

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

| OMB APPROVAL                                 |           |
|--|-----------|
| OMB Number:                                  | 3235-0287 |
| Estimated average burden hours per response: | 0.5       |

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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><u>Watts Ryan J.</u><br><br>(Last) (First) (Middle)<br>C/O DENALI THERAPEUTICS INC.<br>161 OYSTER POINT BLVD.<br><br>(Street)<br>SOUTH SAN FRANCISCO CA 94080<br><br>(City) (State) (Zip) | 2. Issuer Name and Ticker or Trading Symbol<br><u>Denali Therapeutics Inc. [ DNL1 ]</u> | 5. Relationship of Reporting Person(s) to Issuer (Check all applicable)<br><input checked="" type="checkbox"/> Director 10% Owner<br><input checked="" type="checkbox"/> Officer (give title below) Other (specify below)<br><b>President and CEO</b> |
|   | 3. Date of Earliest Transaction (Month/Day/Year)<br>10/01/2021                          |   |
| 4. If Amendment, Date of Original Filed (Month/Day/Year)  |   |   |

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                    | 10/01/2021                           |  | S <sup>(1)</sup>               |   | 3,304   | D          | \$49.38 <sup>(2)</sup> | 2,155,063   | I  | See footnote <sup>(3)</sup>                           |
| Common Stock                    | 10/01/2021                           |  | S <sup>(1)</sup>               |   | 10,782  | D          | \$50.44 <sup>(4)</sup> | 2,144,281   | I  | See footnote <sup>(3)</sup>                           |
| Common Stock                    | 10/01/2021                           |  | S <sup>(1)</sup>               |   | 4,247   | D          | \$50.99 <sup>(5)</sup> | 2,140,034   | I  | See footnote <sup>(3)</sup>                           |
| Common Stock                    |                                      |  |                                |   |   |            |                        | 151,691 <sup>(6)</sup>  | 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 |   |  |  |   |  |

Explanation of Responses:

- The sales reported by the Reporting Person were effected pursuant to a Rule 10b5-1 trading plan.
- The sale price reported in column 4 of Table I represents the weighted average sale price of the shares ranging from \$48.85 to \$49.81 per share. Upon request by the Commission staff, the Issuer, or a security holder of the Issuer, the Reporting Person will provide full information regarding the number of shares sold at each separate price.
- The shares are held of record by the Watts Family 2015 Trust dated July 7, 2015, for which the Reporting Person serves as trustee.
- The sale price reported in column 4 of Table I represents the weighted average sale price of the shares ranging from \$49.90 to \$50.895 per share. Upon request by the Commission staff, the Issuer, or a security holder of the Issuer, the Reporting Person will provide full information regarding the number of shares sold at each separate price.
- The sale price reported in column 4 of Table I represents the weighted average sale price of the shares ranging from \$50.90 to \$51.14 per share. Upon request by the Commission staff, the Issuer, or a security holder of the Issuer, the Reporting Person will provide full information regarding the number of shares sold at each separate price.
- Includes 151,691 restricted stock units.

Remarks:

/s/ Tyler Nielsen, by power of attorney 10/05/2021

\*\* 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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