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

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**SCHEDULE 14A**

**PROXY STATEMENT PURSUANT TO SECTION 14(a) OF THE  
SECURITIES EXCHANGE ACT OF 1934**

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Filed by the Registrant                       Filed by a Party other than the Registrant

Check the appropriate box:

- Preliminary Proxy Statement
- Confidential, for Use of the Commission Only (as permitted by Rule 14a-6(e)(2))**
- Definitive Proxy Statement
- Definitive Additional Materials
- Soliciting Material Pursuant to § 240.14a-12

**Regulus Therapeutics Inc.**

(Exact name of registrant as specified in its charter)

(Name of Person(s) Filing Proxy Statement if Other Than the Registrant)

Payment of Filing Fee (Check the appropriate box)

- No fee required.
- Fee computed on table below per Exchange Act Rules 14a-6(i)(1) and 0-11.

1. Title of each class of securities to which transaction applies:

2. Aggregate number of securities to which transaction applies:

3. Per unit price or other underlying value of transaction computed pursuant to Exchange Act Rule 0-11 (set forth the amount on which the filing fee is calculated and state how it was determined):

4. Proposed maximum aggregate value of transaction:

5. Total fee paid:

- Fee paid previously with preliminary materials.
- Check box if any part of the fee is offset as provided by Exchange Act Rule 0-11(a)(2) and identify the filing for which the offsetting fee was paid previously. Identify the previous filing by registration statement number, or the Form or Schedule and the date of its filing.

1. Amount Previously Paid:

2. Form, Schedule or Registration Statement No.:

3. Filing Party:

4. Date Filed:

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**PRELIMINARY PROXY STATEMENT – SUBJECT TO COMPLETION**

**REGULUS THERAPEUTICS INC.**  
10614 Science Center Drive  
San Diego, CA 92121

**NOTICE OF SPECIAL MEETING OF STOCKHOLDERS**

**To Be Held on September 28, 2018**

Dear Stockholders:

You are cordially invited to a Special Meeting of Stockholders of Regulus Therapeutics Inc., a Delaware corporation. The meeting will be held at 9:00 a.m. local time on September 28, 2018 at the office of Regulus Therapeutics Inc. at 10614 Science Center Drive, San Diego, California 92121.

We are holding the meeting for the following purposes, as more fully described in the accompanying proxy statement:

1. To approve an amendment to our Amended and Restated Certificate of Incorporation to effect a reverse stock split of our common stock at a ratio in the range of 1-for-5 to 1-for-20, with such ratio to be determined in the discretion of our board of directors and with such reverse stock split to be effected at such time and date, if at all, as determined by our board of directors in its sole discretion (Proposal 1);
2. To authorize an adjournment of the meeting, if necessary, to solicit additional proxies if there are not sufficient votes in favor of Proposal 1; and
3. To transact any other business that may be properly brought before the meeting or any continuation, adjournment or postponement thereof.

All of our stockholders of record as of \_\_\_\_\_, 2018 are entitled to attend and vote at the meeting and at any adjournment or postponement thereof.

Our board of directors recommends that you vote **FOR** the reverse split proposal as provided in Proposal 1 and **FOR** the authorization to adjourn the meeting, if necessary, to solicit additional proxies if there are not sufficient votes in favor of Proposal 1, as provided in Proposal 2.

**Important Notice Regarding the Availability of Proxy Materials for the Special Meeting of Stockholders of Regulus Therapeutics Inc. to be Held on September 28, 2018 at 9:00 a.m. local time at the office of Regulus Therapeutics located at 10614 Science Center Drive, San Diego, CA 92121.**

**The proxy statement is available at [www.regulusrx.com](http://www.regulusrx.com).**

By Order of the Board of Directors

Sincerely,

Christopher Aker  
*Corporate Secretary*

San Diego, California  
, 2018

**You are cordially invited to attend the meeting in person. Whether or not you expect to attend the meeting, please vote by proxy pursuant to the instructions set forth herein as promptly as possible in order to ensure your representation at the meeting. Even if you have voted by proxy, you may still vote in person if you attend the meeting. Please note, however, that if your shares are held of record by a broker, bank or other nominee and you wish to vote at the meeting, you must obtain a proxy issued in your name from that record holder.**

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**PRELIMINARY PROXY STATEMENT – SUBJECT TO COMPLETION**

**REGULUS THERAPEUTICS INC.**

10614 Science Center Drive  
San Diego, CA 92121

**PROXY STATEMENT FOR THE  
SPECIAL MEETING OF STOCKHOLDERS  
TO BE HELD ON September 28, 2018**

Our board of directors is soliciting proxies for use at a special meeting of stockholders (the “Special Meeting”) to be held at 9:00 a.m. local time on September 28, 2018 at the office of Regulus Therapeutics Inc. at 10614 Science Center Drive, San Diego, CA 92121, or at any postponement or adjournment of the Special Meeting. Regulus Therapeutics Inc. is sometimes referred to herein as “we”, “us”, “our”, “Regulus Therapeutics” or the “Company.”

**QUESTIONS AND ANSWERS ABOUT THESE PROXY MATERIALS AND VOTING**

**Why did I receive these proxy materials?**

We have sent you this proxy statement and the enclosed proxy card because the board of directors (sometimes referred to as the “Board”) of Regulus Therapeutics is soliciting your proxy to vote at the Special Meeting, including any adjournments or postponements of the Special Meeting. Pursuant to rules adopted by the Securities and Exchange Commission (the “SEC”), we are also providing access to our proxy materials over the internet, which can be accessed at [www.investorvote.com/RGLS](http://www.investorvote.com/RGLS).

We intend to mail these proxy materials on or about September 6, 2018 to all stockholders of record entitled to vote at the Special Meeting.

**Who can vote at the Special Meeting?**

Only stockholders of record at the close of business on \_\_\_\_\_, 2018 (the “record date”) will be entitled to vote at the Special Meeting or any adjournment or postponement thereof. On the record date, there were \_\_\_\_\_ shares of common stock outstanding and entitled to vote.

*Stockholder of Record: Shares Registered in Your Name*

If on the record date your shares were registered directly in your name with Regulus Therapeutics’ transfer agent, Computershare Trust Company, N.A., then you are a stockholder of record. As a stockholder of record, you may vote in person at the meeting or vote by proxy. Whether or not you plan to attend the Special Meeting, we urge you to vote by proxy pursuant to the instructions set forth below to ensure your vote is counted.

*Beneficial Owner: Shares Registered in the Name of a Broker or Bank*

If on the record date your shares were held, not in your name, but rather in an account at a brokerage firm, bank, dealer, or other similar organization, then you are the beneficial owner of shares held in “street name” and these proxy materials are being forwarded to you by that organization. The organization holding your account is considered to be the stockholder of record for purposes of voting at the Special Meeting. As a beneficial owner, you have the right to direct your broker or other agent on how to vote the shares in your account. You are also invited to attend the Special Meeting. However, since you are not the stockholder of record, you may not vote your shares in person at the Special Meeting unless you request and obtain a valid proxy from your broker or other agent.

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**What am I voting on?**

Management is presenting two proposals for stockholder vote:

1. To amend the Company's Amended and Restated Certificate of Incorporation, to effect a reverse stock split (the "Reverse Stock Split") of the Company's common stock at a ratio of between 1-for-5 and 1-for-20, with such ratio to be determined at the sole discretion of the Board and with such Reverse Stock Split to be effected at such time and date, if at all, as determined by the Board in its sole discretion; and
2. To approve the adjournment of the Special Meeting in order to solicit additional proxies if there are insufficient votes at the time of the Special Meeting to approve the Proposal 1 described above (the "Adjournment Proposal").

For each proposal, you may vote "For" such proposal, vote "Against" such proposal or "Abstain" from voting on such proposal.

Our board of directors unanimously recommends a vote FOR both of the foregoing proposals.

**What if another matter is properly brought before the Special Meeting?**

Our board of directors knows of no other matters that will be presented for consideration at the Special Meeting. If any other matters are properly brought before the Special Meeting, it is the intention of the persons named in the proxy to vote on those matters in accordance with their best judgment.

