

| OMB APPROVAL                                 |           |
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**STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP**

Check this box if no longer subject to Section 16. Form 4 or Form 5 obligations may continue. See Instruction 1(b).

Filed pursuant to Section 16(a) of the Securities Exchange Act of 1934 or Section 30(h) of the Investment Company Act of 1940

|  |  |  |
|--|--|--|
| 1. Name and Address of Reporting Person*<br><b>ALTSHULER DAVID</b><br><br>(Last) (First) (Middle)<br><b>C/O VERTEX PHARMACEUTICALS INCORPORATED</b><br><b>50 NORTHERN AVENUE</b><br><br>(Street)<br><b>BOSTON MA 02210</b><br><br>(City) (State) (Zip) | 2. Issuer Name and Ticker or Trading Symbol<br><b>VERTEX PHARMACEUTICALS INC / MA [ VRTX ]</b> | 5. Relationship of Reporting Person(s) to Issuer (Check all applicable)<br><br>Director 10% Owner<br><input checked="" type="checkbox"/> Officer (give title below) Other (specify below)<br><b>EVP, Global Research and CSO</b> |
|  | 3. Date of Earliest Transaction (Month/Day/Year)<br><b>11/07/2022</b>                          |  |
|  |  | 6. Individual or Joint/Group Filing (Check Applicable Line)<br><br><input checked="" type="checkbox"/> Form filed by One Reporting Person<br><input type="checkbox"/> Form filed by More than One Reporting Person               |

**Table I - Non-Derivative Securities Acquired, Disposed of, or Beneficially Owned**

| 1. Title of Security (Instr. 3) | 2. Transaction Date (Month/Day/Year) | 2A. Deemed Execution Date, if any (Month/Day/Year) | 3. Transaction Code (Instr. 8) |   | 4. Securities Acquired (A) or Disposed Of (D) (Instr. 3, 4 and 5) |            |                            | 5. Amount of Securities Beneficially Owned Following Reported Transaction(s) (Instr. 3 and 4) | 6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4) | 7. Nature of Indirect Beneficial Ownership (Instr. 4) |
|---------------------------------|--------------------------------------|--|--------------------------------|---|---|------------|----------------------------|---|--|---|
|                                 |                                      |  | Code                           | V | Amount  | (A) or (D) | Price                      |   |  |   |
| Common Stock                    | 11/07/2022                           |  | M                              |   | 1,303   | A          | \$187.53                   | 37,380  | D  |   |
| Common Stock                    | 11/07/2022                           |  | S <sup>(1)</sup>               |   | 140   | D          | \$299.72 <sup>(2)(3)</sup> | 37,240  | D  |   |
| Common Stock                    | 11/07/2022                           |  | S <sup>(1)</sup>               |   | 210   | D          | \$300.96 <sup>(2)(4)</sup> | 37,030  | D  |   |
| Common Stock                    | 11/07/2022                           |  | S <sup>(1)</sup>               |   | 189   | D          | \$301.99 <sup>(2)(5)</sup> | 36,841  | D  |   |
| Common Stock                    | 11/07/2022                           |  | S <sup>(1)</sup>               |   | 261   | D          | \$303.21 <sup>(2)(6)</sup> | 36,580  | D  |   |
| Common Stock                    | 11/07/2022                           |  | S <sup>(1)</sup>               |   | 280   | D          | \$304.58 <sup>(2)(7)</sup> | 36,300  | D  |   |
| Common Stock                    | 11/07/2022                           |  | S <sup>(1)</sup>               |   | 140   | D          | \$305.39 <sup>(2)(8)</sup> | 36,160  | D  |   |
| Common Stock                    | 11/07/2022                           |  | S <sup>(1)</sup>               |   | 83  | D          | \$306.57 <sup>(2)(9)</sup> | 36,077  | D  |   |

**Table II - Derivative Securities Acquired, Disposed of, or Beneficially Owned (e.g., puts, calls, warrants, options, convertible securities)**

| 1. Title of Derivative Security (Instr. 3) | 2. Conversion or Exercise Price of Derivative Security | 3. Transaction Date (Month/Day/Year) | 3A. Deemed Execution Date, if any (Month/Day/Year) | 4. Transaction Code (Instr. 8) |   | 5. Number of Derivative Securities Acquired (A) or Disposed of (D) (Instr. 3, 4 and 5) |       | 6. Date Exercisable and Expiration Date (Month/Day/Year) |                 | 7. Title and Amount of Securities Underlying Derivative Security (Instr. 3 and 4) | 8. Price of Derivative Security (Instr. 5) | 9. Number of derivative Securities Beneficially Owned Following Reported Transaction(s) (Instr. 4) | 10. Ownership Form: Direct (D) or Indirect (I) (Instr. 4) | 11. Nature of Indirect Beneficial Ownership (Instr. 4) |       |
|--|--|--------------------------------------|--|--------------------------------|---|--|-------|--|-----------------|---|--|--|---|--|-------|
|  |  |                                      |  | Code                           | V | (A)  | (D)   | Date Exercisable   | Expiration Date |   |  |  |   |  | Title |
| Stock Option (Right to Buy)                | \$187.53   | 11/07/2022                           |  | M                              |   |  | 1,303 | (10)   | 02/05/2029      | Common Stock  | 1,303                                      | \$0.00   | 1,304   | D  |       |

**Explanation of Responses:**

- Transaction made pursuant to Mr. Altshuler's company approved trading plan under Rule 10b5-1.
- Mr. Altshuler undertakes to provide (upon request by the SEC staff, the issuer or a security holder of the issuer) full information regarding the number of shares sold at each separate price.
- Open market sales reported on this line occurred at a weighted average price of \$299.72 (range \$299.46 to \$299.97).
- Open market sales reported on this line occurred at a weighted average price of \$300.96 (range \$300.54 to \$301.18).
- Open market sales reported on this line occurred at a weighted average price of \$301.99 (range \$301.84 to \$302.20).
- Open market sales reported on this line occurred at a weighted average price of \$303.21 (range \$302.89 to \$303.45).
- Open market sales reported on this line occurred at a weighted average price of \$304.58 (range \$304.14 to \$304.91).
- Open market sales reported on this line occurred at a weighted average price of \$305.39 (range \$305.38 to \$305.39).
- Open market sales reported on this line occurred at a weighted average price of \$306.57 (range \$306.45 to \$307.22).
- The option vests in 16 quarterly installments from 2/6/2019.

**Remarks:**

/s/ Christiana Stevenson,  
Attorney-in-Fact

11/09/2022

\*\* Signature of Reporting Person Date

Reminder: Report on a separate line for each class of securities beneficially owned directly or indirectly.

\* If the form is filed by more than one reporting person, see Instruction 4 (b)(v).

\*\* Intentional misstatements or omissions of facts constitute Federal Criminal Violations See 18 U.S.C. 1001 and 15 U.S.C. 78ff(a).

Note: File three copies of this Form, one of which must be manually signed. If space is insufficient, see Instruction 6 for procedure.

Persons who respond to the collection of information contained in this form are not required to respond unless the form displays a currently valid OMB Number.