriptions of these risks and uncertainties and other risks and uncertainties applicable to our business that we believe could cause actual results to differ materially from any forward-looking statement in the Risk Factors sections of this prospectus supplement, the accompanying prospectus, and the documents incorporated by reference herein including, but not limited to, the risk factors incorporated by reference from our filings with the SEC. We encourage you to read those descriptions carefully. 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. We caution investors not to place significant reliance on forward-looking statements; such statements need to be evaluated in light of all the information contained and incorporated by reference in this prospectus supplement and the accompanying prospectus. Furthermore, the statements speak only as of the date of each document, and we undertake no obligation to update or revise these statements.

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PROSPECTUS SUPPLEMENT SUMMARY

The following is a summary of selected information contained elsewhere or incorporated by reference. It does not contain all of the information that you should consider before buying our securities. You should read this prospectus supplement and the accompanying prospectus in their entirety, including the information incorporated by reference herein and therein.

The Company

Our Business

AgenuS is 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, cancer vaccine platforms and cell therapy (through our subsidiary, AgenTus Therapeutics). 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 checkpoint modulating (CPM) antibody candidates;

our antibody candidate programs, including our CPM programs;

our vaccine programs, including Prophage , AutoSynVax and PhosPhoSynVax ;

our saponin-based vaccine adjuvants, principally our QS-21 Stimulon[®] adjuvant, or QS-21 Stimulon; and

our cell therapy subsidiary, AgenTus Therapeutics, which is designed to drive the discovery of future adoptive cell therapy, or living drugs (CAR-T and TCR) programs.

We also have our own GMP manufacturing facility with the capacity to support early phase clinical programs.

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. As such, we have recently shifted our strategy for first approval from lung cancer to cervical cancer based on increasing competition and recent data that would have hindered our ability to pursue accelerated pathways for approval. We are currently advancing our own combination of CTLA-4 and PD-1 antibodies

in second line cervical cancer.

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. 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.

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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 NPA, 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.

We have incurred significant losses since our inception. As of March 31, 2018, we had an accumulated deficit of \$1.1 billion. Since our inception, we have financed our operations through the sale of equity and notes, corporate partnerships, and interest income. Based on our current plans, including additional funding we anticipate from multiple sources, including out-licensing and/or partnering opportunities, we believe that our cash resources of \$52.3 million as of March 31, 2018 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. However, in spite of these plans and our ability to control our cash burn, in accordance with the requirements of ASU 2014-15, we are required to disclose the existence of a substantial doubt regarding our ability to continue as a going concern for twelve months from the date on which the financial statements contained in our Quarterly Report on Form 10-Q for the quarterly period ended March 31, 2018, filed with the SEC on May 10, 2018, were issued.

You can find more information about us in our filings with the Securities and Exchange Commission referenced in the sections in this document titled Where You Can Find More Information and Incorporation of Certain Documents by Reference beginning on page S-15.

Corporate Information

Our principal executive office is located at 3 Forbes Road, Lexington, MA, 02421, and our telephone number is (781) 674-4400. Our website address is www.agenusbio.com. Information contained on our website is not a part of this prospectus supplement.

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THE OFFERING

*The following summary contains basic information about our common stock and the offering and is not intended to be complete. It does not contain all the information that may be important to you. For a more complete understanding of our common stock, you should read the section entitled *Description of Common Stock*.*

| | |
|--|--|
| Common stock offered by us | Up to 20,000,000 shares. |
| Common stock outstanding after this offering | Up to 123,999,081 shares. The actual number of shares issued will vary depending on how many shares we choose to sell under this offering. |
| Plan of distribution | At the market offering that may be made from time to time for our common stock in the United States through our agent, BRFBR. See the section entitled <i>Plan of Distribution</i> below. |
| Use of proceeds | We intend to use the net proceeds from this offering for general corporate purposes which may include working capital, capital expenditures, repayment and refinancing of debt, research and development expenditures, clinical trial expenditures, acquisitions of additional companies or technologies and investments. We may temporarily invest the net proceeds in investment-grade, interest-bearing securities until they are used for their stated purpose. We have not determined the amount of net proceeds to be used specifically for such purposes. As a result, management will retain broad discretion over the allocation of net proceeds. |
| Risk factors | Your investment in our common stock involves substantial risks. You should consider the <i>Risk Factors</i> included and incorporated by reference in this prospectus supplement, including the risk factors incorporated by reference from our filings with the SEC. |
| Nasdaq symbol | AGEN |
| The number of shares of our common stock to be outstanding after this offering is based on approximately 103,999,081 shares of our common stock outstanding as of March 31, 2018 and does not include: | |

2,900,000 shares issuable upon the exercise of outstanding or issuable warrants with a weighted-average exercise price of \$4.52 per share;

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14,129,736 shares issuable upon the exercise of outstanding stock options with a weighted-average exercise price of \$4.23 per share;

1,280,750 nonvested shares;

333,333 shares issuable upon the conversion of our outstanding shares of series A-1 convertible preferred stock;

166,666 shares issuable under our Employee Stock Purchase Plan;

252,919 shares issuable under our Directors' Deferred Compensation Plan; and

1,097,726 shares reserved for future issuance under our Amended and Restated 2009 Equity Incentive Plan, or the 2009 Plan, or an additional 9,000,000 shares reserved for future issuance under our 2009 Plan for which our board of directors approved an increase in April 2018, subject to approval by our stockholders at our 2018 annual meeting of stockholders.

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RISK FACTORS

Before purchasing any of the securities you should carefully consider the risk factors relating to Agenesis described below and incorporated by reference in this prospectus supplement and the accompanying prospectus from our Quarterly Report on Form 10-Q for the quarter ended March 31, 2018, as well as the risks, uncertainties and additional information set forth in our reports on Forms 10-K, 10-Q and 8-K filed with the Securities and Exchange Commission, or SEC, and in the other documents incorporated by reference in this prospectus supplement and the accompanying prospectus. For a description of these reports and documents, and information about where you can find them, see [Where You Can Find More Information](#) and [Incorporation of Certain Documents By Reference](#). Additional risks not presently known or that we presently consider to be immaterial could subsequently materially and adversely affect our financial condition, results of operations, business and prospects.

Additional Risks Relating to this Offering and Our Common Stock

We may allocate the net proceeds from this offering in ways that you and other stockholders may not approve.

We currently intend to use the net proceeds of this offering, if any, for general corporate purposes which may include working capital, capital expenditures, repayment and refinancing of debt, research and development expenditures, clinical trial expenditures, acquisitions of additional companies or technologies and investments. This expected use of the net proceeds from this offering represents our intentions based upon our current plans and business conditions. The amounts and timing of our actual expenditures may vary significantly depending on numerous factors, including the progress of our development efforts, the status of and results from clinical trials, as well as any collaborations that we may enter into with third parties for our product candidates, and any unforeseen cash needs. Because of the number and variability of factors that will determine our use of the proceeds from this offering, their ultimate use may vary substantially from their currently intended use. As a result, our management will retain broad discretion over the allocation of the net proceeds from this offering and could spend the proceeds in ways that do not necessarily improve our operating results or enhance the value of our common stock. See [Use of Proceeds](#).

If you purchase our common stock in this offering, you will incur immediate and substantial dilution in the book value of your shares.

The public offering price of our common stock is substantially higher than the net tangible book value per share of our common stock. Therefore, if you purchase shares of our common stock in this offering, you will pay a price per share that substantially exceeds our net tangible book value per share after this offering. Assuming the sale of all 20,000,000 shares of our common stock in this offering, based on the assumed public offering price of \$3.68 per share, which was the last reported sale price of our common stock on Nasdaq on May 4, 2018, and net tangible book value per share of our common stock of \$(1.46) as of March 31, 2018, if you purchase shares in this offering, you will suffer immediate and substantial dilution of \$4.32 per share in the net tangible book value of common stock purchased. To the extent shares are issued under outstanding options or warrants, you will incur further dilution. See [Dilution](#) for a more detailed description of the dilution to new investors in the offering.

You may experience future dilution as a result of future equity offerings.

In order to raise additional capital, we may at any time, including during the pendency of this offering, offer additional shares of our common stock or other securities convertible into or exchangeable for our common stock at prices that may not be the same as the price per share in this offering. We may sell shares or other securities in any other offering at a price per share that is less than the price per share paid by investors in this offering, and investors purchasing shares or other securities in the future could have rights superior to existing stockholders. The price per share at which

we sell additional shares of our common stock, or securities convertible or exchangeable into common stock, in future transactions may be higher or lower than the price per share paid by investors in this offering.

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USE OF PROCEEDS

We intend to use the net proceeds from this offering for general corporate purposes which may include working capital, capital expenditures, repayment and refinancing of debt, research and development expenditures, clinical trial expenditures, acquisitions of additional companies or technologies and investments. We may temporarily invest the net proceeds in investment-grade, interest-bearing securities until they are used for their stated purpose. We have not determined the amount of net proceeds to be used specifically for such purposes. As a result, management will retain broad discretion over the allocation of net proceeds.

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If you invest in our common stock, your ownership interest will be diluted to the extent of the difference between the public offering price per share and the as-adjusted net tangible book value per share after this offering. We calculate net tangible book value per share by dividing the net tangible book value, which is tangible assets less total liabilities, by the number of outstanding shares of our common stock.

Our net tangible book value as of March 31, 2018 was \$(150.6) million, or \$(1.46) per share of common stock. After giving effect to the sale of our common stock in the aggregate amount of \$71.4 million at an assumed offering price of \$3.68 per share, the last reported sale price of our common stock on Nasdaq as of May 4, 2018, and after deducting estimated offering commissions and expenses payable by us, our net tangible book value as of March 31, 2018 would have been \$(79.2) million, or \$(0.64) per share of common stock. This represents an immediate increase in the net tangible book value of \$0.82 per share to our existing stockholders and an immediate and substantial dilution in net tangible book value of \$4.32 per share to new investors. The following table illustrates this per share dilution:

| | |
|--|-----------|
| Assumed offering price per share | \$ 3.68 |
| Net tangible book value per share as of March 31, 2018 | \$ (1.46) |
| Increase in net tangible book value per share after this offering | \$ 0.82 |
| As-Adjusted net tangible book value per share as of March 31, 2018, after giving effect to this offering | \$ (0.64) |
| Dilution per share to new investors in this offering | \$ (4.32) |

The above discussion and table are based on approximately 103,999,081 shares of our common stock outstanding as of March 31, 2018 and does not include:

2,900,000 shares issuable upon the exercise of outstanding or issuable warrants with a weighted-average exercise price of \$4.52 per share;

14,129,736 shares issuable upon the exercise of outstanding stock options with a weighted-average exercise price of \$4.23 per share;

1,280,750 nonvested shares;

333,333 shares issuable upon the conversion of our outstanding shares of series A-1 convertible preferred stock;

166,666 shares issuable under our Employee Stock Purchase Plan;

252,919 shares issuable under our Directors' Deferred Compensation Plan; and

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1,097,726 shares reserved for future issuance under our Amended and Restated 2009 Equity Incentive Plan, or the 2009 Plan, or an additional 9,000,000 shares reserved for future issuance under our 2009 Plan for which our board of directors approved an increase in April 2018, subject to approval by our stockholders at our 2018 annual meeting of stockholders.

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DESCRIPTION OF COMMON STOCK

General

Each share of Agenus common stock has the same relative rights and is identical in all respects with each other share of common stock.

Voting Rights

Subject to preferences that may apply to shares of preferred stock outstanding at the time, the holders of outstanding shares of common stock are entitled to receive dividends out of assets legally available for payment of dividends, as the board may from time to time determine. Each stockholder is entitled to one vote for each share of common stock held on all matters submitted to a vote of stockholders. Our certificate of incorporation does not provide for cumulative voting for the election of directors, which means that the holders of a majority of the shares voted can elect all of the directors then standing for election. The common stock is not entitled to preemptive rights and is not subject to conversion or redemption. Each outstanding share of common stock offered by this prospectus supplement will, when issued, be fully paid and nonassessable.

Dividends

Holders of common stock are entitled to share ratably in any dividends declared by our board of directors, subject to any preferential dividend rights of any outstanding preferred stock. Dividends consisting of shares of common stock may be paid to holders of shares of common stock. We have never declared or paid cash dividends on our common stock. We do not intend to pay cash dividends in the foreseeable future.

Preemptive Rights

Holders of common stock do not have any preemptive rights with respect to any shares that may be issued by Agenus in the future. Thus, Agenus may sell shares of its common stock without first offering them to the then holders of common stock.