**How do I vote?**

The procedures for voting are fairly simple:

*Stockholder of Record: Shares Registered in Your Name*

If you are a stockholder of record, you may vote in person at the Special Meeting or vote by proxy using the enclosed proxy card, vote by proxy over the telephone or vote by proxy through the internet. Whether or not you plan to attend the Special Meeting, we urge you to vote by proxy to ensure your vote is counted. You may still attend the Special Meeting and vote in person if you have already voted by proxy.

- To vote in person, come to the Special Meeting and we will give you a ballot when you arrive.
- To vote using the proxy card, simply complete, sign and date the enclosed proxy card and return it promptly in the envelope provided. Your signed proxy card must be received by 11:59 p.m. Eastern Time on September 27, 2018.
- To vote over the telephone, dial toll-free 1-800-652-8683 using a touch-tone phone and follow the recorded instructions. You will be asked to provide the control number from the enclosed proxy card. Your vote must be received by 11:59 p.m. Eastern Time on September 27, 2018.
- To vote through the internet, go to <http://www.investorvote.com/RGLS> to complete an electronic proxy card. You will be asked to provide the control number from the enclosed proxy card. Your vote must be received by 11:59 p.m. Eastern Time on September 27, 2018.

We provide internet proxy voting to allow you to vote your shares online, with procedures designed to ensure the authenticity and correctness of your proxy vote instructions. However, please be aware that you must bear any costs associated with your internet access, such as usage charges from internet access providers and telephone companies.

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*Beneficial Owner: Shares Registered in the Name of Broker or Bank*

If you are a beneficial owner of shares registered in the name of your broker, bank, or other agent, you should have received a proxy card and voting instructions with this proxy statement from that organization rather than from Regulus Therapeutics. Simply complete and mail the proxy card as directed by the voting instructions to ensure that your vote is counted. Alternatively, you may be able to vote by telephone or over the Internet as instructed by your broker or bank. To vote in person at the Special Meeting, you must obtain a valid proxy from your broker, bank, or other agent. Follow the instructions from your broker or bank included with our proxy materials, or contact your broker or bank to request a proxy form.

**How many votes do I have?**

On each matter to be voted upon, you have one vote for each share of common stock you owned as of the record date.

**What if I return a proxy card or otherwise vote by proxy but do not make specific choices?**

If you voted by proxy without marking any voting selections, then the proxy holders will vote your shares as recommended by our board of directors on all matters presented in this Proxy Statement, and as the proxy holders may determine in their discretion with respect to any other matters properly presented for a vote at the Special Meeting.

**Who is paying for this proxy solicitation?**

We will pay for the entire cost of soliciting proxies. In addition to these proxy materials, our directors and employees may also solicit proxies in person, by telephone, or by other means of communication. Directors and employees will not be paid any additional compensation for soliciting proxies. We will also reimburse brokerage firms, banks and other agents for the cost of forwarding proxy materials to beneficial owners.

**What does it mean if I receive more than one proxy card?**

If you receive more than one proxy card, your shares are registered in more than one name or are registered in different accounts. Please follow the voting instructions with respect to **each** proxy card to ensure that all of your shares are voted.

**Can I change my vote after submitting my proxy?**

Yes. You can revoke your proxy at any time before the final vote at the Special Meeting. If you are a stockholder of record, you may revoke your proxy in any one of the following ways:

- You may send a written notice that you are revoking your proxy to Regulus Therapeutics' Corporate Secretary at 10614 Science Center Drive, San Diego, California 92121.
- You may grant another proxy by telephone or through the internet.
- You may submit another properly completed proxy card with a later date.
- You may attend the Special Meeting and vote in person. Simply attending the Special Meeting will not, by itself, revoke your proxy.

Your most current proxy, whether submitted by proxy card, telephone or internet, is the one that is counted.

If your shares are held by your broker or bank as a nominee or agent, you should follow the instructions provided by your broker or bank.

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**How are votes counted?**

Votes will be counted by the inspector of election appointed for the meeting, who will separately count “For” and “Against” votes, abstentions and broker non-votes. Abstentions will be counted towards the vote total for each proposal, and will have the same effect as “Against” votes. Broker non-votes will also have the same effect as “Against” votes for Proposal 1, but will not be counted towards the vote total for Proposal 2.

**What are “broker non-votes”?**

Broker non-votes occur when a beneficial owner of shares held in street name does not give instructions to the broker or nominee holding the shares as to how to vote on matters deemed “non-routine.” Generally, if shares are held in street name (shares are held by your broker as your nominee), the beneficial owner of the shares is entitled to give voting instructions to the broker or nominee holding the shares. If you do not give instructions to your broker, your broker can vote your shares with respect to matters that are considered to be “routine,” but not with respect to “non-routine” matters. Under the rules and interpretations of the New York Stock Exchange, “non-routine” matters are generally those involving a contest or a matter that may substantially affect the rights or privileges of stockholders. We believe Reverse Split Proposal and the Adjournment Proposal are routine matters. However, it is possible that brokers will not have discretionary authority with respect to the Reverse Split Proposal, in which case, if you do not instruct your broker how to vote with respect to the Reverse Stock Split Proposal, your broker may not vote with respect to such proposal.

**How many votes are needed to approve the Reverse Stock Split?**

A majority of the shares outstanding on the record date must vote “FOR” approval. Abstentions and broker non-votes will have the same effect as “AGAINST” votes.

**How many votes are needed to approve the Adjournment Proposal?**

Approval of the Adjournment Proposal requires the affirmative vote of a majority of the shares of our common stock present or represented by proxy at the Special Meeting and entitled to vote on such matter. Abstentions will have the same effect as “AGAINST” votes. Broker non-votes will have no effect on this proposal.

**What is the quorum requirement?**

A quorum of stockholders is necessary to hold a valid Special Meeting. A quorum will be present if at least a majority of the outstanding shares entitled to vote are represented by stockholders present at the Special Meeting or by proxy. On the record date, there were \_\_\_\_\_ shares outstanding and entitled to vote. Thus, \_\_\_\_\_ shares must be represented by stockholders present at the Special Meeting or by proxy to have a quorum.

Your shares will be counted towards the quorum only if you submit a valid proxy (or one is submitted on your behalf by your broker, bank or other nominee) or if you vote in person at the Special Meeting. Abstentions and broker non-votes will be counted towards the quorum requirement. If there is no quorum, the chairman of the Special Meeting or a majority of the votes present at the Special Meeting may adjourn the meeting to another date.

**How can I find out the results of the voting at the Special Meeting?**

Preliminary voting results will be announced at the Special Meeting. Final voting results will be published in a current report on Form 8-K that we expect to file no later than four business days after the conclusion of the Special Meeting. If final voting results are not available to us in time to file a Form 8-K on or before the fourth business day after the Special Meeting, we intend to file a Form 8-K to publish preliminary results and, within four business days after the final results are known to us, file an additional Form 8-K to publish the final results.

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**When are stockholder proposals due for the next annual meeting?**

To be considered for inclusion in the Company's proxy materials for next year's annual meeting of stockholders, your proposal must be submitted in writing by December 21, 2018 to the attention of the Secretary of Regulus Therapeutics Inc. at 10614 Science Center Drive, San Diego, California 92121. If you wish to submit a proposal (including a director nomination) at the meeting that is not to be included in the Company's proxy materials for next year's annual meeting, your written request must be received by the Secretary for Regulus Therapeutics Inc. at 10614 Science Center Drive, San Diego, California 92121 between February 1, 2019 and March 3, 2019. You are also advised to review the Company's Bylaws, which contain additional requirements about advance notice of stockholder proposals and director nominations.

**Whom should I contact with other questions?**

If you have additional questions about this Proxy Statement or the Special Meeting, or if you would like additional copies of this Proxy Statement, please contact: Regulus Therapeutics Inc., 10614 Science Center Drive, San Diego, California 92121, Attn: Corporate Secretary.

