FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

OMB APPROVAL

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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 Add | ress of Reporting F | erson* | 2. Issuer Name and Ticker or Trading Symbol Denali Therapeutics Inc. [DNLI] | 5. Relationship of Reporting Person(s) to Issuer (Check all applicable) Director 10% Owner Officer (give title X Other (specify | | | | | | | | |
| (Last) 245 SUMMER | (Last) (First) (Middle) 245 SUMMER STREET | | 3. Date of Earliest Transaction (Month/Day/Year) 12/12/2017 | below) A below) See Remark 1 | | | | | | | | |
| (Street) BOSTON | MA | 02210 | 4. If Amendment, Date of Original Filed (Month/Day/Year) | G. Individual or Joint/Group Filing (Check Applicable Line) X Form filed by One Reporting Person Form filed by More than One Reporting Person | | | | | | | | |
| (City) (State) (Zip) Table I - Non-Derivative Securities Acquired, Disposed of, or Beneficially Owned | | | | | | | | | | | | |

| (Last) (Hirst) (Middle) 245 SUMMER STREET | | | | 2/12/2017 | | | | | See Remark 1 | | | | |
|--|----------|-----------|---|-------------------------------|---|---------|--|--------|--|---|---|---|--|
| (Street) BOSTON MA 02210 (City) (State) (Zip) | | | 4. | If Amendment, Date of | Origina | I Filed | (Month/Day/Ye | Line | Individual or Joint/Group Filing (Check Applicable Line) X Form filed by One Reporting Person Form filed by More than One Reporting Person | | | | |
| | | | on-Derivati | ve Securities Ac | nuired | l Die | enosed of a | or Ben | eficially | Owned | | | |
| 1. Title of Security (Instr. 3) | | | 2. Transaction Date (Month/Day/Ye | 2A. Deemed Execution Date, | 3. Transaction Code (Instr. 8) | | 4. Securities Acquired (A) of Disposed Of (D) (Instr. 3, 4 | | A) or | 5. Amount of Securities Beneficially Owned Following | 6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4) | 7. Nature of Indirect Beneficial Ownership | |
| | | | | Code | v | Amount | (A) or (D) | Price | Reported Transaction(s) (Instr. 3 and 4) | | (Instr. 4) | | |
| Common Stock | | | 12/12/201 | 7 | С | | 2,188,869 | A | (1) | 2,841,896 | I | Impresa Fund III Limited Partnership | |
| Common Stock | | | 12/12/201 | .7 | С | | 45,035 | A | (1) | 2,886,931 | I | Impresa Fund III Limited Partnership | |
| Common Stock | | | 12/12/201 | 7 | С | | 112,590 | A | (1) | 2,999,521 | I | Impresa Fund III Limited Partnership | |
| Common Stock | · | | 12/12/201 | .7 | С | | 808,013 | A | (1) | 1,049,072 | I | F-Prime Capital Partners Healthcare Fund IV LP | |
| Common Stock | s. | | 12/12/201 | .7 | С | | 16,624 | A | (1) | 1,065,696 | I | F-Prime Capital Partners Healthcare Fund IV LP | |
| Common Stock | S | | 12/12/201 | .7 | С | | 41,561 | A | (1) | 1,107,257 | I | F-Prime Capital Partners Healthcare Fund IV LP | |
| Common Stock | S | | 12/12/201 | 7 | С | | 24,240 | A | (1) | 31,471 | I | F-Prime Capital Partners Healthcare Advisors Fund IV LP | |
| Common Stock | | 12/12/201 | .7 | С | | 498 | A | (1) | 31,969 | I | F-Prime Capital Partners Healthcare Advisors Fund IV LP | | |

| 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) | | | | | 5. Amount of Securities Beneficially Owned Following | 6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4) | 7. Nature of Indirect Beneficial Ownership | | |
| | | | Code | v | Amount | (A) or (D) | Price | Reported Transaction(s) (Instr. 3 and 4) | | (Instr. 4) | | |
| Common Stock | 12/12/2017 | | С | | 1,246 | A | (1) | 33,215 | I | F-Prime Capital Partners Healthcare Advisors Fund IV LP | | |
| Common Stock | | | | | | | | 750,001 | I | F-Prime Inc. | | |