Liquidation

In the event of any liquidation or dissolution of Agenus, whether voluntary or involuntary, the holders of Agenus common stock would be entitled to receive pro rata, after payment of all debts and liabilities of Agenus, all assets of Agenus available for distribution, subject to the rights of the holders of any preferred stock which may be issued with a priority in liquidation or dissolution over the holders of common stock.

Listing

Our common stock is listed on Nasdaq under the symbol AGEN. On May 9, 2018, the last reported sale price for our common stock on Nasdaq was \$3.41 per share. As of May 9, 2018, we had approximately 394 stockholders of record.

Anti-Takeover Provisions

Statutory Business Combination Provision

Delaware has adopted a business combination statute (Section 203 of the DGCL) that may also have additional anti-takeover effects to provisions in Agenus' restated certificate of incorporation and bylaws. Section 203 of the DGCL, which, subject to certain exceptions, prohibits a Delaware corporation from engaging in any business combination with an interested stockholder for a period of three years following the time that such stockholder became an interested stockholder, unless:

the board of directors of the corporation approves either the business combination or the transaction that resulted in the stockholder becoming an interested stockholder, prior to the time the interested stockholder attained that status;

upon the closing of the transaction that resulted in the stockholder becoming an interested stockholder, the interested stockholder owned at least 85% of the voting stock of the corporation outstanding at the time the transaction commenced, excluding, for purposes of determining the number of shares outstanding, those shares owned by persons who are directors and also officers and by employee stock plans in which employee participants do not have the right to determine confidentially whether shares held subject to the plan will be tendered in a tender or exchange offer; or

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at or subsequent to such time, the business combination is approved by the board of directors and authorized at an annual or special meeting of stockholders, and not by written consent, by the affirmative vote of at least 66 2/3% of the outstanding voting stock that is not owned by the interested stockholder.

With certain exceptions, an interested stockholder is a person or group who or which owns 15% or more of the corporation's outstanding voting stock (including any rights to acquire stock pursuant to an option, warrant, agreement, arrangement or understanding, or upon the exercise of conversion or exchange rights, and stock with respect to which the person has voting rights only), or is an affiliate or associate of the corporation and was the owner of 15% or more of such voting stock at any time within the previous three years.

In general, Section 203 defines a business combination to include:

any merger or consolidation involving the corporation and the interested stockholder;

any sale, transfer, pledge or other disposition of 10% or more of the assets of the corporation involving the interested stockholder;

subject to certain exceptions, any transaction that results in the issuance or transfer by the corporation of any stock of the corporation to the interested stockholder;

any transaction involving the corporation that has the effect of increasing the proportionate share of the stock or any class or series of the corporation beneficially owned by the interested stockholder; or

the receipt by the interested stockholder of the benefit of any loans, advances, guarantees, pledges or other financial benefits provided by or through the corporation.

A Delaware corporation may opt out of this provision with an express provision in its original certificate of incorporation or an express provision in its amended and restated certificate of incorporation or bylaws resulting from a stockholders' amendment approved by at least a majority of the outstanding voting shares. However, Agenus has not opted out of this provision. Section 203 could prohibit or delay mergers or other takeover or change-in-control attempts and, accordingly, may discourage attempts to acquire Agenus.

Size of the Board and Vacancies

Our bylaws provide that the exact number of directors is determined by resolution of the board of directors or by the stockholders at the annual meeting. Our board of directors has the right to fill any vacancies resulting from death, resignation, disqualification or removal, as well as any newly created directorships arising from an increase in the size of the board.

Amendment of Charter Provisions

The affirmative vote of the holders of at least a majority of the voting power of all then outstanding shares of our voting stock, voting together as a single class, is required to, among other things, amend, alter, change or repeal certain provisions of our restated certificate of incorporation. Our bylaws may only be amended (or new bylaws

adopted) by the board of directors or the affirmative vote of the holders of at least a majority of the voting power of all then outstanding shares of our voting stock represented in person or by proxy at the meeting at which the amendment is voted on.

Undesignated Preferred Stock

We could issue preferred stock that could have other rights, including economic rights senior to our common stock, so that the issuance of the preferred stock could adversely affect the market value of our common stock. The issuance of the preferred stock may also have the effect of delaying, deferring or preventing a change in control of Agenus without any action by the stockholders. The effects of issuing preferred stock could include one or more of the following:

restricting dividends on the common stock;

diluting the voting power of the common stock;

impairing the liquidation rights of the common stock; or

discouraging, delaying or preventing changes in control or management of Agenus.

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Transfer Agent

The transfer agent and registrar for Agenus common stock is American Stock Transfer & Trust Company. Its telephone number is (800) 937-5449.

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PLAN OF DISTRIBUTION

We have entered into an At Market Issuance Sales Agreement with BRFBR, under which we may issue and sell an aggregate of up to 20,000,000 shares of our common stock from time to time through BRFBR acting as agent. The sales agreement will be filed with the SEC and will be incorporated by reference into this prospectus supplement. This is a brief summary of the material terms of the sales agreement and does not purport to be a complete statement of its terms and conditions.

Each time that we wish to sell common stock under the sales agreement, we will provide BRFBR with a placement notice describing the number or dollar value of shares to be issued, the time period during which sales are requested to be made, any limitation on the number of shares that may be sold in any one day and any minimum price below which sales may not be made.

Upon receipt of a placement notice from us, and subject to the terms and conditions of the sales agreement, BRFBR has agreed to use its commercially reasonable efforts consistent with its normal trading and sales practices and applicable state and federal laws, rules and regulations, and the rules of Nasdaq to sell such shares up to the amount specified. The settlement between us and BRFBR of each sale will occur on the second trading day following the date on which the sale was made or on some other date that is agreed upon by us and BRFBR in connection with a particular transaction. The obligation of BRFBR under the sales agreement to sell our common stock pursuant to a placement notice is subject to a number of conditions.

We will pay BRFBR a commission of up to 3.0% of the gross proceeds of the sale price per share. We estimate that the total expenses for the offering, excluding compensation and reimbursements payable to BRFBR under the terms of the sales agreement, will be approximately \$50,000.00.

The actual proceeds to us will vary depending on the number of share sold and the prices of such sales. Because there is no minimum offering amount required as a condition to close this offering, the actual total public offering amount, commissions and proceeds to us, if any, are not determinable at this time.

In connection with the sale of our common stock contemplated in this prospectus supplement, BRFBR will be deemed to be an underwriter within the meaning of the Securities Act, and the compensation paid to BRFBR will be deemed to be underwriting commissions or discounts. We have agreed to indemnify BRFBR against certain civil liabilities, including liabilities under the Securities Act.

Sales of our common stock as contemplated in this prospectus supplement will be settled through the facilities of The Depository Trust Company or by such other means as we and BRFBR may agree upon. There is no arrangement for funds to be received in escrow, trust or similar arrangement.

The offering of our common stock pursuant to the sales agreement will terminate on the earlier of (1) the issuance and sale of all of our common stock subject to the sales agreement, or (2) termination of the sales agreement by us or BRFBR.

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LEGAL MATTERS

The validity of the securities that may be offered hereby will be passed upon for us by Ropes & Gray LLP, Boston, Massachusetts. B. Riley FBR, Inc. is being represented in connection with this offering by Duane Morris LLP, New York, New York.

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EXPERTS

The consolidated financial statements of Agenus Inc., as of December 31, 2017 and 2016, and for each of the years in the three-year period ended December 31, 2017, and management's assessment of the effectiveness of internal control over financial reporting as of December 31, 2017 have been incorporated by reference herein and in the registration statement in reliance upon the reports of KPMG LLP, independent registered public accounting firm, incorporated by reference herein, and upon the authority of said firm as experts in accounting and auditing. The audit report of KPMG LLP on the aforementioned consolidated financial statements contains an explanatory paragraph that states that the Company's recurring losses from operations and net capital deficiency raise substantial doubt about the entity's ability to continue as a going concern. The consolidated financial statements do not include any adjustments that might result from the outcome of that uncertainty.

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WHERE YOU CAN FIND MORE INFORMATION

This prospectus supplement and the accompanying prospectus are part of a registration statement we filed with the SEC. This prospectus supplement and the accompanying prospectus do not contain all of the information set forth in the registration statement and the exhibits to the registration statement. For further information with respect to us and the securities we are offering under this prospectus supplement and the accompanying prospectus, we refer you to the registration statement and the exhibits and schedules filed as a part of the registration statement. Neither we nor any agent, underwriter or dealer has authorized any person to provide you with information that is different from that contained in this prospectus supplement and the accompanying prospectus or in any free writing prospectus we may authorize to be delivered or made available to you. We take no responsibility for, and can provide no assurance as to the reliability of, any other information that others may give you. We are not making an offer of these securities in any state where the offer is not permitted. You should not assume that the information in this prospectus supplement and the accompanying prospectus is accurate as of any date other than the date on the front page of such documents, regardless of the time of delivery of this prospectus supplement and the accompanying prospectus or any sale of the securities offered by this prospectus supplement and the accompanying prospectus.

Agenus Inc. is subject to the information requirements of the Exchange Act and files annual, quarterly and special reports, proxy statements and other information with the SEC. You may read and copy any materials we file with the SEC at the Public Reference Room of the SEC at Room 1580, 100 F Street, N.E., Washington, D.C. 20549. You may obtain information on the operation of the Public Reference Room by calling the SEC at 1-800-SEC-0330. In addition, we file many of our documents electronically with the SEC, and you may access those documents over the Internet. The SEC maintains a web site that contains reports, proxy and information statements and other information regarding issuers that file electronically with the SEC. The address of the SEC's website is <http://www.sec.gov>. Documents we have filed with the SEC are also available on our website through the investor link at www.agenusbio.com. Information contained on our web site does not constitute a part of this prospectus supplement and is not incorporated by reference herein.

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INCORPORATION OF CERTAIN DOCUMENTS BY REFERENCE

The SEC allows us to incorporate by reference in this prospectus supplement and the accompanying prospectus the information we file with the SEC. This helps us disclose certain important information to you by referring you to the documents we file. The information we incorporate by reference is an important part of this prospectus supplement and the accompanying prospectus. Because we are incorporating by reference future filings with the SEC, this prospectus supplement and the accompanying prospectus are continually updated and those future filings may modify or supersede some of the information included or incorporated in this prospectus supplement and the accompanying prospectus. This means that you must look at all of the SEC filings that we incorporate by reference to determine if any of the statements in this prospectus supplement, the accompanying prospectus or in any document previously incorporated by reference have been modified or superseded. We incorporate by reference each of the documents listed below.

our Annual Report on Form 10-K for the year ended December 31, 2017 (File No. 000-29089);

our Quarterly Report on Form 10-Q for the quarter ended March 31, 2018 (File No. 000-29089);

our Current Reports on Form 8-K filed on January 8, 2018, January 22, 2018 and May 8, 2018 (except, with respect to each of the foregoing, for portions of such reports which were deemed to be furnished and not filed) (File No. 000-29089);

our Proxy Statement on Schedule 14A filed with the SEC on April 26, 2018 (File No. 000-29089); and

the description of our common stock contained in our registration statement on Form 8-A filed under the Exchange Act on January 24, 2000, including any amendment or reports filed for the purpose of updating such descriptions (File No. 000-29089).

All filings by Agenus pursuant to the Exchange Act subsequent to the date hereof and prior to effectiveness of this registration statement are incorporated in this registration statement and deemed to be a part hereof from the date of filing of such documents or reports. In addition, all documents and reports filed by Agenus subsequent to the date hereof pursuant to Sections 13(a), 13(c), 14 and 15(d) of the Exchange Act prior to the filing of a post-effective amendment which indicates that all securities offered have been sold or which deregisters all securities remaining unsold, shall be deemed to be incorporated by reference in this registration statement and to be a part hereof from the date of filing of such documents or reports. Any statement contained in a document incorporated by reference herein shall be deemed to be modified or superseded to the extent that a statement contained herein or in any other subsequently filed document which also is incorporated by reference herein modifies or supersedes such statement. Statements in this prospectus supplement and the accompanying prospectus concerning any document we filed as an exhibit to the registration statement or that we otherwise filed with the SEC are not intended to be comprehensive and are qualified by reference to these filings. You should review the complete document to evaluate these statements.

We will provide each person to whom this prospectus supplement and the accompanying prospectus is delivered a copy of all of the information that has been incorporated by reference in this prospectus supplement and the accompanying prospectus but not delivered with this prospectus supplement and the accompanying prospectus. You

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may obtain copies of these filings, at no cost, through the Investor section of our website (www.agenusbio.com), and you may request copies of these filings, at no cost, by writing or telephoning us at:

Agenus Inc.