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## PROPOSAL 1

### BACKGROUND

Our Board of Directors has unanimously approved a series of alternate amendments to our Amended and Restated Certificate of Incorporation, which would:

- effect a reverse stock split, or Reverse Stock Split, of all issued and outstanding shares of our common stock, at a ratio ranging from 1-for-5 to 1-for-20, inclusive.

Accordingly, effecting a Reverse Stock Split would reduce the number of outstanding shares of our common stock. The effectiveness of any one of these amendments and the abandonment of the other amendments, or the abandonment of all of these amendments, will be determined by our Board of Directors following the Special Meeting and prior to the date of our 2019 Annual Meeting of Stockholders. Our Board of Directors has recommended that these proposed amendments be presented to our stockholders for approval.

Our stockholders are being asked to approve these proposed amendments pursuant to Proposal 1, and to grant authorization to our Board of Directors to determine, at its option, whether to implement a Reverse Stock Split, including its specific timing and ratio.

Should we receive the required stockholder approvals for Proposal 1, our Board of Directors will have the sole authority to elect, at any time on or prior to one-year anniversary of the Special Meeting, or September 27, 2019, and without the need for any further action on the part of our stockholders, whether to effect a Reverse Stock Split and the number of whole shares of our common stock, between and including five and twenty, that will be combined into one share of our common stock.

Notwithstanding approval of Proposal 1 by our stockholders, our Board of Directors may, at its sole option, abandon the proposed amendments and determine prior to the effectiveness of any filing with the Secretary of State of the State of Delaware not to effect any Reverse Stock Split, as permitted under Section 242(c) of the General Corporation Law of the State of Delaware. If our Board of Directors does not implement a Reverse Stock Split on or prior to the one-year anniversary of the Special Meeting, or September 28, 2019, stockholder approval would again be required prior to implementing any Reverse Stock Split.

**By approving Proposal 1, our stockholders will: (a) approve a series of alternate amendments to our Amended and Restated Certificate of Incorporation pursuant to which any whole number of outstanding shares of common stock between and including five (5) and twenty (20) could be combined into one share of common stock; and (b) authorize our Board of Directors to file only one such amendment, as determined by the Board at its sole option, and to abandon each amendment not selected by the Board. Our Board of Directors may also elect not to undertake any Reverse Stock Split and therefore abandon all amendments.**

### APPROVAL OF REVERSE STOCK SPLIT OF OUR COMMON STOCK (PROPOSAL 1)

Our Board of Directors has adopted and is recommending that our stockholders approve a series of alternate amendments to our Amended and Restated Certificate of Incorporation to effect a Reverse Stock Split. The text of the proposed form of Certificate of Amendment to our Amended and Restated Certificate of Incorporation, which we refer to as the Certificate of Amendment, is attached hereto as Appendix A.

We are proposing that our Board of Directors have the discretion to select the Reverse Stock Split ratio from within a range between and including 1-for-5 and 1-for-20, rather than proposing that stockholders approve a specific ratio at this time, in order to give our Board of Directors the flexibility to implement a Reverse Stock



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Split at a ratio that reflects the Board's then-current assessment of the factors described below under "Criteria to be Used for Determining Whether to Implement the Reverse Stock Split." If the Board decides to implement a Reverse Stock Split, we will file the Certificate of Amendment with the Secretary of State of the State of Delaware and the Reverse Stock Split will be effective when it is filed with the Secretary of State of the State of Delaware, or such later time as is chosen by the Board and set forth in the Certificate of Amendment. Except for adjustments that may result from the treatment of fractional shares as described below, each of our stockholders will hold the same percentage of our outstanding common stock immediately following the Reverse Stock Split as such stockholder holds immediately prior to the Reverse Stock Split.

### **Reasons for Reverse Stock Split**

*To maintain our listing on The Nasdaq Global Market.* By potentially increasing our stock price, the Reverse Stock Split would reduce the risk that our common stock could be delisted from The Nasdaq Global Market. To continue our listing on The Nasdaq Global Market, we must comply with Nasdaq Marketplace Rules, which requirements include a minimum bid price of \$1.00 per share. On April 23, 2018, we were notified by the Nasdaq Listing Qualifications Department that we do not comply with the \$1.00 minimum bid price requirement as our common stock had traded below the \$1.00 minimum bid price for 30 consecutive business days. We were automatically provided with a 180 calendar day period, ending on October 22, 2018, within which to regain compliance. To regain compliance, our common stock must close at or above the \$1.00 minimum bid price for at least 10 consecutive business days or more at the discretion of Nasdaq. If we do not regain compliance by October 22, 2018, it is possible that we would be eligible for an additional 180 calendar day compliance period. To qualify, we would need to meet, on October 22, 2018, the continued listing requirement for market value of publicly held shares and all other applicable standards for initial listing on The Nasdaq Capital Market, with the exception of the bid price requirement, and would need to provide written notice of our intention to cure the deficiency during the second compliance period by effecting a reverse stock split, if necessary. It is possible that we would not meet one or more of the standards required for the transfer of our common stock to the Nasdaq Capital Market on October 22, 2018, including the minimum stockholders' equity requirement. If we do not regain compliance by October 22, 2018 and are not granted a second 180-day compliance period, Nasdaq will notify us that our common stock will be subject to delisting. In that event, we may appeal the decision to a Nasdaq Listing Qualifications Panel. In the event of an appeal, our common stock would remain listed on The Nasdaq Global Market pending a written decision by the Panel following a hearing. In the event that the Nasdaq Listing Qualifications Panel determines not to continue our listing and we are delisted from The Nasdaq Global Market, our common stock may be delisted and trade on the OTC Bulletin Board or other small trading markets, such as the pink sheets.

The Board of Directors has considered the potential harm to us and our stockholders should Nasdaq delist our common stock from The Nasdaq Global Market. Delisting could adversely affect the liquidity of our common stock since alternatives, such as the OTC Bulletin Board and the pink sheets, are generally considered to be less efficient markets. An investor likely would find it less convenient to sell, or to obtain accurate quotations in seeking to buy, our common stock on an over-the-counter market. Many investors likely would not buy or sell our common stock due to difficulty in accessing over-the-counter markets, policies preventing them from trading in securities not listed on a national exchange or for other reasons. In addition, the delisting of our common stock from The Nasdaq Global Market would constitute an event of default under our Loan and Security Agreement with Oxford Finance, LLC.

The Board of Directors believes that the proposed Reverse Stock Split is a potentially effective means for us to maintain compliance with the \$1.00 minimum bid requirement and to avoid, or at least mitigate, the likely adverse consequences of our common stock being delisted from The Nasdaq Global Market by producing the immediate effect of increasing the bid price of our common stock.

*To potentially improve the marketability and liquidity of our common stock.* Our Board of Directors believes that the increased market price of our common stock expected as a result of implementing a Reverse Stock Split

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could improve the marketability and liquidity of our common stock and encourage interest and trading in our common stock.

- **Stock Price Requirements:** We understand that many brokerage houses, institutional investors and funds have internal policies and practices that either prohibit them from investing in low-priced stocks or tend to discourage individual brokers from recommending low-priced stocks to their customers or by restricting or limiting the ability to purchase such stocks on margin. Additionally, a Reverse Stock Split could help increase analyst and broker interest in our common stock as their internal policies might discourage them from following or recommending companies with low stock prices.
- **Stock Price Volatility:** Because of the trading volatility often associated with low-priced stocks, many brokerage houses and institutional investors have internal policies and practices that either prohibit them from investing in low-priced stocks or tend to discourage individual brokers from recommending low-priced stocks to their customers. Some of those policies and practices may make the processing of trades in low-priced stocks economically unattractive to brokers.
- **Transaction Costs:** Investors may be dissuaded from purchasing stocks below certain prices because brokers' commissions, as a percentage of the total transaction value, can be higher for low-priced stocks.