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

| (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. Transa Code (8) | action (Instr. | Deri Sec Acq or D | umber of vative urities uired (A) visposed of (Instr. 3, 4 5) | 6. Date Exerc Expiration Day/\(\text{Month/Day/\}\) | ate | 7. Title and Amount of Securities Underlying Derivative Security (Instr. 3 and 4) | | 8. Price of Derivative Security (Instr. 5) | Securities Beneficially Owned Following Reported | 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 | Amount or Number of Shares | | Transaction(s) (Instr. 4) | | |
| Series A-1 Convertible Preferred Stock | (1) | 12/12/2017 | | С | | | 2,188,869 | (1) | (1) | Common Stock | 2,188,869 | (1) | 0 | I | Impresa Fund III Limited Partnership |
| Series A-1 Convertible Preferred Stock | (1) | 12/12/2017 | | С | | | 808,013 | (1) | (1) | Common Stock | 808,013 | (1) | 0 | I | F-Prime Capital Partners Healthcare Fund IV LP |
| Series A-1 Convertible Preferred Stock | (1) | 12/12/2017 | | С | | | 24,240 | (1) | (1) | Common Stock | 24,240 | (1) | 0 | I | F-Prime Capital Partners Healthcare Advisors Fund IV LP |
| Series A-2 Convertible Preferred Stock | (1) | 12/12/2017 | | С | | | 45,035 | (1) | (1) | Common Stock | 45,035 | (1) | 0 | I | Impresa Fund III Limited Partnership |
| Series A-2 Convertible Preferred Stock | (1) | 12/12/2017 | | С | | | 16,624 | (1) | (1) | Common Stock | 16,624 | (1) | 0 | I | F-Prime Capital Partners Healthcare Fund IV LP |
| Series A-2 Convertible Preferred Stock | (1) | 12/12/2017 | | С | | | 498 | (1) | (1) | Common Stock | 498 | (1) | 0 | I | F-Prime Capital Partners Healthcare Advisors Fund IV LP |
| Series B-1 Convertible Preferred Stock | (1) | 12/12/2017 | | С | | | 112,590 | (1) | (1) | Common Stock | 112,590 | (1) | 0 | I | Impresa Fund III Limited Partnership |
| Series B-1 Convertible Preferred Stock | (1) | 12/12/2017 | | С | | | 41,561 | (1) | (1) | Common Stock | 41,561 | (1) | 0 | I | F-Prime Capital Partners Healthcare Fund IV LP |
| Series B-1 Convertible Preferred Stock | (1) | 12/12/2017 | | С | | | 1,246 | (1) | (1) | Common Stock | 1,246 | (1) | 0 | I | F-Prime Capital Partners Healthcare Advisors Fund IV LP |

Explanation of Responses:

1. On December 12, 2017, in connection with the completion of the issuer's initial public offering, each share of Convertible Preferred Stock converted into one share of Common Stock.

Remarks

Remark 1: Abigail P. Johnson is a Director, the Chairman and the Chief Executive Officer 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. The address of Abigail P. Johnson is c/o FMR LLC, 245 Summer Street, Boston, MA 02110. Remark 2: The filing of this statement shall not be deemed to be an admission that, for purposes of Section 16 of the Securities Exchange Act of 1934 or otherwise, the undersigned are the beneficial owners of any securities reported herein. Remark 3: The general partner of F-Prime Capital Partners Healthcare Fund IV LP is F-Prime Capital Partners Healthcare Fund IV LP is F-Prime Capital Partners and its investment manager. Impress Management LLC, its general partner and its investment manager. Impress Management LLC, its general partner and investment manager. Impress Management LLC is owned, directly or indirectly, by various shareholders and employees of FMR LLC, including certain members of the Johnson family. F-Prime Inc. is a wholly-owned subsidiary of FMR LLC.

and on behalf of FMR LLC and its direct and indirect subsidiaries, and Abigail P. Johnson

** 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.

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