Attention: Secretary

3 Forbes Road

Lexington, MA 02421

Telephone: (781) 674-4400

The information contained on our website is not a part of this prospectus supplement.

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PROSPECTUS

\$250,000,000

Common Stock

Preferred Stock

Warrants

Debt Securities

Units

We may offer to sell to the public, from time to time in one or more offerings for an aggregate initial offering price of up to \$250,000,000:

shares of our common stock;

shares of our preferred stock;

warrants to purchase shares of our common stock, preferred stock and/or debt securities;

debt securities consisting of debentures, notes, or other evidences of indebtedness; or

units consisting of any combination of the foregoing securities.

We may offer and sell any combination of the securities in amounts, at prices and on terms that we will determine at the time of any particular offering, to or through one or more agents, dealers or underwriters, or directly to purchasers, including through subscription rights offerings, on a continuous or delayed basis. If agents, underwriters or dealers are used to sell the securities, we will name them and describe their compensation in a prospectus supplement.

This prospectus describes the general terms of these securities and the general manner in which these securities will be offered. We will provide the specific terms of these securities in supplements to this prospectus. The prospectus supplements will also describe the specific manner in which these securities will be offered and may also supplement, update or amend information contained in this document. You should read this prospectus and any applicable prospectus supplement before you invest.

Our common stock is listed on The NASDAQ Capital Market and trades under the symbol AGEN. On October 17, 2017, the last sale price of our common stock as reported on the NASDAQ Capital Market was \$4.67 per share. The other securities that may be offered are not listed on any securities exchange or included in any automated quotation system.

You should read carefully and consider the Risk Factors referenced on page 3 of this prospectus and the risk factors described in other documents incorporated by reference herein.

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

The date of this prospectus is October 18, 2017.

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ABOUT THIS PROSPECTUS

This prospectus is part of a registration statement that we filed with the Securities and Exchange Commission (the SEC), using a shelf registration process for the delayed offering and sale of securities pursuant to Rule 415 under the Securities Act of 1933, as amended (the Securities Act). Under the shelf process, we may, from time to time, sell any of the securities described in this prospectus in one or more offerings for an aggregate initial offering price of up to \$250,000,000.

This prospectus provides you with a general description of the securities we may offer. Each time we sell securities, we will provide one or more prospectus supplements that will contain specific information about the terms of the offering. The prospectus supplement may also add, update or change information contained in this prospectus. You should read both this prospectus and the accompanying prospectus supplement together with the additional information described under the heading **Where You Can Find More Information** beginning on page 36 of this prospectus.

We have not authorized anyone to provide you with any additional information. This prospectus and any accompanying prospectus supplement do not constitute an offer to sell or the solicitation of an offer to buy any securities other than the securities described in the accompanying prospectus supplement or an offer to sell or the solicitation of an offer to buy such securities in any circumstances in which such offer or solicitation is unlawful. You should assume that the information appearing in this prospectus, any prospectus supplement, the documents incorporated by reference and any related free writing prospectus is accurate only as of their respective dates. Our business, financial condition, results of operations and prospects may have changed materially since those dates.

As used in this prospectus, unless the context otherwise requires, the terms **we**, **us**, **our**, **the Company** and **Agenus** mean, collectively, Agenus Inc. and its subsidiaries and their predecessors.

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CAUTIONARY NOTE ABOUT FORWARD-LOOKING STATEMENTS

This prospectus, any prospectus supplement, and any information incorporated by reference into this prospectus or prospectus supplement may contain certain forward-looking statements within the meaning of Section 27A of the Securities Act of 1933, as amended (the Securities Act) and Section 21E of the Securities Exchange Act of 1934, as amended (the Exchange Act). 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, future and other words and terms of similar meaning and expression in connection with any discussion of future operating or financial performance. You can also identify forward-looking statements by the fact that they do not relate strictly to historical or current facts. Such forward-looking statements are based on current expectations and involve inherent risks and uncertainties, including factors that could delay, divert or change any of them, and could cause actual outcomes to differ materially from current expectations. These statements are likely to 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, uncertainty regarding our future operating results and our profitability, anticipated sources of funds as well as our plans, objectives, expectations, and intentions. will,

We have included more detailed descriptions of these risks and uncertainties and other risks and uncertainties applicable to our business that we believe could cause actual results to differ materially from any forward-looking statement in the Risk Factors sections of this prospectus and the documents incorporated by reference herein including, but not limited to, the risk factors incorporated by reference from our filings with the SEC. We encourage you to read those descriptions carefully. 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. We caution investors not to place significant reliance on forward-looking statements; such statements need to be evaluated in light of all the information contained and incorporated by reference in this prospectus. Furthermore, the statements speak only as of the date of each document, and we undertake no obligation to update or revise these statements, except as required by law.

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

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RISK FACTORS

Before purchasing any of the securities you should carefully consider the risk factors relating to Agenus incorporated by reference in this prospectus from our Quarterly Report on Form 10-Q for the quarter ended June 30, 2017, as well as the risks, uncertainties and additional information set forth in our SEC reports on Forms 10-K, 10-Q and 8-K and in the other documents incorporated by reference in this prospectus. For a description of these reports and documents, and information about where you can find them, see [Where You Can Find More Information](#) and [Incorporation of Certain Documents By Reference](#). Additional risks not presently known or that we presently consider to be immaterial could subsequently materially and adversely affect our financial condition, results of operations, business and prospects.

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THE COMPANY

Agenus is a clinical-stage immuno-oncology (I-O) company focused on the discovery and development of therapies that engage the body's immune system to fight cancer. Our approach to cancer immunotherapy involves a diverse portfolio consisting of antibody-based therapeutics, adjuvants and cancer vaccine platforms. We, in collaboration with our partners, are developing a number of immuno-modulatory antibodies against important nodes of immune regulation. These include antibodies targeting CTLA-4, GITR, OX40, and PD-1 that are in clinical development. Our discovery pipeline includes a number of proprietary checkpoint modulating (CPM) antibodies against innovative targets such as TIGIT and CD-137. We believe that tailored combination therapies are essential to combat some of the most resistant cancers. Accordingly, our immune education strategy focuses on pursuing antibodies as well as vaccine candidates in conjunction with adjuvants.

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 produce quality human antibodies;

our antibody candidate programs, including our CPM programs;

our vaccine programs, including Prophage[™], AutoSynVax[™] and PhosphoSynVax[™]; and

our saponin-based vaccine adjuvants, principally our QS-21 Stimulon[®] adjuvant, or QS-21 Stimulon. We also have our own good manufacturing practices manufacturing facility with the capacity to support early phase clinical programs.

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, 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), all of which are in Phase 1 clinical trials. 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 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 our antibodies targeting GITR and OX40. Concurrent with the execution of the amendment, we and Incyte also entered into a separate 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 will evaluate 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 providing pembrolizumab and the NCI and Brain Tumor Trials Collaborative member sites are recruiting patients and conducting the trial. Earlier this year, we also initiated a Phase 1 clinical trial for our neoantigen vaccine candidate, AutoSynVax (ASV), in patients with advanced cancer. ASV is being used in combination with our QS-21 Stimulon adjuvant.

Our QS-21 Stimulon adjuvant is also partnered with GlaxoSmithKline (GSK) and is a key component in multiple GSK vaccine programs that target prophylactic or therapeutic impact in a variety of infectious diseases and cancer. These programs are in various stages, with the most advanced being GSK s shingles and malaria programs, which GSK first announced positive Phase 3 results for in December 2014 and October 2013, respectively. In 2015, we monetized a portion of the future royalties we are 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 \$78.2 million. In 2016, GSK filed for approval of its shingles vaccine candidate in the United States, European Union and Canada, and in 2017 it filed for approval in Japan. Assuming regulatory approval, the first products containing QS-21 Stimulon are anticipated to be launched by GSK in 2018. We do not incur clinical development costs for products partnered with GSK.

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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.

We have incurred significant losses since our inception. As of June 30, 2017, we had an accumulated deficit of \$954.6 million. Since our inception, we have successfully financed our operations primarily through the sale of equity and convertible and other notes, corporate partnerships, and interest income earned on cash, cash equivalents, and short-term investments balances. Based on our current plans and activities, our cash, cash equivalents and short-term investments balance of \$96.8 million as of June 30, 2017 would only be sufficient to satisfy our liquidity requirements through the first quarter of 2018 without any additional funding before that time, which we anticipate. Regardless of this anticipated funding, in accordance with ASU 2014-15 this is deemed to be a condition which raises substantial doubt regarding our ability to continue as a going concern for at least one year from when the June 30, 2017 financial statements were issued. In order to continue as a going concern, we expect to raise additional funding from currently contemplated transactions before year end. We also 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, if necessary.

You can find more information about us in our filings with the Securities and Exchange Commission referenced in the sections in this document titled *Where You Can Find More Information* and *Incorporation of Certain Documents by Reference* beginning on pages 36 and 37, respectively.

Corporate Information

Our principal executive office is located at 3 Forbes Road, Lexington, MA, 02421, and our telephone number is (781) 674-4400. Our website address is www.agenusbio.com. Information contained on our website is not a part of this prospectus.

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USE OF PROCEEDS

We intend to use the net proceeds from the sale of the securities from offerings under this prospectus for general corporate purposes unless the applicable prospectus supplement states otherwise. General corporate purposes may include working capital, capital expenditures, repayment and refinancing of debt, research and development expenditures, clinical trial expenditures, acquisitions of additional companies or technologies and investments. We may temporarily invest the net proceeds in investment-grade, interest-bearing securities until they are used for their stated purpose. We have not determined the amount of net proceeds to be used specifically for such purposes. As a result, management will retain broad discretion over the allocation of net proceeds.

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| | Six Months | | Years Ended December 31, | | | |
|---|-------------------|--------------|---------------------------------|-------------|-------------|-------------|
| | Ended | | | | | |
| | June 30, | 2016 | 2015 | 2014 | 2013 | 2012 |
| | 2017 | | | | | |
| Ratio of Earnings to Fixed Charges | | | | | | |
| Deficiency of Earnings to Cover Fixed Charges | \$ (48,816) | \$ (126,995) | \$ (93,268) | \$ (42,486) | \$ (30,073) | \$ (11,325) |
| Ratio of Earnings to Combined Fixed Charges and Preferred Stock Dividends | | | | | | |
| Deficiency of Earnings to Cover Combined Fixed Charges and Preferred Stock Dividends | \$ (48,919) | \$ (127,199) | \$ (93,471) | \$ (42,690) | \$ (33,233) | \$ (12,117) |

For purposes of the ratio of earnings to fixed charges and the ration of combined fixed charges and preferred stock dividends to earnings, earnings consist of income before income taxes, interest and the portions of rentals representative of the interest factor. Fixed charges consist of interest expense and the portions of rentals representative of the interest factor.

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DILUTION

If there is a material dilution of the investor's equity interest from the sale of common equity securities offered under this prospectus, we will set forth in any prospectus supplement the following information regarding any such material dilution of the equity interests of purchasers purchasing securities in an offering under this prospectus:

the net tangible book value per share of our equity securities before and after the offering;

the amount of the increase in such net tangible book value per share attributable to the cash payments made by purchasers in the offering; and

the amount of the immediate dilution from the public offering price which will be absorbed by such purchasers.

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DESCRIPTION OF SECURITIES

The descriptions of the securities contained in this prospectus, together with the applicable prospectus supplements, summarize the material terms and provisions of the various types of securities that we may offer. We will describe in the applicable prospectus supplement relating to any securities the particular terms of the securities offered by that prospectus supplement. If we so indicate in the applicable prospectus supplement, the terms of the securities may differ from the terms we have summarized below. We will also include in the prospectus supplement information, where applicable, about material U.S. federal income tax considerations relating to the securities, and the securities exchange, if any, on which the securities will be listed.

We may sell from time to time common stock, preferred stock, debt securities, warrants to purchase any such securities or any combination of the foregoing.

In this prospectus, we refer to the common stock, preferred stock, debt securities and warrants to be sold by us collectively as securities. The total dollar amount of all securities that we may issue under the registration statement of which this prospectus forms a part will not exceed \$250,000,000.

If we issue debt securities at a discount from their original stated principal amount, then we will use the issue price, and not the principal amount, of such debt securities for purposes of calculating the total dollar amount of all securities issued under this prospectus.

This prospectus may not be used to consummate a sale of securities unless it is accompanied by a prospectus supplement.