#### **Criteria to be Used for Determining Whether to Implement Reverse Stock Split**

In determining whether to implement the Reverse Stock Split and which Reverse Stock Split ratio to implement, if any, following receipt of stockholder approval of Proposal 1, our Board of Directors may consider, among other things, various factors, such as:

- the historical trading price and trading volume of our common stock;
- the then-prevailing trading price and trading volume of our common stock and the expected impact of the Reverse Stock Split on the trading market for our common stock in the short- and long-term;
- our ability to maintain our listing on The Nasdaq Global Market;
- which Reverse Stock Split ratio would result in the least administrative cost to us;
- prevailing general market and economic conditions; and
- whether and when our Board of Directors desires to have the additional authorized but unissued shares of common stock that will result from the implementation of a Reverse Stock Split available to provide the flexibility to use our common stock for business and/or financial purposes, as well as to accommodate the shares of our common stock to be authorized and reserved for future equity awards.

#### **Certain Risks and Potential Disadvantages Associated with Reverse Stock Split**

*We cannot assure you that the proposed Reverse Stock Split will increase our stock price and have the desired effect of maintaining compliance with Nasdaq Marketplace Rules.* We expect that the Reverse Stock Split will increase the market price of our common stock so that we may be able to regain and maintain compliance with the Nasdaq \$1.00 minimum bid price requirement. However, the effect of the Reverse Stock Split upon the market price of our common stock cannot be predicted with any certainty, and the history of similar reverse stock splits for companies in like circumstances is varied, particularly since some investors may view a reverse stock split negatively. It is possible that the per share price of our common stock after the Reverse Stock Split will not rise in proportion to the reduction in the number of shares of our common stock outstanding resulting from the Reverse Stock Split, and the market price per post-Reverse Stock Split share may not exceed or remain in excess of the \$1.00 minimum bid price for a sustained period of time, and the Reverse Stock Split may not result in a per share price that would attract brokers and investors who do not trade in lower priced stocks. In addition, although we believe the Reverse Stock Split may enhance the desirability of our common stock to certain potential

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investors, we cannot assure you that, if implemented, our common stock will be more attractive to institutional and other long term investors. Even if we implement the Reverse Stock Split, the market price of our common stock may decrease due to factors unrelated to the Reverse Stock Split. In any case, the market price of our common stock may also be based on other factors which may be unrelated to the number of shares outstanding, including our future performance. If the Reverse Stock Split is consummated and the trading price of the common stock declines, the percentage decline as an absolute number and as a percentage of our overall market capitalization may be greater than would occur in the absence of the Reverse Stock Split. Even if the market price per post-Reverse Stock Split share of our common stock remains in excess of \$1.00 per share, we may be delisted due to a failure to meet other continued listing requirements, including Nasdaq requirements related to the minimum number of shares that must be in the public float, the minimum market value of the public float and the minimum number of “round lot” holders.

*The proposed Reverse Stock Split may decrease the liquidity of our common stock and result in higher transaction costs.* The liquidity of our common stock may be negatively impacted by a Reverse Stock Split, given the reduced number of shares that would be outstanding after the Reverse Stock Split, particularly if the stock price does not increase as a result of the Reverse Stock Split. In addition, if a Reverse Stock Split is implemented, it will increase the number of our stockholders who own “odd lots” of fewer than 100 shares of common stock. Brokerage commission and other costs of transactions in odd lots are generally higher than the costs of transactions of more than 100 shares of common stock. Accordingly, a Reverse Stock Split may not achieve the desired results of increasing marketability and liquidity of our common stock described above.

*The effective increase in the authorized number of shares of our common stock as a result of the Reverse Stock Split could have anti-takeover implications.* The implementation of a Reverse Stock Split will result in an effective increase in the authorized number of shares of our common stock, which could, under certain circumstances, have anti-takeover implications. The additional shares of common stock that would become available for issuance if this Proposal 1 is approved and a Reverse Stock Split is implemented could be used by us to oppose a hostile takeover attempt or to delay or prevent changes in control or our management. For example, without further stockholder approval, the Board could adopt a “poison pill” which would, under certain circumstances related to an acquisition of our securities that is not approved by the Board, give certain holders the right to acquire additional shares of our common stock at a low price. The Board also could strategically sell shares of common stock in a private transaction to purchasers who would oppose a takeover or favor the current Board. Although this Proposal 1 has been prompted by business and financial considerations and not by the threat of any hostile takeover attempt (nor is the Board currently aware of any such attempts directed at us), stockholders should be aware that approval of this Proposal 1 could facilitate future efforts by us to deter or prevent changes in control, including transactions in which the stockholders might otherwise receive a premium for their shares over then current market prices.

#### **Effects of Reverse Stock Split**

After the effective date of any Reverse Stock Split that our Board of Directors elects to implement, each stockholder will own a reduced number of shares of common stock. However, any Reverse Stock Split will affect all of our stockholders uniformly and will not affect any stockholder’s percentage ownership interests in Regulus Therapeutics, except to the extent that the Reverse Stock Split results in any of our stockholders owning a fractional share as described below. Voting rights and other rights and preferences of the holders of our common stock will not be affected by a Reverse Stock Split (other than as a result of the payment of cash in lieu of fractional shares). For example, a holder of 2% of the voting power of the outstanding shares of our common stock immediately prior to a Reverse Stock Split would continue to hold 2% (assuming there is no impact as a result of the payment of cash in lieu of issuing fractional shares) of the voting power of the outstanding shares of our common stock immediately after such Reverse Stock Split. The number of stockholders of record will not be affected by a Reverse Stock Split (except to the extent that any stockholder holds only a fractional share interest and receives cash for such interest after such Reverse Stock Split).

The principal effects of a Reverse Stock Split will be that:

- depending on the Reverse Stock Split ratio selected by the Board, each five to twenty shares of our common stock owned by a stockholder will be combined into one new share of our common stock;
- no fractional shares of common stock will be issued in connection with any Reverse Stock Split; instead, holders of common stock who would otherwise receive a fractional share of common stock pursuant to the Reverse Stock Split will receive cash in lieu of the fractional share as explained more fully below;
- the total number of authorized shares of our common stock will remain at 200,000,000, resulting in an effective increase in the authorized number of shares of our common stock;
- based upon the Reverse Stock Split ratio selected by the Board, proportionate adjustments will be made to the per share exercise price and/or the number of shares issuable upon the exercise or vesting of all then outstanding stock options, restricted stock units and warrants (if any), which will result in a proportional decrease in the number of shares of our common stock reserved for issuance upon exercise or vesting of such stock options, restricted stock units and warrants, and, in the case of stock options and warrants, a proportional increase in the exercise price of all such stock options and warrants; and
- the number of shares then reserved for issuance under our equity compensation plans will be reduced proportionately based upon the Reverse Stock Split ratio selected by the Board.

The following table contains approximate information, based on share information as of June 30, 2018, relating to our outstanding common stock based on the proposed Reverse Stock Split ratios (without giving effect to the treatment of fractional shares):

<u>Status</u>	<u>Number of Shares of Common Stock Authorized</u>	<u>Number of Shares of Common Stock Issued and Outstanding</u>	<u>Number of Shares of Common Stock Reserved for Future Issuance</u>	<u>Number of Shares of Common Stock Authorized but Unissued and Unreserved</u>
Pre-Reverse Stock Split	200,000,000	104,424,492	2,584,077	92,991,431
Post-Reverse Stock Split 1:5	200,000,000	20,884,898	516,815	178,598,287
Post-Reverse Stock Split 1:6	200,000,000	17,404,082	430,679	182,165,239
Post-Reverse Stock Split 1:7	200,000,000	14,917,784	369,153	184,713,063
Post-Reverse Stock Split 1:8	200,000,000	13,053,061	323,009	186,623,930
Post-Reverse Stock Split 1:9	200,000,000	11,602,721	287,119	188,110,160
Post-Reverse Stock Split 1:10	200,000,000	10,442,449	258,407	189,299,144
Post-Reverse Stock Split 1:11	200,000,000	9,493,135	234,916	190,271,949
Post-Reverse Stock Split 1:12	200,000,000	8,702,041	215,339	191,082,620
Post-Reverse Stock Split 1:13	200,000,000	8,032,653	198,775	191,768,572
Post-Reverse Stock Split 1:14	200,000,000	7,458,892	184,576	192,356,532
Post-Reverse Stock Split 1:15	200,000,000	6,961,632	172,271	192,866,097
Post-Reverse Stock Split 1:16	200,000,000	6,526,530	161,504	193,311,966
Post-Reverse Stock Split 1:17	200,000,000	6,142,617	152,004	193,705,379
Post-Reverse Stock Split 1:18	200,000,000	5,801,360	143,559	194,055,081
Post-Reverse Stock Split 1:19	200,000,000	5,496,025	136,004	194,367,971
Post-Reverse Stock Split 1:20	200,000,000	5,221,224	129,203	194,649,573

After the effective date of any Reverse Stock Split that our Board of Directors elects to implement, our common stock would have a new committee on uniform securities identification procedures, or CUSIP number, a number used to identify our common stock.