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DESCRIPTION OF CAPITAL STOCK

Agenus is authorized to issue up to 240,000,000 shares of common stock, par value \$0.01 per share, with 99,743,257 issued and outstanding as of September 30, 2017. Agenus is also authorized to issue up to 5,000,000 shares of preferred stock, par value \$0.01 per share, with 31,620 shares of Series A-1 convertible preferred stock issued and outstanding as of September 30, 2017.

The following description of our capital stock does not purport to be complete and is qualified in all respects by reference to our restated certificate of incorporation, as amended, and restated bylaws, and the Delaware General Corporation Law (DGCL).

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DESCRIPTION OF COMMON STOCK

General

Each share of Agenus common stock has the same relative rights and is identical in all respects with each other share of common stock.

Voting Rights

Subject to preferences that may apply to shares of preferred stock outstanding at the time, the holders of outstanding shares of common stock are entitled to receive dividends out of assets legally available for payment of dividends, as the board may from time to time determine. Each stockholder is entitled to one vote for each share of common stock held on all matters submitted to a vote of stockholders. Our certificate of incorporation does not provide for cumulative voting for the election of directors, which means that the holders of a majority of the shares voted can elect all of the directors then standing for election. The common stock is not entitled to preemptive rights and is not subject to conversion or redemption. Each outstanding share of common stock offered by this prospectus will, when issued, be fully paid and nonassessable.

Dividends

Holders of common stock are entitled to share ratably in any dividends declared by our board of directors, subject to any preferential dividend rights of any outstanding preferred stock. Dividends consisting of shares of common stock may be paid to holders of shares of common stock. We have never declared or paid cash dividends on our common stock. We do not intend to pay cash dividends in the foreseeable future.

Preemptive Rights

Holders of common stock do not have any preemptive rights with respect to any shares that may be issued by Agenus in the future. Thus, Agenus may sell shares of its common stock without first offering them to the then holders of common stock.

Liquidation

In the event of any liquidation or dissolution of Agenus, whether voluntary or involuntary, the holders of Agenus common stock would be entitled to receive pro rata, after payment of all debts and liabilities of Agenus, all assets of Agenus available for distribution, subject to the rights of the holders of any preferred stock which may be issued with a priority in liquidation or dissolution over the holders of common stock.

Listing

Our common stock is listed on The NASDAQ Capital Market under the symbol AGEN. On October 17, 2017, the last reported sale price for our common stock on The NASDAQ Capital Market was \$4.67 per share. As of October 11, 2017, we had approximately 407 stockholders of record.

Anti-Takeover Provisions

Statutory Business Combination Provision

Delaware has adopted a business combination statute (Section 203 of the DGCL) that may also have additional anti-takeover effects to provisions in Agenus' restated certificate of incorporation and bylaws. Section 203 of the DGCL, which, subject to certain exceptions, prohibits a Delaware corporation from engaging in any business combination with an interested stockholder for a period of three years following the time that such stockholder became an interested stockholder, unless:

the board of directors of the corporation approves either the business combination or the transaction that resulted in the stockholder becoming an interested stockholder, prior to the time the interested stockholder attained that status;

upon the closing of the transaction that resulted in the stockholder becoming an interested stockholder, the interested stockholder owned at least 85% of the voting stock of the corporation outstanding at the time the transaction commenced, excluding, for purposes of determining the number of shares outstanding, those shares owned by persons who are directors and also officers and by employee stock plans in which employee participants do not have the right to determine confidentially whether shares held subject to the plan will be tendered in a tender or exchange offer; or

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at or subsequent to such time, the business combination is approved by the board of directors and authorized at an annual or special meeting of stockholders, and not by written consent, by the affirmative vote of at least 66 2/3% of the outstanding voting stock that is not owned by the interested stockholder.

With certain exceptions, an interested stockholder is a person or group who or which owns 15% or more of the corporation's outstanding voting stock (including any rights to acquire stock pursuant to an option, warrant, agreement, arrangement or understanding, or upon the exercise of conversion or exchange rights, and stock with respect to which the person has voting rights only), or is an affiliate or associate of the corporation and was the owner of 15% or more of such voting stock at any time within the previous three years.

In general, Section 203 defines a business combination to include:

any merger or consolidation involving the corporation and the interested stockholder;

any sale, transfer, pledge or other disposition of 10% or more of the assets of the corporation involving the interested stockholder;

subject to certain exceptions, any transaction that results in the issuance or transfer by the corporation of any stock of the corporation to the interested stockholder;

any transaction involving the corporation that has the effect of increasing the proportionate share of the stock or any class or series of the corporation beneficially owned by the interested stockholder; or

the receipt by the interested stockholder of the benefit of any loans, advances, guarantees, pledges or other financial benefits provided by or through the corporation.

A Delaware corporation may opt out of this provision with an express provision in its original certificate of incorporation or an express provision in its amended and restated certificate of incorporation or bylaws resulting from a stockholders' amendment approved by at least a majority of the outstanding voting shares. However, Agenus has not opted out of this provision. Section 203 could prohibit or delay mergers or other takeover or change-in-control attempts and, accordingly, may discourage attempts to acquire Agenus.

Size of the Board and Vacancies

Our bylaws provide that the exact number of directors is determined by resolution of the board of directors or by the stockholders at the annual meeting. Our board of directors has the right to fill any vacancies resulting from death, resignation, disqualification or removal, as well as any newly created directorships arising from an increase in the size of the board.

Amendment of Charter Provisions

The affirmative vote of the holders of at least a majority of the voting power of all then outstanding shares of our voting stock, voting together as a single class, is required to, among other things, amend, alter, change or repeal certain provisions of our restated certificate of incorporation. Our bylaws may only be amended (or new bylaws

adopted) by the board of directors or the affirmative vote of the holders of at least a majority of the voting power of all then outstanding shares of our voting stock represented in person or by proxy at the meeting at which the amendment is voted on.

Undesignated Preferred Stock

We could issue preferred stock that could have other rights, including economic rights senior to our common stock, so that the issuance of the preferred stock could adversely affect the market value of our common stock. The issuance of the preferred stock may also have the effect of delaying, deferring or preventing a change in control of Agenus without any action by the stockholders. The effects of issuing preferred stock could include one or more of the following:

restricting dividends on the common stock;

diluting the voting power of the common stock;

impairing the liquidation rights of the common stock; or

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discouraging, delaying or preventing changes in control or management of Agenus.

Transfer Agent

The transfer agent and registrar for Agenus common stock is American Stock Transfer & Trust Company. Its telephone number is (800) 937-5449.

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DESCRIPTION OF PREFERRED STOCK

We currently have authorized 5,000,000 shares of preferred stock, of which 31,620 shares have been designated as series A-1 convertible preferred stock. As of the date of this prospectus, all of the series A-1 convertible preferred stock is issued and outstanding in the amount described in the preceding sentence. The remaining 4,968,380 authorized shares of preferred stock are undesignated and not issued or outstanding as of the date of this prospectus. As of the date of this prospectus, we do not have any equity securities that would be senior to, or on par with, our authorized preferred stock.

Series A Preferred Stock

On September 24, 2003, we sold 31,620 shares of series A convertible preferred stock, par value \$.01 per share, which we refer to as series A preferred stock. Under the terms and conditions of the Certificate of Designation creating the series A preferred stock, the stock is convertible by the holder at any time into shares of our common stock, is non-voting, carries a 2.5 percent annual dividend yield, has an initial conversion price of \$94.86, and is redeemable by us at its face amount on or after September 24, 2013. In February 2013, we entered into a Securities Exchange Agreement with the holder of our series A preferred stock pursuant to which the holder exchanged all 31,620 of the then outstanding shares of our series A preferred stock for an equivalent number of shares of series A-1 convertible preferred stock (series A-1 preferred stock). The terms of the series A-1 preferred stock are materially identical to the series A preferred stock, except that shares of the series A-1 preferred stock accrue a 0.63% annual dividend, as compared to a 2.5% annual dividend for the series A preferred stock. After giving effect to the transaction, no shares of the series A preferred stock remain outstanding. The liquidation value of the series A-1 preferred stock is equal to \$1,000 per share outstanding plus any accrued and unpaid dividends. The Certificate of Designation does not restrict the repurchase or redemption of shares by us while there is an arrearage in the payment of dividends. The Certificate of Designation does not contemplate a sinking fund. This description of the series A-1 preferred stock is qualified in its entirety by reference to the Certificate of Designation.

Undesignated Preferred Stock

This section describes the general terms and provisions of the preferred stock that we may offer by this prospectus. The prospectus supplement will describe the specific terms of the series of the preferred stock offered through that prospectus supplement. Those terms may differ from the terms discussed below. Any series of preferred stock we will issue will be governed by our restated certificate of incorporation, as amended, including the certificate of designations relating to such series of preferred stock, and our bylaws, as amended. In this section entitled Description of Preferred Stock, references to Agenus, we, our and us refer only to Agenus Inc. and not to its consolidated subsidiaries.

We will fix the rights, preferences, privileges and restrictions of the preferred stock of each series in the certificate of designations relating to that series. We will incorporate by reference as an exhibit to the registration statement that includes this prospectus the form of any certificate of designations which describes the terms of the series of preferred stock we are offering before the issuance of the related series of preferred stock. This description will include the following, to the extent applicable:

the title and stated value;

the number of shares we are offering;

the liquidation preference per share;

the purchase price;

the dividend rate, period and payment date, and method of calculation for dividends, if any;

whether any dividends will be cumulative or non-cumulative and, if cumulative, the date from which dividends will accumulate;

the provisions for a sinking fund, if any;

the provisions for redemption or repurchase, if applicable, and any restrictions on our ability to exercise those redemption and repurchase rights;

any listing of the preferred stock on any securities exchange or market;

whether the preferred stock will be convertible into our common stock and, if applicable, the conversion price, or how it will be calculated, and the conversion period;

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whether the preferred stock will be exchangeable into debt securities and, if applicable, the exchange price, or how it will be calculated, and the exchange period;

voting rights, if any, of the preferred stock;

preemptive rights, if any;

restrictions on transfer, sale or other assignment, if any;

whether interests in the preferred stock will be represented by depositary shares;

a discussion of any material or special United States federal income tax considerations applicable to the preferred stock;

the relative ranking and preferences of the preferred stock as to dividend rights and rights if we liquidate, dissolve or wind up our affairs; any limitations on issuance of any class or series of preferred stock ranking senior to or on a parity with the series of preferred stock as to dividend rights and rights if we liquidate, dissolve or wind up our affairs; and

any other specific terms, preferences, rights or limitations of, or restrictions on, the preferred stock.

When we issue shares of preferred stock under this prospectus, the shares, when issued in accordance with the terms of the applicable agreement, will be validly issued, fully paid and non-assessable and will not have, or be subject to, any preemptive or similar rights.

Section 242 of DGCL provides that the holders of each class or series of stock will have the right to vote separately as a class on certain amendments to our restated certificate of incorporation that would affect the class or series of preferred stock, as applicable. This right is in addition to any voting rights that may be provided for in the applicable certificate of designation.

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DESCRIPTION OF WARRANTS

We may issue warrants or other rights. We may issue these securities in such amounts or in as many distinct series as we wish. This section summarizes the terms of these securities that apply generally. We will describe the financial and other specific terms of any such series of securities in the prospectus supplement accompanying this prospectus. Those terms may vary from the terms described here.

When we refer to a series of securities in this section, we mean all securities issued as part of the same series under any applicable indenture, agreement or other instrument. When we refer to the prospectus supplement, we mean the applicable prospectus supplement describing the specific terms of the security you purchase. The terms used in the prospectus supplement will have the meanings described in this prospectus, unless otherwise specified.

In this section entitled Description of Warrants, references to Agenus, we, our and us refer only to Agenus Inc. and its consolidated subsidiaries. Also, in this section, references to holders mean those who own warrants or other rights registered in their own names, on the books that we or any applicable trustee or warrant or rights agent maintain for this purpose, and not those who own beneficial interests in warrants registered in street name or in warrants issued in book-entry form through one or more depositories. Owners of beneficial interests in warrants should also read the section entitled Legal Ownership and Book-Entry Issuance.

Warrants

The following description of warrants does not purport to be complete and is qualified in its entirety by reference to the description of a particular series of warrants contained in an applicable prospectus supplement. For information relating to common stock and preferred stock, see Description of Common Stock and Description of Preferred Stock, respectively.