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Our common stock is currently registered under Section 12(b) of the Securities Exchange Act, and we are subject to the periodic reporting and other requirements of the Securities Exchange Act of 1934, as amended, or the Exchange Act. The implementation of any proposed Reverse Stock Split will not affect the registration of our common stock under the Exchange Act. Our common stock would continue to be listed on The Nasdaq Global Market under the symbol "RGLS" immediately following the Reverse Stock Split, although it is likely that Nasdaq would add the letter "D" to the end of the trading symbol for a period of twenty trading days after the effective date of the Reverse Stock Split to indicate that the Reverse Stock Split had occurred.

#### **Effective Date**

The proposed Reverse Stock Split would become effective at 5:00 p.m., Eastern time, on the date of filing of a Certificate of Amendment with the office of the Secretary of State of the State of Delaware, or such later date as is chosen by the Board and set forth in the Certificate of Amendment, which date we refer to in this Proposal 1 as the Effective Date. Except as explained below with respect to fractional shares, effective as of 5:00 p.m., Eastern time, on the Effective Date, shares of common stock issued and outstanding immediately prior thereto will be combined, automatically and without any action on the part of us or our stockholders, into a lesser number of new shares of our common stock in accordance with the Reverse Stock Split ratio determined by our Board of Directors within the limits set forth in this Proposal 1.

#### **Cash Payment In Lieu of Fractional Shares**

No fractional shares of common stock will be issued as a result of any Reverse Stock Split. Instead, in lieu of any fractional shares to which a stockholder of record would otherwise be entitled as a result of the Reverse Stock Split, Regulus Therapeutics will pay cash (without interest) equal to such fraction multiplied by the average of the closing sales prices of the common stock on The Nasdaq Global Market during regular trading hours for the five consecutive trading days immediately preceding the Effective Date (with such average closing sales prices being adjusted to give effect to the Reverse Stock Split). After the Reverse Stock Split, a stockholder otherwise entitled to a fractional interest will not have any voting, dividend or other rights with respect to such fractional interest except to receive payment as described above.

As of June 30, 2018, there were four stockholders of record of our common stock. Upon stockholder approval of this Proposal 1, if our Board of Directors elects to implement the proposed Reverse Stock Split, stockholders owning, prior to the Reverse Stock Split, less than the number of whole shares of common stock that will be combined into one share of common stock in the Reverse Stock Split would no longer be stockholders. For example, if a stockholder held 10 shares of common stock immediately prior to the Reverse Stock Split and the Reverse Stock Split ratio selected by the Board was 1-for-15, then such stockholder would cease to be a stockholder of Regulus Therapeutics following the Reverse Stock Split and would not have any voting, dividend or other rights except to receive payment for the fractional share as described above. Based on our stockholders of record as of June 30, 2018, and assuming a Reverse Stock Split ratio of 1-for-15, we expect that cashing out fractional stockholders would not reduce the number of stockholders of record. In addition, we do not intend for this transaction to be the first step in a series of plans or proposals of a "going private transaction" within the meaning of Rule 13e-3 of the Exchange Act.

#### **Record and Beneficial Stockholders**

If this Proposal 1 is approved by our stockholders and our Board of Directors elects to implement a Reverse Stock Split, stockholders of record holding all of their shares of our common stock electronically in book-entry form under the direct registration system for securities will be automatically exchanged by the exchange agent and will receive a transaction statement at their address of record indicating the number of new post-split shares of our common stock they hold after the Reverse Stock Split along with payment in lieu of any fractional shares. Non-registered stockholders holding common stock through a bank, broker or other nominee should note that such banks, brokers or other nominees may have different procedures for processing the Reverse Stock Split and

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making payment for fractional shares than those that would be put in place by us for registered stockholders. If you hold your shares with such a bank, broker or other nominee and if you have questions in this regard, you are encouraged to contact your nominee.

If this Proposal 1 is approved by our stockholders and our Board of Directors elects to implement a Reverse Stock Split, stockholders of record holding some or all of their shares in certificate form will receive a letter of transmittal from Regulus Therapeutics or its exchange agent, as soon as practicable after the effective date of the Reverse Stock Split. Our transfer agent is expected to act as “exchange agent” for the purpose of implementing the exchange of stock certificates. Holders of pre-Reverse Stock Split shares will be asked to surrender to the exchange agent certificates representing pre-Reverse Stock Split shares in exchange for post-Reverse Stock Split shares and payment in lieu of fractional shares (if any) in accordance with the procedures to be set forth in the letter of transmittal. No new post-Reverse Stock Split share certificates will be issued to a stockholder until such stockholder has surrendered such stockholder’s outstanding certificate(s) together with the properly completed and executed letter of transmittal to the exchange agent.

STOCKHOLDERS SHOULD NOT DESTROY ANY PRE-SPLIT STOCK CERTIFICATE AND SHOULD NOT SUBMIT ANY CERTIFICATES UNTIL THEY ARE REQUESTED TO DO SO.

#### **Accounting Consequences**

The par value per share of our common stock would remain unchanged at \$0.001 per share after any Reverse Stock Split. As a result, on the Effective Date, the stated capital on our balance sheet attributable to the common stock would be reduced proportionally, based on the actual Reverse Stock Split ratio, from its present amount, and the additional paid-in capital account would be credited with the amount by which the stated capital would be reduced. The net income or loss per share of common stock would be increased because there would be fewer shares of common stock outstanding. The Reverse Stock Split would be reflected retroactively in our consolidated financial statements. We do not anticipate that any other accounting consequences would arise as a result of any Reverse Stock Split.

#### **No Appraisal Rights**

Our stockholders are not entitled to dissenters’ or appraisal rights under the General Corporation Law of the State of Delaware with respect to the proposed alternate amendments to our Amended and Restated Certificate of Incorporation to allow for a Reverse Stock Split.

#### **Material Federal Income Tax Consequences**

The following is a summary of the material U.S. federal income tax consequences of a Reverse Stock Split to our stockholders. The summary is based on the Internal Revenue Code of 1986, as amended, or the Code, applicable Treasury Regulations promulgated thereunder, judicial authority and current administrative rulings and practices as in effect on the date of this proxy statement. Changes to the laws could alter the tax consequences described below, possibly with retroactive effect. We have not sought and will not seek an opinion of counsel or a ruling from the Internal Revenue Service regarding the federal income tax consequences of a Reverse Stock Split. This discussion is for general information only and does not discuss the tax consequences which may apply to special classes of taxpayers (e.g., non-resident aliens, broker/dealers or insurance companies). The state and local tax consequences of a Reverse Stock Split may vary significantly as to each stockholder, depending upon the jurisdiction in which such stockholder resides. Stockholders are urged to consult their own tax advisors to determine the particular consequences to them.

In general, the federal income tax consequences of a Reverse Stock Split will vary among stockholders depending upon whether they receive cash for fractional shares or solely a reduced number of shares of common stock in exchange for their old shares of common stock. We believe that because the Reverse Stock Split is not

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part of a plan to increase periodically a stockholder's proportionate interest in our assets or earnings and profits, the Reverse Stock Split should have the following federal income tax effects. A stockholder who receives solely a reduced number of shares of common stock will not recognize gain or loss. In the aggregate, such a stockholder's basis in the reduced number of shares of common stock will equal the stockholder's basis in its old shares of common stock and such stockholder's holding period in the reduced number of shares will include the holding period in its old shares exchanged. A stockholder who receives cash in lieu of a fractional share as a result of the Reverse Stock Split should generally be treated as having received the payment as a distribution in redemption of the fractional share, as provided in Section 302(a) of the Code. Generally, if redemption of the fractional shares of all stockholders reduces the percentage of the total voting power held by a particular redeemed stockholder (determined by including the voting power held by certain related persons), the particular stockholder should recognize gain or loss equal to the difference, if any, between the amount of cash received and the stockholder's basis in the fractional share. In the aggregate, such a stockholder's basis in the reduced number of shares of common stock will equal the stockholder's basis in its old shares of common stock decreased by the basis allocated to the fractional share for which such stockholder is entitled to receive cash, and the holding period of the reduced number of shares received will include the holding period of the old shares exchanged. If the redemption of the fractional shares of all stockholders leaves the particular redeemed stockholder with no reduction in the stockholder's percentage of total voting power (determined by including the voting power held by certain related persons), it is likely that cash received in lieu of a fractional share would be treated as a distribution under Section 301 of the Code. Stockholders should consult their own tax advisors regarding the tax consequences to them of a payment for fractional shares.

We will not recognize any gain or loss as a result of the proposed Reverse Stock Split.

**Required Vote**

Stockholder approval of this Proposal 1 requires a "FOR" vote from at least a majority of the outstanding shares of our common stock on the record date.

**THE BOARD OF DIRECTORS RECOMMENDS THAT STOCKHOLDERS VOTE "FOR"  
THE APPROVAL OF A REVERSE STOCK SPLIT AS SET FORTH IN PROPOSAL 1.**

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**AUTHORIZATION TO ADJOURN THE SPECIAL MEETING (PROPOSAL 2)**

**General**

If the Special Meeting is convened and a quorum is present, but there are not sufficient votes to approve Proposal 1, our proxy holders may move to adjourn the Special Meeting at that time in order to enable the our board of directors to solicit additional proxies.

In this proposal, we are asking our stockholders to authorize the holder of any proxy solicited by our board of directors to vote in favor of granting discretionary authority to the proxy holders, and each of them individually, to adjourn the Special Meeting to another time and place, if necessary, to solicit additional proxies in the event there are not sufficient votes to approve Proposal 1. If our stockholders approve this proposal, we could adjourn the Special Meeting and any adjourned session of the Special Meeting and use the additional time to solicit additional proxies, including the solicitation of proxies from our stockholders that have previously voted. Among other things, approval of this proposal could mean that, even if we had received proxies representing a sufficient number of votes to defeat Proposal 1, we could adjourn the Special Meeting without a vote on such proposal and seek to convince our stockholders to change their votes in favor of such proposal.

If it is necessary to adjourn the Special Meeting, no notice of the adjourned meeting is required to be given to our stockholders, other than an announcement at the Special Meeting of the time and place to which the Special Meeting is adjourned, so long as the meeting is adjourned for 30 days or less and no new record date is fixed for the adjourned meeting. At the adjourned meeting, we may transact any business which might have been transacted at the original meeting.

**Vote Required**

Approval of this proposal requires the affirmative vote of a majority of the shares of our common stock present or represented by proxy at the Special Meeting and entitled to vote on such matter.

**THE BOARD OF DIRECTORS RECOMMENDS THAT STOCKHOLDERS VOTE “FOR”  
THE AUTHORIZATION TO ADJOURN THE SPECIAL MEETING AS SET FORTH IN  
PROPOSAL 2.**



## SECURITY OWNERSHIP OF CERTAIN BENEFICIAL OWNERS AND MANAGEMENT

The following table sets forth the beneficial ownership of our common stock as of July 31, 2018 by:

- each person, entity or group known to the Company to be the beneficial owner of more than 5% of the Company's common stock;
- each of our directors;
- each of our named executive officers; and
- all current executive officers and directors of the Company as a group.

Unless otherwise indicated in the footnotes to the table below and subject to community property laws where applicable, we believe that each of the stockholders named in the table has sole voting and investment power with respect to the shares indicated as beneficially owned.

Applicable percentages are based on 104,797,517 shares of common stock outstanding as of July 31, 2018, adjusted as required by rules promulgated by the SEC. These rules generally attribute beneficial ownership of securities to persons who possess sole or shared voting power or investment power with respect to those securities. In addition, the rules include shares of common stock issuable pursuant to the exercise of stock options or warrants that are exercisable within 60 days following July 31, 2018. These shares are deemed to be outstanding and beneficially owned by the person holding those options for the purpose of computing the percentage ownership of that person, but they are not treated as outstanding for the purpose of computing the percentage ownership of any other person. Certain of the options in this table are exercisable at any time but, if exercised, are subject to a lapsing right of repurchase until the options are fully vested. Unless otherwise indicated, the address for each person or entity listed in the table is c/o Regulus Therapeutics Inc., 10614 Science Center Drive, San Diego, CA 92121.

Beneficial Owner	Beneficial Ownership	
	Number of Shares	Percent of Total
<b>Greater than 5% Stockholders</b>		
FMR LLC <sup>(1)</sup> 245 Summer Street Boston, Massachusetts 02210	15,592,272	14.9%
New Enterprise Associates <sup>(2)</sup> 1954 Greenspring Dr., Suite 600 Timonium, Maryland 21093	10,989,010	10.5%
BVF Partners, LP <sup>(3)</sup> 1 Sansome Street, 30 <sup>th</sup> Floor San Francisco, California 94104	6,140,322	5.9%
Aventisub LLC <sup>(4)</sup> c/o Sanofi 54, rue La Boétie 75414 Paris—France	8,706,526	8.3%
<b>Named Executive Officers and Directors</b>		
David Baltimore, Ph.D. <sup>(5)</sup>	426,207	*
Kathryn J. Collier <sup>(6)</sup>	45,234	*
Daniel R. Chevallard <sup>(7)</sup>	375,440	*
Mark Deeg, M.D., Ph.D. <sup>(8)</sup>	475,000	*
Paul C. Grint, M.D. <sup>(9)</sup>	—	—
Joseph P. Hagan. <sup>(10)</sup>	1,164,162	1.1%
Stelios Papadopoulos, Ph.D. <sup>(11)</sup>	1,908,684	1.8%
William H. Rastetter, Ph.D. <sup>(12)</sup>	847,426	*
Hugh Rosen, M.D., Ph.D. <sup>(13)</sup>	150,052	*
Timothy Wright, M.D. <sup>(14)</sup>	853,020	*
Pascale Witz, MBA, MSc <sup>(15)</sup>	94,087	*
All current executive officers and directors as a group (9 Persons) <sup>(16)</sup>	5,684,702	5.4%

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\* Less than 1%.