We may offer by means of this prospectus warrants for the purchase of our preferred stock or common stock. We may issue warrants separately or together with any other securities offered by means of this prospectus, and the warrants may be attached to or separate from such securities. Each series of warrants will be issued under a separate warrant agreement to be entered into between us and a warrant agent specified therein. The warrant agent will act solely as our agent in connection with the warrants of such series and will not assume any obligation or relationship of agency or trust for or with any holders or beneficial owners of warrants.

Agreements

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any warrants we issue pursuant to this prospectus. Each series of warrants may be evidenced by certificates and may be issued under a separate indenture, agreement or other instrument to be entered into between us and a bank that we select as agent with respect to such series. The agent, if any, will have its principal office in the U.S. and have a combined capital and surplus of at least \$50,000,000. Warrants in book-entry form will be represented by a global security registered in the name of a depository, which will be the holder of all the securities represented by the global security. Those who own beneficial interests in a global security will do so through participants in the depository's system, and the rights of these indirect owners will be governed solely by the applicable procedures of the depository and its participants. We describe book-entry securities under Legal Ownership and Book-Entry Issuance.

General Terms of Warrants

The prospectus supplement relating to a series of warrants will identify the name and address of the warrant agent, if any. The prospectus supplement will describe the following terms, where applicable, of the warrants in respect of which this prospectus is being delivered:

the title and issuer of the warrants;

the aggregate number of warrants;

the price or prices at which the warrants will be issued;

the currencies in which the price or prices of the warrants may be payable;

the designation, amount and terms of the securities purchasable upon exercise of the warrants;

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the designation and terms of the other securities with which the warrants are issued and the number of warrants issued with each such security or each principal amount of security;

if applicable, the date on and after which the warrants and any related securities will be separately transferable;

any securities exchange or quotation system on which the warrants or any securities deliverable upon exercise of such securities may be listed;

the price or prices at which and currency or currencies in which the securities purchasable upon exercise of the warrants may be purchased;

the date on which the right to exercise the warrants shall commence and the date on which such right shall expire;

the minimum or maximum amount of warrants that may be exercised at any one time;

whether the warrants will be issued in fully registered for or bearer form, in global or non-global form or in any combination of these forms;

information with respect to book-entry procedures, if any;

a discussion of certain U.S. federal income tax considerations; and

any other material terms of the warrants, including terms, procedures and limitations relating to the exchange and exercise of the warrants.

Exercise of Warrants

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any warrants we issue pursuant to this prospectus. If any warrant is exercisable for other securities or other property, the following provisions will apply. Each such warrant may be exercised at any time up to any expiration date and time mentioned in the prospectus supplement relating to those warrants. After the close of business on any applicable expiration date, unexercised warrants will become void.

Warrants may be exercised by delivery of the certificate representing the securities to be exercised, or in the case of global securities, as described below under Legal Ownership and Book-Entry Issuance, by delivery of an exercise notice for those warrants, together with certain information, and payment to any agent in immediately available funds, as provided in the prospectus supplement, of the required purchase amount, if any. Upon receipt of payment and the certificate or exercise notice properly executed at the office indicated in the prospectus supplement, we will, in the

time period the relevant agreement provides, issue and deliver the securities or other property purchasable upon such exercise. If fewer than all of the warrants represented by such certificates are exercised, a new certificate will be issued for the remaining amount of warrants.

If mentioned in the prospectus supplement, securities may be surrendered as all or part of the exercise price for warrants.

Antidilution Provisions

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any warrants we issue pursuant to this prospectus. In the case of warrants to purchase common stock, the exercise price payable and the number of shares of common stock purchasable upon warrant exercise may be adjusted in certain events, including:

the issuance of a stock dividend to common shareholders or a combination, subdivision or reclassification of common stock;

the issuance of rights, warrants or options to all common and preferred shareholders entitling them to purchase common stock for an aggregate consideration per share less than the current market price per share of common stock;

any distribution to our common shareholders of evidences of our indebtedness of assets, excluding cash dividends or distributions referred to above; and

any other events mentioned in the prospectus supplement.

The prospectus supplement will describe which, if any, of these provisions shall apply to a particular series of warrants. Unless otherwise specified in the applicable prospectus supplement, no adjustment in the number of shares purchasable upon warrant exercise will be required until cumulative adjustments require an adjustment of at least 1% of such number and no fractional shares will be issued upon warrant exercise, but we will pay the cash value of any fractional shares otherwise issuable.

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Modification

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any warrants we issue pursuant to this prospectus. We and any agent for any series of warrants may amend any warrant or rights agreement and the terms of the related warrants by executing a supplemental agreement, without any such warrant holders' consent, for the purpose of:

curing any ambiguity, any defective or inconsistent provision contained in the agreement, or making any other corrections to the agreement that are not inconsistent with the provisions of the warrant certificates;

evidencing the succession of another corporation to us and its assumption of our covenants contained in the agreement and the securities;

appointing a successor depository, if the securities are issued in the form of global securities;

evidencing a successor agent's acceptance of appointment with respect to any securities;

adding to our covenants for the benefit of securityholders or surrendering any right or power we have under the agreement;

issuing warrants in definitive form, if such securities are initially issued in the form of global securities; or

amending the agreement and the warrants as we deem necessary or desirable and that will not adversely affect the interests of the applicable warrant holders in any material respect.

We and any agent for any series of warrants may also amend any agreement and the related warrants by a supplemental agreement with the consent of the holders of a majority of the warrants of any series affected by such amendment, for the purpose of adding, modifying or eliminating any of the agreement's provisions or of modifying the rights of the holders of warrants. However, no such amendment that:

reduces the number or amount of securities receivable upon any exercise of any such security;

shortens the time period during which any such security may be exercised;

otherwise adversely affects the exercise rights of warrant holders in any material respect; or

reduces the number of securities the consent of holders of which is required for amending the agreement or the related warrants;
may be made without the consent of each holder affected by that amendment.

Consolidation, Merger and Sale of Assets

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any warrants we issue pursuant to this prospectus. Any agreement with respect to warrants will provide that we are generally permitted to merge or consolidate with another corporation or other entity. Any such agreement will also provide that we are permitted to sell our assets substantially as an entirety to another corporation or other entity or to have another entity sell its assets substantially as an entirety to us. With regard to any series of warrants, however, we may not take any of these actions unless all of the following conditions are met:

if we are not the successor entity, the person formed by the consolidation or into or with which we merge or the person to which our properties and assets are conveyed, transferred or leased must be an entity organized and existing under the laws of the United States, any state or the District of Columbia and must expressly assume the performance of our covenants under any relevant indenture, agreement or other instrument; and

we or that successor corporation must not immediately be in default under that agreement.

Enforcement by Holders of Warrants

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any warrants we issue pursuant to this prospectus. Any agent for any series of warrants will act solely as our agent under the relevant agreement

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and will not assume any obligation or relationship of agency or trust for any securityholder. A single bank or trust company may act as agent for more than one issue of securities. Any such agent will have no duty or responsibility in case we default in performing our obligations under the relevant agreement or warrant, including any duty or responsibility to initiate any legal proceedings or to make any demand upon us. Any securityholder may, without the agent's consent or consent of any other securityholder, enforce by appropriate legal action its right to exercise any warrant exercisable for any property.

Replacement of Certificates

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any warrants we issue pursuant to this prospectus. We will replace any destroyed, lost, stolen or mutilated warrant or rights certificate upon delivery to us and any applicable agent of satisfactory evidence of the ownership of that certificate and of its destruction, loss, theft or mutilation, and (in the case of mutilation) surrender of that certificate to us or any applicable agent, unless we have, or the agent has, received notice that the certificate has been acquired by a bona fide purchaser. That securityholder will also be required to provide indemnity satisfactory to us and the relevant agent before a replacement certificate will be issued.

Title

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any warrants we issue pursuant to this prospectus. We, any agents for any series of warrants and any of their agents may treat the registered holder of any certificate as the absolute owner of the securities evidenced by that certificate for any purpose and as the person entitled to exercise the rights attaching to the warrants so requested, despite any notice to the contrary. See Legal Ownership and Book-Entry Issuance.

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DESCRIPTION OF DEBT SECURITIES

We will issue the debt securities offered by this prospectus and any accompanying prospectus supplement under an indenture to be entered into between Agenus and the trustee identified in the applicable prospectus supplement. The terms of the debt securities will include those stated in the indenture and those made part of the indenture by reference to the Trust Indenture Act of 1939, as in effect on the date of the indenture. We have filed a copy of the form of indenture as an exhibit to the registration statement in which this prospectus is included. The indenture will be subject to and governed by the terms of the Trust Indenture Act of 1939.

We may offer under this prospectus up to an aggregate principal amount of \$250,000,000 in debt securities; or if debt securities are issued at a discount, or in a foreign currency, foreign currency units or composite currency, the principal amount as may be sold for an initial public offering price of up to \$250,000,000. Unless otherwise specified in the applicable prospectus supplement, the debt securities will represent direct, unsecured obligations of Agenus and will rank equally with all of our other unsecured indebtedness.

The following statements relating to the debt securities and the indenture are summaries, qualified in their entirety to the detailed provisions of the indenture.

General

We may issue the debt securities in one or more series with the same or various maturities, at par, at a premium, or at a discount. We will describe the particular terms of each series of debt securities in a prospectus supplement relating to that series, which we will file with the SEC.

The prospectus supplement will set forth, to the extent required, the following terms of the debt securities in respect of which the prospectus supplement is delivered:

the title of the series;

the aggregate principal amount;

the issue price or prices, expressed as a percentage of the aggregate principal amount of the debt securities;

any limit on the aggregate principal amount;

the date or dates on which principal is payable;

the interest rate or rates (which may be fixed or variable) or, if applicable, the method used to determine such rate or rates;

the date or dates from which interest, if any, will be payable and any regular record date for the interest payable;

the place or places where principal and, if applicable, premium and interest, is payable;

the terms and conditions upon which we may, or the holders may require us to, redeem or repurchase the debt securities;

the denominations in which such debt securities may be issuable, if other than denominations of \$1,000, or any integral multiple of that number;

whether the debt securities are to be issuable in the form of certificated debt securities (as described below) or global debt securities (as described below);

the portion of principal amount that will be payable upon declaration of acceleration of the maturity date if other than the principal amount of the debt securities;

the currency of denomination;

the designation of the currency, currencies or currency units in which payment of principal and, if applicable, premium and interest, will be made;

if payments of principal and, if applicable, premium or interest, on the debt securities are to be made in one or more currencies or currency units other than the currency of denomination, the manner in which the exchange rate with respect to such payments will be determined;

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if amounts of principal and, if applicable, premium and interest may be determined by reference to an index based on a currency or currencies, or by reference to a commodity, commodity index, stock exchange index, or financial index, then the manner in which such amounts will be determined;

the provisions, if any, relating to any collateral provided for such debt securities;

any addition to or change in the covenants and/or the acceleration provisions described in this prospectus or in the indenture;

any events of default, if not otherwise described below under **Events of Default** ;

the terms and conditions, if any, for conversion into or exchange for shares of common stock or preferred stock;

any depositaries, interest rate calculation agents, exchange rate calculation agents, or other agents; and

the terms and conditions, if any, upon which the debt securities shall be subordinated in right of payment to other indebtedness of Agenus.

We may issue discount debt securities that provide for an amount less than the stated principal amount to be due and payable upon acceleration of the maturity of such debt securities in accordance with the terms of the indenture. We may also issue debt securities in bearer form, with or without coupons. If we issue discount debt securities or debt securities in bearer form, we will describe material U.S. federal income tax considerations and other material special considerations which apply to these debt securities in the applicable prospectus supplement.

We may issue debt securities denominated in or payable in a foreign currency or currencies or a foreign currency unit or units. If we do, we will describe the restrictions, elections, and general tax considerations relating to the debt securities and the foreign currency or currencies or foreign currency unit or units in the applicable prospectus supplement.

Exchange and/or Conversion Rights

We may issue debt securities which can be exchanged for or converted into shares of common stock or preferred stock. If we do, we will describe the terms of exchange or conversion in the prospectus supplement relating to these debt securities.

Transfer and Exchange

We may issue debt securities that will be represented by either:

book-entry securities, which means that there will be one or more global securities registered in the name of a depositary or a nominee of a depositary; or

certificated securities, which means that they will be represented by a certificate issued in definitive registered form.

We will specify in the prospectus supplement applicable to a particular offering whether the debt securities offered will be book-entry or certificated securities.

Certificated Debt Securities

Those who hold certificated debt securities may transfer or exchange such debt securities at the trustee's office or at the paying agent's office or agency in accordance with the terms of the indenture. There will be no service charge for any transfer or exchange of certificated debt securities but there may be a requirement to pay an amount sufficient to cover any tax or other governmental charge payable in connection with such transfer or exchange.