- (1) Consists of shares beneficially owned, or that may be deemed to be beneficially owned, by FMR LLC, certain of its subsidiaries and affiliates, and other companies (collectively, the “FMR Reporters”). Does not reflect securities, if any, beneficially owned by certain other companies whose beneficial ownership of securities is disaggregated from that of the FMR Reporters in accordance with Securities and Exchange Commission Release No. 34-39538 (January 12, 1998). Abigail P. Johnson is a Director, the Vice Chairman, the Chief Executive Officer and the President of FMR LLC. Members of the Johnson family, including Abigail P. Johnson, are the predominant owners, directly or through trusts, of Series B voting common shares of FMR LLC, representing 49% of the voting power of FMR LLC. The Johnson family group and all other Series B shareholders have entered into a shareholders’ voting agreement under which all Series B voting common shares will be voted in accordance with the majority vote of Series B voting common shares. Accordingly, through their ownership of voting common shares and the execution of the shareholders’ voting agreement, members of the Johnson family may be deemed, under the Investment Company Act of 1940, to form a controlling group with respect to FMR LLC. Neither FMR LLC nor Abigail P. Johnson has the sole power to vote or direct the voting of the shares owned directly by the various investment companies registered under the Investment Company Act (“Fidelity Funds”) advised by Fidelity Management & Research Company (“FMR Co”), a wholly owned subsidiary of FMR LLC, which power resides with the Fidelity Funds’ Boards of Trustees. Fidelity Management & Research Company carries out the voting of the shares under written guidelines established by the Fidelity Funds’ Boards of Trustees. On August 10, 2018, FMR, LLC filed an amended Schedule 13G indicating it had reduced its ownership in our stock to 8,537,762 shares, reducing FRM LLC’s percentage ownership to 8.1%.
- (2) Consists of shares beneficially owned, or that may be deemed to be beneficially owned by, (a) Growth Equity Opportunities Fund V, LLC (“GEO”); (b) New Enterprise Associates 16, L.P. (“NEA 16”), which is the sole member of GEO, NEA Partners 16, L.P. (“NEA Partners 16”), which is the sole general partner of NEA 16; and NEA 16 GP, LLC (“NEA 16 LLC” and, together with NEA Partners 16, the “Control Entities”), which is the sole general partner of NEA Partners 16; and (c) Peter J. Barris (“Barris”), Forest Baskett (“Baskett”), Anthony A. Florence, Jr. (“Florence”), Mohamad H. Makhzoumi (“Makhzoumi”), Joshua Makower (“Makower”), David M. Mott (“Mott”), Chetan Puttagunta (“Puttagunta”), Jon M. Sakoda (“Sakoda”), Scott D. Sandell (“Sandell”), Peter W. Sonsini (“Sonsini”) and Ravi Viswanathan (“Viswanathan”) (together, the “Managers”). The Managers are the managers of NEA 16 LLC. The persons named herein are referred to individually herein as a “Reporting Person” and collectively as the “Reporting Persons.” GEO is the record owner of the GEO Shares. As the sole member of GEO, NEA 16 may be deemed to own beneficially the GEO Shares. As the general partner of NEA 16, NEA Partners 16 may be deemed to own beneficially the GEO Shares. As the sole general partner of NEA Partners 16, NEA 16 LLC may be deemed to own beneficially the GEO Shares. As members of NEA 16 LLC, each of the Managers may be deemed to own beneficially the GEO Shares. Each Reporting Person disclaims beneficial ownership of the GEO Shares other than those shares which such person owns of record.
- (3) Consists of shares beneficially owned, or that may be deemed to be beneficially owned, by BVF Inc., as the general partner of BVF, BVF2, and Trading Fund OS. BVF Partners, L.P., as the general partner of BVF, BVF2, the investment manager of Trading Fund OS, and the sole member of Partners OS, may be deemed to beneficially own the 13,974,249 shares of Common Stock beneficially owned in the aggregate by BVF, BVF2, Trading Fund OS, and certain BVF Partners, L.P. managed accounts (the “Partners Managed Accounts”), including 2,865,734 shares of Common Stock held in the Partners Managed Accounts. Mr. Mark M. Lampert, as a director and officer of BVF Inc., may be deemed to beneficially own the 13,974,249 Common Stock beneficially owned by BVF Inc. The foregoing should not be construed in and of itself as an admission by any Reporting Person as to beneficial ownership of any Common Stock owned by another Reporting Person. Partners OS disclaims beneficial ownership of the Common Stock beneficially owned by Trading Fund OS. Each of Partners, BVF Inc. and Mr. Lampert disclaims beneficial ownership of the Common Stock beneficially owned by BVF, BVF2, Trading Fund OS, and the Partners Managed

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Accounts, and the filing of this statement shall not be construed as an admission that any such person or entity is the beneficial owner of any such securities.

- (4) Aventisub LLC is a subsidiary of Sanofi. Sanofi has the ability to exercise voting and dispositive power over the shares held by Aventisub LLC.
- (5) Includes 392,870 shares that Dr. Baltimore has the right to acquire from us within 60 days of July 31, 2018 pursuant to the exercise of stock options.
- (6) Includes 26,250 shares that Ms. Collier has the right to acquire from us within 60 days of July 31, 2018, pursuant to the exercise of stock options.
- (7) Includes 313,920 shares that Mr. Chevallard has the right to acquire from us within 60 days of July 31, 2018 pursuant to the exercise of stock options.
- (8) Includes 475,000 shares that Dr. Deeg has the right to acquire from us within 60 days of July 31, 2018 pursuant to the exercise of stock options. Dr. Deeg's employment ended on July 13, 2018 and his options will cancel on October 13, 2018 unless exercised.
- (9) Dr. Grint's employment ended on May 4, 2017.
- (10) Includes 912,865 shares that Mr. Hagan has the right to acquire from us within 60 days of July 31, 2018.
- (11) Includes 159,693 shares that Dr. Papadopoulos has the right to acquire from us within 60 days of July 31, 2018 pursuant to the exercise of stock options.
- (12) Includes 187,870 shares that Dr. Rastetter has the right to acquire from us within 60 days of July 31, 2018 pursuant to the exercise of stock options.
- (13) Includes 121,963 shares that Dr. Rosen has the right to acquire from us within 60 days of July 31, 2018.
- (14) Includes 558,295 shares that Dr. Wright has the right to acquire from us within 60 days of July 31, 2018.
- (15) Includes 63,750 shares that Ms. Witz has the right to acquire from us within 60 days of July 31, 2018.
- (16) Includes the shares described in notes (5) through (7) and (10) through (15).

#### **STOCKHOLDERS SHARING THE SAME ADDRESS**

SEC rules permit companies, brokers, banks or other agents to deliver a single copy of a proxy statement to households at which two or more stockholders reside. This practice, known as "householding," is designed to reduce duplicate mailings and save significant printing and postage costs as well as natural resources. Stockholders sharing an address who have been previously notified by their broker, bank or other agent and have consented to householding will receive only one copy of our proxy statement.

If you would like to opt out of this practice for future mailings and receive separate proxy statements and, if applicable, annual reports for each stockholder sharing the same address, please contact your broker, bank or other agent. You may also obtain a separate proxy statement without charge by contacting us at Regulus Therapeutics Inc., Attn: Investor Relations, 10614 Science Center Drive, San Diego, California 92121, or contact our Corporate Secretary by telephone at (858) 202-6300. We will promptly send additional copies of the proxy statement.

Stockholders sharing an address that are receiving multiple copies of the proxy statement can request delivery of a single copy of the proxy statement by contacting their broker, bank or other intermediary or by contacting us as indicated above.

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**OTHER MATTERS**

We do not know of any business other than that described in this Proxy Statement that will be presented for consideration or action by the stockholders at the Special Meeting. If, however, any other business is properly brought before the Special Meeting, shares represented by proxies will be voted in accordance with the best judgment of the persons named in the proxies or their substitutes.

By Order of the Board of Directors

Christopher Aker  
*Vice President, Legal Affairs and Secretary*

San Diego, California  
, 2018

APPENDIX A

CERTIFICATE OF AMENDMENT OF

AMENDED AND RESTATED CERTIFICATE OF INCORPORATION  
OF  
REGULUS THERAPEUTICS INC.

**REGULUS THERAPEUTICS INC.**, a corporation organized and existing under the laws of the State of Delaware, hereby certifies as follows:

**FIRST:** The name of this corporation is Regulus Therapeutics Inc. (the "Company").

**SECOND:** The date on which the Company's Certificate of Incorporation was originally filed with the Secretary of State of the State of Delaware is January 2, 2009.