Those who hold certificated debt securities may effect the transfer of certificated debt securities and of the right to receive the principal of, premium, and/or interest, if any, on the certificated debt securities only by surrendering the certificate representing the certificated debt securities and having us or the trustee issue a new certificate to the new holder.

Global Securities

If we decide to issue debt securities in the form of one or more global securities, then we will register the global securities in the name of the depositary for the global securities or the nominee of the depositary, and the global securities will be delivered by the trustee to the depositary for credit to the accounts of the holders of beneficial interests in the debt securities.

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The prospectus supplement will describe the specific terms of the depositary arrangement for debt securities of a series that are issued in global form. None of the Company, the trustee, any payment agent, or the security registrar will have any responsibility or liability for any aspect of the records relating to or payments made on account of beneficial ownership interests in a global debt security or for maintaining, supervising, or reviewing any records relating to these beneficial ownership interests.

No Protection in the Event of Change of Control

The indenture does not have any covenants or other provisions providing for a put or increased interest or otherwise that would afford holders of debt securities additional protection in the event of a recapitalization transaction, a change of control of Agenus, or a highly leveraged transaction. If we offer any covenants or provisions of this type with respect to any debt securities covered by this prospectus, we will describe them in the applicable prospectus supplement.

Covenants

Unless otherwise indicated in this prospectus or a prospectus supplement, the debt securities will not have the benefit of any covenants that limit or restrict our business or operations, the pledging of our assets, or the incurrence by us of indebtedness. We will describe in the applicable prospectus supplement any material covenants in respect of a series of debt securities.

Consolidation, Merger, and Sale of Assets

We have agreed in the indenture that we will not consolidate with or merge into any other person, or convey, transfer, sell, or lease our properties and assets substantially as an entirety to any person, unless:

the person formed by the consolidation or into or with which we are merged or the person to which our properties and assets are conveyed, transferred, sold, or leased, is a corporation organized and existing under the laws of the U.S., any state or the District of Columbia, or a corporation or comparable legal entity organized under the laws of a foreign jurisdiction and, if we are not the surviving person, the surviving person has expressly assumed all of our obligations, including the payment of the principal of, and premium, if any, and interest on the debt securities and the performance of the other covenants under the indenture; and

immediately after giving effect to the transaction, no event of default, and no event which, after notice or lapse of time or both, would become an event of default, has occurred and is continuing under the indenture.

Events of Default

Unless otherwise specified in the applicable prospectus supplement, the following events will be events of default under the indenture with respect to debt securities of any series:

we fail to pay any principal or premium, if any, when it becomes due;

we fail to pay any interest within 30 days after it becomes due;

we fail to comply with any other covenant in the debt securities or the indenture for 60 days after written notice specifying the failure from the trustee or the holders of not less than 25% in aggregate principal amount of the outstanding debt securities of that series; and

certain events involving bankruptcy, insolvency, or reorganization of Agenus or any of our significant subsidiaries.

The trustee may withhold notice to the holders of the debt securities of any series of any default, except in payment of principal of, or premium, if any, or interest on the debt securities of a series, if the trustee considers it to be in the best interest of the holders of the debt securities of that series to do so.

If an event of default (other than an event of default resulting from certain events of bankruptcy, insolvency, or reorganization) occurs, and is continuing, then the trustee or the holders of not less than 25% in aggregate principal amount of the outstanding debt securities of any series may accelerate the maturity of the debt securities. If this happens, the entire principal amount, plus the premium, if any, of all the outstanding debt securities of the affected series plus accrued interest to

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the date of acceleration will be immediately due and payable. At any time after the acceleration, but before a judgment or decree based on such acceleration is obtained by the trustee, the holders of a majority in aggregate principal amount of outstanding debt securities of such series may rescind and annul such acceleration if:

all events of default (other than nonpayment of accelerated principal, premium, or interest) have been cured or waived;

all lawful interest on overdue interest and overdue principal has been paid; and

the rescission would not conflict with any judgment or decree.

In addition, if the acceleration occurs at any time when Agenus has outstanding indebtedness which is senior to the debt securities, the payment of the principal amount of outstanding debt securities may be subordinated in right of payment to the prior payment of any amounts due under the senior indebtedness, in which case the holders of debt securities will be entitled to payment under the terms prescribed in the instruments evidencing the senior indebtedness and the indenture.

If an event of default resulting from certain events of bankruptcy, insolvency, or reorganization occurs, the principal, premium, and interest amount with respect to all of the debt securities of any series will be due and payable immediately without any declaration or other act on the part of the trustee or the holders of the debt securities of that series.

The holders of a majority in principal amount of the outstanding debt securities of a series will have the right to waive any existing default or compliance with any provision of the indenture or the debt securities of that series and to direct the time, method, and place of conducting any proceeding for any remedy available to the trustee, subject to certain limitations specified in the indenture.

No holder of any debt security of a series will have any right to institute any proceeding with respect to the indenture or for any remedy under the indenture, unless:

the holder gives to the trustee written notice of a continuing event of default;

the holders of at least 25% in aggregate principal amount of the outstanding debt securities of the affected series make a written request and offer reasonable indemnity to the trustee to institute a proceeding as trustee;

the trustee fails to institute a proceeding within 60 days after such request; and

the holders of a majority in aggregate principal amount of the outstanding debt securities of the affected series do not give the trustee a direction inconsistent with such request during such 60-day period.

These limitations do not, however, apply to a suit instituted for payment on debt securities of any series on or after the due dates expressed in the debt securities.

Modification and Waiver

From time to time, we and the trustee may, without the consent of holders of the debt securities of one or more series, amend the indenture or the debt securities of one or more series, or supplement the indenture, for certain specified purposes, including:

to provide that the surviving entity following a change of control of Agenus permitted under the indenture will assume all of our obligations under the indenture and debt securities;

to provide for certificated debt securities in addition to uncertificated debt securities;

to comply with any requirements of the SEC under the Trust Indenture Act of 1939;

to cure any ambiguity, defect, or inconsistency, or make any other change that does not materially and adversely affect the rights of any holder; and

to appoint a successor trustee under the indenture with respect to one or more series.

From time to time, we and the trustee may, with the consent of holders of at least a majority in principal amount of the outstanding debt securities, amend or supplement the indenture or the debt securities, or waive compliance in a particular instance by us with any provision of the indenture or the debt securities. We may not, however, without the consent of each holder affected by such action, modify or supplement the indenture or the debt securities, or waive compliance with any

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provision of the indenture or the debt securities in order to:

reduce the amount of debt securities whose holders must consent to an amendment, supplement, or waiver to the indenture or such debt security;

reduce the rate of or change the time for payment of interest;

reduce the principal of or change the stated maturity of the debt securities;

make any debt security payable in money other than that stated in the debt security;

change the amount or time of any payment required, or reduce the premium payable upon any redemption, or change the time before which no such redemption may be made;

waive a default in the payment of the principal of, premium, if any, or interest on the debt securities or a redemption payment; or

take any other action otherwise prohibited by the indenture to be taken without the consent of each holder affected by the action.

Defeasance of Debt Securities and Certain Covenants in Certain Circumstances

The indenture permits us, at any time, to elect to discharge our obligations with respect to one or more series of debt securities by following certain procedures described in the indenture. These procedures will allow us either:

to defease and be discharged from any and all of our obligations with respect to any debt securities except for the following obligations (which discharge is referred to as legal defeasance):

- (1) to register the transfer or exchange of such debt securities;
- (2) to replace temporary or mutilated, destroyed, lost, or stolen debt securities;
- (3) to compensate and indemnify the trustee; or
- (4) to maintain an office or agency in respect of the debt securities and to hold monies for payment in trust; or

to be released from our obligations with respect to the debt securities under certain covenants contained in the indenture, as well as any additional covenants which may be contained in the applicable supplemental

indenture (which release is referred to as covenant defeasance).

In order to exercise either defeasance option, we must deposit with the trustee or other qualifying trustee, in trust for that purpose:

money;

U.S. Government Obligations (as described below) or Foreign Government Obligations (as described below) which through the scheduled payment of principal and interest in accordance with their terms will provide money; or

a combination of money and/or U.S. Government Obligations and/or Foreign Government Obligations sufficient in the written opinion of a nationally-recognized firm of independent accountants to provide money;

which in each case specified above, provides a sufficient amount to pay the principal of, premium, if any, and interest, if any, on the debt securities of the series, on the scheduled due dates or on a selected date of redemption in accordance with the terms of the indenture.

In addition, defeasance may be effected only if, among other things:

in the case of either legal or covenant defeasance, we deliver to the trustee an opinion of counsel, as specified in the indenture, stating that as a result of the defeasance neither the trust nor the trustee will be required to register as an investment company under the Investment Company Act of 1940;

in the case of legal defeasance, we deliver to the trustee an opinion of counsel stating that we have received from, or there has been published by, the Internal Revenue Service a ruling to the effect that, or there has been a change in any applicable federal income tax law with the effect that (and the opinion shall confirm that), the holders of outstanding debt securities will not recognize income, gain, or loss for U.S. federal income tax purposes solely as a result of such legal defeasance and will be subject to U.S. federal income tax on the same amounts, in the same manner, including as a result of prepayment, and at the same times as would have been the case if legal defeasance had not occurred;

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in the case of covenant defeasance, we deliver to the trustee an opinion of counsel to the effect that the holders of the outstanding debt securities will not recognize income, gain, or loss for U.S. federal income tax purposes as a result of covenant defeasance and will be subject to U.S. federal income tax on the same amounts, in the same manner, and at the same times as would have been the case if covenant defeasance had not occurred; and

certain other conditions described in the indenture are satisfied.

If we fail to comply with our remaining obligations under the indenture and applicable supplemental indenture after a covenant defeasance of the indenture and applicable supplemental indenture, and the debt securities are declared due and payable because of the occurrence of any undefeased event of default, the amount of money and/or U.S. Government Obligations and/or Foreign Government Obligations on deposit with the trustee could be insufficient to pay amounts due under the debt securities of the affected series at the time of acceleration. We will, however, remain liable in respect of these payments.

The term "U.S. Government Obligations" as used in the above discussion means securities which are direct obligations of or non-callable obligations guaranteed by the United States of America for the payment of which obligation or guarantee the full faith and credit of the United States of America is pledged.

The term "Foreign Government Obligations" as used in the above discussion means, with respect to debt securities of any series that are denominated in a currency other than U.S. dollars (1) direct obligations of the government that issued or caused to be issued such currency for the payment of which obligations its full faith and credit is pledged or (2) obligations of a person controlled or supervised by or acting as an agent or instrumentality of such government the timely payment of which is unconditionally guaranteed as a full faith and credit obligation by that government, which in either case under clauses (1) or (2), are not callable or redeemable at the option of the issuer.

Regarding the Trustee

We will identify the trustee with respect to any series of debt securities in the prospectus supplement relating to the applicable debt securities. You should note that if the trustee becomes a creditor of Agenus, the indenture and the Trust Indenture Act of 1939 limit the rights of the trustee to obtain payment of claims in certain cases, or to realize on certain property received in respect of any such claim, as security or otherwise. The trustee and its affiliates may engage in, and will be permitted to continue to engage in, other transactions with us and our affiliates. If, however, the trustee acquires any conflicting interest within the meaning of the Trust Indenture Act of 1939, it must eliminate such conflict or resign.

The holders of a majority in principal amount of the then outstanding debt securities of any series may direct the time, method, and place of conducting any proceeding for exercising any remedy available to the trustee. If an event of default occurs and is continuing, the trustee, in the exercise of its rights and powers, must use the degree of care and skill of a prudent person in the conduct of his or her own affairs. Subject to that provision, the trustee will be under no obligation to exercise any of its rights or powers under the indenture at the request of any of the holders of the debt securities, unless they have offered to the trustee reasonable indemnity or security.

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DESCRIPTION OF UNITS

This section outlines some of the provisions of the units and the unit agreements that we may enter into. This information may not be complete in all respects and is qualified entirely by reference to the unit agreement with respect to the units of any particular series. The specific terms of any series of units will be described in the applicable prospectus supplement. If so described in a particular supplement, the specific terms of any series of units may differ from the general description of terms presented below.

In this section entitled Description of Units, references to Agenus, we, our and us refer only to Agenus Inc. and its consolidated subsidiaries. Also, in this section, references to holders mean those who own units registered in their own names, on the books that we or our agent maintain for this purpose, and not those who own beneficial interests in units registered in street name or in units issued in book-entry form through one or more depositories. Owners of beneficial interests in the units should read the section below entitled Legal Ownership and Book-Entry Issuance.

We may issue units comprised of one or more debt securities, shares of common stock, shares of preferred stock, and warrants in any combination. Each unit will be issued so that the holder of the unit is also the holder of each security included in the unit. Thus, the holder of a unit will have the rights and obligations of a holder of each included security. The unit agreement under which a unit is issued may provide that the securities included in the unit may not be held or transferred separately, at any time or at any time before a specified date.

The applicable prospectus supplement may describe:

the designation and terms of the units and of the securities comprising the units, including whether and under what circumstances those securities may be held or transferred separately;

any provisions of the governing unit agreement that differ from those described below; and

any provisions for the issuance, payment, settlement, transfer or exchange of the units or of the securities comprising the units.

The provisions described in this section, as well as those described under Description of Common Stock, Description of Preferred Stock, Description of Warrants and Description of Debt Securities will apply to the securities included in each unit, to the extent relevant.

Issuance in Series

We may issue units in such amounts and in as many distinct series as we wish. This section summarizes terms of the units that apply generally to all series. Most of the financial and other specific terms of your series will be described in the applicable prospectus supplement.

Unit Agreements

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any units we issue pursuant to this prospectus. We will issue the units under one or more unit agreements to be entered into between us and a bank or other financial institution, as unit agent. We may add, replace or terminate unit agents from

time to time. We will identify the unit agreement under which each series of units will be issued and the unit agent under that agreement in the applicable prospectus supplement.

The following provisions will generally apply to all unit agreements unless otherwise stated in the applicable prospectus supplement.

Enforcement of Rights

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any units we issue pursuant to this prospectus. The unit agent under a unit agreement will act solely as our agent in connection with the units issued under that agreement. The unit agent will not assume any obligation or relationship of agency or trust for or with any holders of those units or of the securities comprising those units. The unit agent will not be obligated to take any action on behalf of those holders to enforce or protect their rights under the units or the included securities.

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Except as indicated in the next paragraph, a holder of a unit may, without the consent of the unit agent or any other holder, enforce its rights as holder under any security included in the unit, in accordance with the terms of that security and the indenture, warrant agreement, rights agreement or other instrument under which that security is issued. Those terms are described elsewhere in this prospectus under the sections relating to debt securities, preferred stock, common stock or warrants, as relevant.

Notwithstanding the foregoing, a unit agreement may limit or otherwise affect the ability of a holder of units issued under that agreement to enforce its rights, including any right to bring a legal action, with respect to those units or any securities, other than debt securities, that are included in those units. Limitations of this kind will be described in the applicable prospectus supplement.

Modification without Consent of Holders

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any units we issue pursuant to this prospectus. We and the applicable unit agent may amend any unit or unit agreement without the consent of any holder:

to cure any ambiguity;

to correct or supplement any defective or inconsistent provision; or

to make any other change that we believe is necessary or desirable and will not adversely affect the interests of the affected holders in any material respect.

We do not need any approval to make changes that affect only units to be issued after the changes take effect. We may also make changes that do not adversely affect a particular unit in any material respect, even if they adversely affect other units in a material respect. In those cases, we do not need to obtain the approval of the holder of the unaffected unit; we need only obtain any required approvals from the holders of the affected units.

Modification with Consent of Holders

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any units we issue pursuant to this prospectus. We may not amend any particular unit or a unit agreement with respect to any particular unit unless we obtain the consent of the holder of that unit, if the amendment would:

impair any right of the holder to exercise or enforce any right under a security included in the unit if the terms of that security require the consent of the holder to any changes that would impair the exercise or enforcement of that right; or

reduce the percentage of outstanding units or any series or class the consent of whose holders is required to amend that series or class, or the applicable unit agreement with respect to that series or class, as described below.

Any other change to a particular unit agreement and the units issued under that agreement would require the following approval:

if the change affects only the units of a particular series issued under that agreement, the change must be approved by the holders of a majority of the outstanding units of that series; or

if the change affects the units of more than one series issued under that agreement, it must be approved by the holders of a majority of all outstanding units of all series affected by the change, with the units of all the affected series voting together as one class for this purpose.

These provisions regarding changes with majority approval also apply to changes affecting any securities issued under a unit agreement, as the governing document.

In each case, the required approval must be given by written consent.

Unit Agreements Will Not Be Qualified Under Trust Indenture Act

No unit agreement will be qualified as an indenture, and no unit agent will be required to qualify as a trustee, under the Trust Indenture Act. Therefore, holders of units issued under unit agreements will not have the protections of the Trust Indenture Act with respect to their units.

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Mergers and Similar Transactions Permitted; No Restrictive Covenants or Events of Default

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any units we issue pursuant to this prospectus. The unit agreements will not restrict our ability to merge or consolidate with, or sell our assets to, another corporation or other entity or to engage in any other transactions. If at any time we merge or consolidate with, or sell our assets substantially as an entirety to, another corporation or other entity, the successor entity will succeed to and assume our obligations under the unit agreements. We will then be relieved of any further obligation under these agreements.

The unit agreements will not include any restrictions on our ability to put liens on our assets, including our interests in our subsidiaries, nor will they restrict our ability to sell our assets. The unit agreements also will not provide for any events of default or remedies upon the occurrence of any events of default.

Governing Law

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any units we issue pursuant to this prospectus. The unit agreements and the units will be governed by New York law.

Form, Exchange and Transfer

Unless otherwise provided in the applicable prospectus supplement, the following provisions will apply to any units we issue pursuant to this prospectus. We will issue each unit in global that is, book-entry form only. Units in book-entry form will be represented by a global security registered in the name of a depository, which will be the holder of all the units represented by the global security. Those who own beneficial interests in a unit will do so through participants in the depository's system, and the rights of these indirect owners will be governed solely by the applicable procedures of the depository and its participants. We describe book-entry securities below under Legal Ownership and Book-Entry Issuance.

In addition, we will issue each unit in registered form, unless we say otherwise in the applicable prospectus supplement. Bearer securities would be subject to special provisions, as we describe below under Securities Issued in Bearer Form.

Each unit and all securities comprising the unit will be issued in the same form.

If we issue any units in registered, non-global form, the following will apply to them.

The units will be issued in the denominations stated in the applicable prospectus supplement. Holders may exchange their units for units of smaller denominations or combined into fewer units of larger denominations, as long as the total amount is not changed.

Holders may exchange or transfer their units at the office of the unit agent. Holders may also replace lost, stolen, destroyed or mutilated units at that office. We may appoint another entity to perform these functions or perform them ourselves.

Holders will not be required to pay a service charge to transfer or exchange their units, but they may be required to pay for any tax or other governmental charge associated with the transfer or exchange. The transfer or exchange, and any replacement, will be made only if our transfer agent is satisfied with the holder's proof of legal ownership. The transfer agent may also require an indemnity before replacing any units.

If we have the right to redeem, accelerate or settle any units before their maturity, and we exercise our right as to less than all those units or other securities, we may block the exchange or transfer of those units during the period beginning 15 days before the day we mail the notice of exercise and ending on the day of that mailing, in order to freeze the list of holders to prepare the mailing. We may also refuse to register transfers of or exchange any unit selected for early settlement, except that we will continue to permit transfers and exchanges of the unsettled portion of any unit being partially settled. We may also block the transfer or exchange of any unit in this manner if the unit includes securities that are or may be selected for early settlement.

Only the depositary will be entitled to transfer or exchange a unit in global form, since it will be the sole holder of the unit.

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Payments and Notices

In making payments and giving notices with respect to our units, we will follow the procedures we plan to use with respect to our debt securities, where applicable.

LEGAL OWNERSHIP AND BOOK-ENTRY ISSUANCE

We can issue securities in registered form or in the form of one or more global securities. We describe global securities in greater detail below. We refer to those persons who have securities registered in their own names on the books that we or any applicable depository or warrant agent maintain for this purpose as the holders of those securities. These persons are the legal holders of the securities. We refer to those persons who, indirectly through others, own beneficial interests in securities that are not registered in their own names, as indirect holders of those securities. As we discuss below, indirect holders are not legal holders, and investors in securities issued in book-entry form or in street name will be indirect holders.

Book-Entry Holders

Except as described below, we will issue securities in book-entry form only, as we will specify in the applicable prospectus supplement. This means securities may be represented by one or more global securities registered in the name of a financial institution that holds them as depository on behalf of other financial institutions that participate in the depository's book-entry system. These participating institutions, which are referred to as participants, in turn, hold beneficial interests in the securities on behalf of themselves or their customers.

Only the person in whose name a security is registered is recognized as the holder of that security. Global securities will be registered in the name of the depository or its participants. Consequently, for global securities, we will recognize only the depository as the holder of the securities, and we will make all payments on the securities to the depository. The depository passes along the payments it receives to its participants, which in turn pass the payments along to their customers who are the beneficial owners. The depository and its participants do so under agreements they have made with one another or with their customers; they are not obligated to do so under the terms of the securities.

As a result, investors in a book-entry security will not own securities directly. Instead, they will own beneficial interests in a global security, through a bank, broker or other financial institution that participates in the depository's book-entry system or holds an interest through a participant. As long as the securities are issued in global form, investors will be indirect holders, and not legal holders, of the securities.

Street Name Holders

We may terminate a global security or issue securities that are not issued in global form. In these cases, investors may choose to hold their securities in their own names or in street name. Securities held by an investor in street name would be registered in the name of a bank, broker or other financial institution that the investor chooses, and the investor would hold only a beneficial interest in those securities through an account he or she maintains at that institution.

For securities held in street name, we or any applicable depository will recognize only the intermediary banks, brokers and other financial institutions in whose names the securities are registered as the holders of those securities, and we or any such depository will make all payments on those securities to them. These institutions pass along the payments they receive to their customers who are the beneficial owners, but only because they agree to do so in their customer

agreements or because they are legally required to do so. Investors who hold securities in street name will be indirect holders, not legal holders, of those securities.

Legal Holders

Our obligations, as well as the obligations of any applicable depositary or warrant agent or other third party employed by us or any of the foregoing, run only to the legal holders of the securities. We do not have obligations to investors who hold beneficial interests in global securities, in street name or by any other indirect means. This will be the case whether an investor chooses to be an indirect holder of a security or has no choice because we are issuing the securities only in global form.

For example, once we make a payment or give a notice to the holder, we have no further responsibility for the payment or notice even if that holder is required, under agreements with depositary participants or customers or by law, to pass it along to the indirect holders but does not do so. Similarly, we may want to obtain the approval of the holders to amend an instrument defining the rights of security holders, to relieve us of the consequences of a breach or of our or its obligation to comply with a particular provision of such an instrument or for other purposes. In such an event, we would seek approval only from the legal holders, and not the indirect holders, of the securities. Whether and how the holders contact the indirect holders is up to the legal holders.

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Special Considerations for Indirect Holders

If you hold securities through a bank, broker or other financial institution, either in book-entry form or in street name, you should check with your own institution to find out:

how it handles securities payments and notices;

whether it imposes fees or charges;

how it would handle a request for the holders' consent, if ever required;

whether and how you can instruct it to send you securities registered in your own name so you can be a holder, if that is permitted in the future;

how it would exercise rights under the securities if there were a default or other event triggering the need for holders to act to protect their interests; and

if the securities are in book-entry form, how the depository's rules and procedures will affect these matters.

Global Securities

A global security is a security that represents one or any other number of individual securities held by a depository. Generally, all securities represented by the same global securities will have the same terms.

Each security issued in book-entry form will be represented by a global security that we deposit with and register in the name of a financial institution or its nominee that we select. The financial institution that we select for this purpose is called the depository. Unless specified otherwise in the applicable prospectus supplement, The Depository Trust Company, New York, New York ("DTC"), will be the depository for all securities issued in book-entry form.

A global security may not be transferred to or registered in the name of anyone other than the depository, its nominee or a successor depository, unless special termination situations arise. We describe those situations below under "Special Situations When a Global Security Will Be Terminated." As a result of these arrangements, the depository, or its nominee, will be the sole registered owner and legal holder of all securities represented by a global security, and investors will be permitted to own only beneficial interests in a global security. Beneficial interests must be held by means of an account with a broker, bank or other financial institution that in turn has an account with the depository or with another institution that does. Thus, an investor whose security is represented by a global security will not be a legal holder of the security, but only an indirect holder of a beneficial interest in the global security.