**THIRD:** The Board of Directors of the Company, acting in accordance with the provisions of Sections 141 and 242 of the General Corporation Law of the State of Delaware, adopted resolutions amending its Amended and Restated Certificate of Incorporation, as heretofore amended (the "Certificate of Incorporation"), as follows:<sup>1</sup>

Effective as of the effective time of 5:00 p.m., Eastern time, on the date this Certificate of Amendment is filed with the Secretary of State of the State of Delaware (the "Effective Time"), each [ five (5), six (6), seven (7), eight (8), nine (9), ten (10), eleven (11), twelve (12), thirteen (13), fourteen (14), fifteen (15), sixteen (16), seventeen (17), eighteen (18), nineteen (19), twenty (20)] shares of the Company's Common Stock, par value \$0.001 per share, issued and outstanding immediately prior to the Effective Time shall, automatically and without any action on the part of the Company or the respective holders thereof, be combined into one (1) share of Common Stock without increasing or decreasing the par value of each share of Common Stock (the "Reverse Split"); *provided, however*, no fractional shares of Common Stock shall be issued as a result of the Reverse Split and, in lieu thereof, upon receipt after the Effective Time by the exchange agent selected by the Company of a properly completed and duly executed transmittal letter and, where shares are held in certificated form, the surrender of the stock certificate(s) formerly representing shares of pre-Reverse Split Common Stock, any stockholder who would otherwise be entitled to a fractional share of post-Reverse Split Common Stock as a result of the Reverse Split, following the Effective Time (after taking into account all fractional shares of post-Reverse Split Common Stock otherwise issuable to such stockholder), shall be entitled to receive a cash payment (without interest) equal to the fractional share of post-Reverse Split Common Stock to which such stockholder would otherwise be entitled multiplied by the average of the closing sales prices of a share of the Company's Common Stock (as adjusted to give effect to the Reverse Split) on The Nasdaq Global Market during regular trading hours for the five (5) consecutive trading days immediately preceding the date this Certificate of Amendment is filed with the Secretary of State of the State of Delaware. Each stock certificate that, immediately prior to the Effective Time, represented shares of pre-Reverse Split Common Stock shall, from and after the Effective Time, automatically and without any action on the part of the Company or the respective holders thereof, represent

<sup>1</sup> These amendments approve the combination of any whole number of shares of Regulus's common stock between and including five (5) and twenty (20) into one (1) share of Regulus's common stock. By these amendments, the stockholders would approve each of the alternate amendments proposed by Regulus's Board of Directors. If the reverse stock split proposal is approved by stockholders, the Certificate of Amendment filed with the Secretary of State of the State of Delaware will include only that reverse stock split ratio determined by Regulus's Board of Directors to be in the best interests of Regulus and its stockholders. The other amendments will be abandoned pursuant to Section 242(c) of the General Corporation Law of the State of Delaware. Regulus's Board of Directors may also elect not to effect any reverse stock split, in which case all proposed alternate amendments will be abandoned.

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that number of whole shares of post-Reverse Split Common Stock into which the shares of pre-Reverse Split Common Stock represented by such certificate shall have been combined (as well as the right to receive cash in lieu of any fractional shares of post-Reverse Split Common Stock as set forth above; provided, however, that each holder of record of a certificate that represented shares of pre-Reverse Split Common Stock shall receive, upon surrender of such certificate, a new certificate representing the number of whole shares of post-Reverse Split Common Stock into which the shares of pre-Reverse Split Common Stock represented by such certificate shall have been combined pursuant to the Reverse Split, as well as any cash in lieu of fractional shares of post-Reverse Split Common Stock to which such holder may be entitled as set forth above. The Reverse Split shall be effected on a record holder-by-record holder basis, such that any fractional shares of post-Reverse Split Common Stock resulting from the Reverse Split and held by a single record holder shall be aggregated.

**FOURTH:** The foregoing amendment was submitted to the stockholders of the Company for their approval, and was duly adopted in accordance with the provisions of Section 242 of the General Corporation Law of the State of Delaware and shall be effective as of 5:00 p.m., Eastern time, on the date this Certificate of Amendment is filed with the Secretary of State of the State of Delaware.

**IN WITNESS WHEREOF, REGULUS THERAPEUTICS INC.** has caused this Certificate of Amendment to be signed by its President and Chief Executive Officer this     day of     , 2018.

**REGULUS THERAPEUTICS INC.**

By: \_\_\_\_\_  
Joseph Hagan  
President and Chief Executive Officer



IMPORTANT SPECIAL MEETING INFORMATION

Electronic Voting Instructions

Available 24 hours a day, 7 days a week!

Instead of mailing your proxy, you may choose one of the voting methods outlined below to vote your proxy.

VALIDATION DETAILS ARE LOCATED BELOW IN THE TITLE BAR.

Proxies submitted by the Internet or telephone must be received by 11:59 p.m. Eastern, on September 27, 2018.

Vote by Internet

- Go to www.investorvote.com/RGLS
• Or scan the QR code with your smartphone
• Follow the steps outlined on the secure website

Vote by telephone

- Call toll free 1-800-652-VOTE (8683) within the USA, US territories & Canada on a touch tone telephone
• Follow the instructions provided by the recorded message

Using a black ink pen, mark your votes with an X as shown in this example. Please do not write outside the designated areas.



Special Meeting Proxy Card

IF YOU HAVE NOT VOTED VIA THE INTERNET OR TELEPHONE, FOLD ALONG THE PERFORATION, DETACH AND RETURN THE BOTTOM PORTION IN THE ENCLOSED ENVELOPE.

A Proposals — The Board of Directors recommends a vote FOR Proposal 1 and FOR Proposal 2.

1. To approve an amendment to the Company's Amended and Restated Certificate of Incorporation to effect a reverse stock split of its common stock at a ratio in the range of 1-for-5 to 1-for-20, with such ratio to be determined in the discretion of Board and with such reverse stock split to be effected at such time and date, if at all, as determined by Board in its sole discretion

For Against Abstain
[ ] [ ] [ ]



2. To authorize an adjournment of the meeting, if necessary, to solicit additional proxies if there are not sufficient votes in favor of Proposal 1.

For Against Abstain
[ ] [ ] [ ]

B Authorized Signatures — This section must be completed for your vote to be counted. — Date and Sign Below

Please sign exactly as name(s) appears hereon. Joint owners should each sign. When signing as attorney, executor, administrator, corporate officer, trustee, guardian, or custodian, please give full title.

Date (mm/dd/yyyy) — Please print date below.

Signature 1 — Please keep signature within the box.

Signature 2 — Please keep signature within the box.

[ / / ]

[ ]

[ ]

IF VOTING BY MAIL, YOU MUST COMPLETE SECTIONS A - C ON BOTH SIDES OF THIS CARD.



1 U P X



**REGULUS THERAPEUTICS INC.**

Special Meeting of  
Regulus Therapeutics Inc. Stockholders  
September 28, 2018 9:00 A.M. PDT

**Important Notice Regarding the Availability of Proxy Materials for the Special Meeting:**  
The Proxy Statement is available at [www.edocumentview.com/RGLS](http://www.edocumentview.com/RGLS).

▼ IF YOU HAVE NOT VOTED VIA THE INTERNET OR TELEPHONE, FOLD ALONG THE PERFORATION, DETACH AND RETURN THE BOTTOM PORTION IN THE ENCLOSED ENVELOPE. ▼



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**Proxy – Regulus Therapeutics Inc.**

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**Notice of Special Meeting of Stockholders**

**Proxy Solicited by Board of Directors for Special Meeting – September 28, 2018**

The undersigned hereby appoint(s) Joseph P. Hagan and Christopher Aker and each of them, with power to act without the other and with power of substitution, as proxies and attorneys-in-fact and hereby authorizes them to represent and vote, as provided on the other side, all the shares of Regulus Therapeutics Inc. Common Stock which the undersigned is entitled to vote and, in their discretion, to vote upon such other business as may properly come before the Special Meeting of Stockholders of the Company to be held September 28, 2018 at the company's offices located at 10614 Science Center Drive, San Diego, CA 92121, with all powers which the undersigned would possess if present at the Meeting.

This proxy, when properly executed, will be voted in the manner directed herein. If no direction is made but the card is signed, this proxy card will be voted FOR Proposal 1 and FOR for Proposal 2, and in the discretion of the proxies with respect to such other business as may properly come before the meeting.

Continued and to be signed on reverse side

**C Non-Voting Items**

**Change of Address** — Please print your new address below.

**Comments** — Please print your comments below.

**Meeting Attendance**

Mark the box to the right if you plan to attend the Special Meeting.



IF VOTING BY MAIL, YOU MUST COMPLETE SECTIONS A - C ON BOTH SIDES OF THIS CARD.