If the prospectus supplement for a particular security indicates that the security will be issued in global form only, then the security will be represented by a global security at all times unless and until the global security is terminated. If termination occurs, we may issue the securities through another book-entry clearing system or decide that the securities may no longer be held through any book-entry clearing system.

Special Considerations for Global Securities

As an indirect holder, an investor's rights relating to a global security will be governed by the account rules of the investor's financial institution and of the depository, as well as general laws relating to securities transfers. We do not recognize an indirect holder as a legal holder of securities and instead deal only with the depository that holds the global security.

If securities are issued only in the form of a global security, an investor should be aware of the following:

An investor cannot cause the securities to be registered in his or her name, and cannot obtain non-global certificates for his or her interest in the securities, except in the special situations we describe below.

An investor will be an indirect holder and must look to his or her own bank or broker for payments on the securities and protection of his or her legal rights relating to the securities, as we describe above.

An investor may not be able to sell interests in the securities to some insurance companies and to other institutions that are required by law to own their securities in non-book-entry form.

An investor may not be able to pledge his or her interest in a global security in circumstances where certificates representing the securities must be delivered to the lender or other beneficiary of the pledge in order for the pledge to be effective.

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The depositary's policies, which may change from time to time, will govern payments, transfers, exchanges and other matters relating to an investor's interest in a global security. We and any applicable agent have no responsibility for any aspect of the depositary's actions or for its records of ownership interests in a global security. We and any applicable agent also will not supervise the depositary in any way.

The depositary may, and we understand that DTC will, require that those who purchase and sell interests in the global security within its book-entry system use immediately available funds, and your broker or bank may require you to do so as well.

Financial institutions that participate in the depositary's book-entry system, and through which an investor holds its interest in the global security, may also have their own policies affecting payments, notices and other matters relating to the securities.

There may be more than one financial intermediary in the chain of ownership for an investor. We do not monitor and are not responsible for the actions of any of those intermediaries.

Special Situations When A Global Security Will Be Terminated

In a few special situations described below, the global security will terminate, and interests in it will be exchanged for physical certificates representing those interests. After that exchange, the choice of whether to hold securities directly or in street name will be up to the investor. Investors must consult their own banks or brokers to find out how to have their interests in securities transferred to their own name, so that they will be direct holders. We have described the rights of holders and street name investors above.

The global security will terminate when the following special situations occur:

if the depositary notifies us that it is unwilling, unable or no longer qualified to continue as depositary for that global security and we do not appoint another institution to act as depositary within 90 days;

if we notify any applicable depositary or warrant agent that we wish to terminate that global security; or

if an event of default has occurred with regard to securities represented by that global security and has not been cured or waived.

The applicable prospectus supplement may also list additional situations for terminating a global security that would apply only to the particular series of securities covered by the prospectus supplement. When a global security terminates, the depositary, and not us or any applicable agent, is responsible for deciding the names of the institutions that will be the initial direct holders.

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PLAN OF DISTRIBUTION

We may sell the securities covered by this prospectus directly to purchasers or through underwriters, broker-dealers or agents, who may receive compensation in the form of discounts, concessions or commissions from us. These discounts, concessions or commissions as to any particular underwriter, broker-dealer or agent may be in excess of those customary in the types of transactions involved. In addition, we may issue the securities as a dividend or distribution or in a subscription rights offering to our existing security holders.

The securities may be sold in one or more transactions at fixed prices, at prevailing market prices at the time of sale, at varying prices determined at the time of sale or at negotiated prices. These sales may be effected in transactions which may involve crosses or block transactions.

If underwriters are used in an offering of securities, such offered securities may be resold in one or more transactions:

on any national securities exchange or quotation service on which the preferred stock or the common stock may be listed or quoted at the time of sale, including, as of the date of this prospectus, the NASDAQ Capital Market in the case of the common stock;

in the over-the-counter market;

in transactions otherwise than on these exchanges or services or in the over-the-counter market; or

through the writing of options, whether the options are listed on an options exchange or otherwise. Each prospectus supplement will state the terms of the offering, including, but not limited to:

the names of any underwriters, dealers, or agents;

the public offering or purchase price of the securities and the net proceeds that we will receive from the sale;

any underwriting discounts and commissions or other items constituting underwriters' compensation;

any discounts, commissions, or fees allowed or paid to dealers or agents; and

any securities exchange on which the offered securities may be listed.

If we sell securities to underwriters, we will execute an underwriting agreement with them at the time of the sale and will name them in the applicable prospectus supplement. In connection with these sales, the underwriters may be deemed to have received compensation in the form of underwriting discounts and commissions. The underwriters also

may receive commissions from purchasers of securities for whom they may act as agent. Unless we specify otherwise in the applicable prospectus supplement, the underwriters will not be obligated to purchase the securities unless the conditions set forth in the underwriting agreement are satisfied, and if the underwriters purchase any of the securities offered by such prospectus supplement, they will be required to purchase all of such offered securities. The underwriters may acquire the securities for their own account and may resell the securities from time to time in one or more transactions, including negotiated transactions, at a fixed public offering price or varying prices determined at the time of sale. The underwriters may sell the securities to or through dealers, and those dealers may receive discounts, concessions, or commissions from the underwriters as well as from the purchasers for whom they may act as agent.

We may designate agents who agree to use their reasonable efforts to solicit purchasers for the period of their appointment or to sell securities on a continuing basis. We may also sell securities directly to one or more purchasers without using underwriters or agents.

Under agreements entered into with us, underwriters and agents may be entitled to indemnification by us against certain civil liabilities, including liabilities under the Securities Act, or to contribution for payments the underwriters or agents may be required to make. The underwriters, agents, and their affiliates may engage in financial or other business transactions with us and our subsidiaries in the ordinary course of business.

The aggregate proceeds to us from the sale of the securities will be the purchase price of the securities less discounts and commissions, if any.

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In order to comply with the securities laws of certain states, if applicable, any securities covered by this prospectus must be sold in such jurisdictions only through registered or licensed brokers or dealers. In addition, in certain states securities may not be sold unless they have been registered or qualified for sale in the applicable state or an exemption from the registration or qualification requirement is available and is complied with.

In order to facilitate the offering of the securities, any underwriters may engage in transactions that stabilize, maintain or otherwise affect the price of the securities or any other securities the prices of which may be used to determine payments on such securities. Specifically, any underwriters may overallocate in connection with the offering, creating a short position for their own accounts. In addition, to cover overallocations or to stabilize the price of the securities or of any such other securities, the underwriters may bid for, and purchase, the securities or any such other securities in the open market. Finally, in any offering of the securities through a syndicate of underwriters, the underwriting syndicate may reclaim selling concessions allowed to an underwriter or a dealer for distributing the securities in the offering if the syndicate repurchases previously distributed securities in transactions to cover syndicate short positions, in stabilization transactions or otherwise. Any of these activities may stabilize or maintain the market price of the securities above independent market levels. Any such underwriters are not required to engage in these activities and may end any of these activities at any time.

The applicable prospectus supplement may provide that the original issue date for your securities may be more than three scheduled business days after the trade date for your securities. Accordingly, in such a case, if you wish to trade securities on any date prior to the third business day before the original issue date for your securities, you will be required, by virtue of the fact that your securities initially are expected to settle in more than three scheduled business days after the trade date for your securities, to make alternative settlement arrangements to prevent a failed settlement.

The securities may be new issues of securities and may have no established trading market. The securities may or may not be listed on a national securities exchange. We can make no assurance as to the liquidity of or the existence of trading markets for any of the securities.

In order to comply with the securities laws of some states, if applicable, the shares of common stock offered by this prospectus must be sold in such jurisdictions only through registered or licensed brokers or dealers. In addition, in some states the shares of common stock may not be sold unless they have been registered or qualified for sale in the applicable state or an exemption from the registration or qualification requirement is available and is complied with.

To the extent required, this prospectus may be amended or supplemented from time to time to describe a specific plan of distribution.

Transfer Agent

The transfer agent and registrar for Agenus common stock is American Stock Transfer & Trust Company. Its telephone number is (800) 937-5449.

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LEGAL MATTERS

The validity of the securities that may be offered hereby will be passed upon for us by Ropes & Gray LLP, Boston, Massachusetts. Additional legal matters may be passed upon for us or any underwriters, dealers or agents, by counsel that we will name in the applicable prospectus supplement.

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EXPERTS

The consolidated financial statements of Agenus Inc., as of December 31, 2016 and 2015, and for each of the years in the three-year period ended December 31, 2016, and management's assessment of the effectiveness of internal control over financial reporting as of December 31, 2016 have been incorporated by reference herein and in the registration statement in reliance upon the reports of KPMG LLP, independent registered public accounting firm, incorporated by reference herein, and upon the authority of said firm as experts in accounting and auditing.

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WHERE YOU CAN FIND MORE INFORMATION

Agenus Inc. is subject to the information requirements of the Exchange Act and files annual, quarterly and special reports, proxy statements and other information with the SEC. You may read and copy any materials we file with the SEC at the Public Reference Room of the SEC at Room 1580, 100 F Street, N.E., Washington, D.C. 20549. You may obtain information on the operation of the Public Reference Room by calling the SEC at 1-800-SEC-0330. In addition, we file many of our documents electronically with the SEC, and you may access those documents over the Internet. The SEC maintains a web site that contains reports, proxy and information statements and other information regarding issuers that file electronically with the SEC. The address of the SEC's website is <http://www.sec.gov>. Documents we have filed with the SEC are also available on our website through the investor link at www.agenusbio.com. Information contained on our web site does not constitute a part of this prospectus and is not incorporated by reference herein.

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INCORPORATION OF CERTAIN DOCUMENTS BY REFERENCE

The SEC allows us to incorporate by reference in this prospectus the information we file with the SEC. This helps us disclose certain important information to you by referring you to the documents we file. The information we incorporate by reference is an important part of this prospectus. Because we are incorporating by reference future filings with the SEC, this prospectus is continually updated and those future filings may modify or supersede some of the information included or incorporated in this prospectus. This means that you must look at all of the SEC filings that we incorporate by reference to determine if any of the statements in this prospectus or in any document previously incorporated by reference have been modified or superseded. We incorporate by reference each of the documents listed below.

our Annual Report on Form 10-K for the year ended December 31, 2016 (File No. 000-29089);

our Quarterly Report on Form 10-Q for the quarter ended March 31, 2017 (File No. 000-29089);

our Quarterly Report on Form 10-Q for the quarter ended June 30, 2017 (File No. 000-29089);

our Current Reports on Form 8-K filed on January 17, 2017, February 14, 2017, March 30, 2017, April 20, 2017 and June 30, 2017 (except, with respect to each of the foregoing, for portions of such reports which were deemed to be furnished and not filed) (File No. 000-29089);

our Proxy Statement on Schedule 14A filed with the SEC on May 8, 2017 (File No. 000-29089); and

the description of our common stock contained in our registration statement on Form 8-A filed under the Exchange Act on January 24, 2000, including any amendment or reports filed for the purpose of updating such descriptions (File No. 000-29089).

All filings by Agenus pursuant to the Exchange Act subsequent to the date hereof and prior to effectiveness of this registration statement are incorporated in this registration statement and deemed to be a part hereof from the date of filing of such documents or reports. In addition, all documents and reports filed by Agenus subsequent to the date hereof pursuant to Sections 13(a), 13(c), 14 and 15(d) of the Exchange Act prior to the filing of a post-effective amendment which indicates that all securities offered have been sold or which deregisters all securities remaining unsold, shall be deemed to be incorporated by reference in this registration statement and to be a part hereof from the date of filing of such documents or reports. Any statement contained in a document incorporated by reference herein shall be deemed to be modified or superseded to the extent that a statement contained herein or in any other subsequently filed document which also is incorporated by reference herein modifies or supersedes such statement. Statements in this prospectus concerning any document we filed as an exhibit to the registration statement or that we otherwise filed with the SEC are not intended to be comprehensive and are qualified by reference to these filings. You should review the complete document to evaluate these statements.

We will provide each person to whom this prospectus is delivered a copy of all of the information that has been incorporated by reference in this prospectus but not delivered with this prospectus. You may obtain copies of these

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filings, at no cost, through the Investor section of our website (www.agenusbio.com), and you may request copies of these filings, at no cost, by writing or telephoning us at:

Agenus Inc.

Attention: Secretary

3 Forbes Road

Lexington, MA 02421

Telephone: (781) 674-4400

The information contained on our website is not a part of this prospectus.

Table of Contents

Up to 20,000,000 Shares

Common Stock

PROSPECTUS SUPPLEMENT

B. RILEY FBR

May 11, 2018