

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* <u>Tessier-Lavigne Marc</u> (Last) (First) (Middle) C/O DENALI THERAPEUTICS INC. 161 OYSTER POINT BLVD. (Street) SOUTH SAN FRANCISCO CA 94080 (City) (State) (Zip)	2. Issuer Name and Ticker or Trading Symbol <u>Denali Therapeutics Inc. [DNL1]</u>	5. Relationship of Reporting Person(s) to Issuer (Check all applicable) <input checked="" type="checkbox"/> Director 10% Owner Officer (give title below) Other (specify below)
	3. Date of Earliest Transaction (Month/Day/Year) <u>12/28/2021</u>	
	4. If Amendment, Date of Original Filed (Month/Day/Year)	6. Individual or Joint/Group Filing (Check Applicable Line) <input checked="" type="checkbox"/> Form filed by One Reporting Person 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	10/04/2021		G	V	300,000	D	\$0 ⁽¹⁾	2,244,370 ⁽²⁾	D	
Common Stock	10/04/2021		G	V	100,000	A	\$0 ⁽¹⁾	100,000	I	The Tessier-Lavigne/Hynes Irrevocable Trust 1 ⁽³⁾
Common Stock	12/28/2021		S		3,732	D	\$44.37 ⁽⁴⁾	96,268	I	The Tessier-Lavigne/Hynes Irrevocable Trust 1 ⁽³⁾
Common Stock	12/29/2021		S		10,700	D	\$43.88 ⁽⁵⁾	85,568	I	The Tessier-Lavigne/Hynes Irrevocable Trust 1 ⁽³⁾
Common Stock	12/30/2021		S		3,343	D	\$45.13 ⁽⁶⁾	82,225	I	The Tessier-Lavigne/Hynes Irrevocable Trust 1 ⁽³⁾
Common Stock	12/30/2021		S		3,377	D	\$45.59 ⁽⁷⁾	78,848	I	The Tessier-Lavigne/Hynes Irrevocable Trust 1 ⁽³⁾
Common Stock	10/04/2021		G	V	100,000	A	\$0 ⁽¹⁾	100,000	I	The Tessier-Lavigne/Hynes Irrevocable Trust 2 ⁽⁸⁾
Common Stock	12/28/2021		S		3,602	D	\$44.38 ⁽⁹⁾	96,398	I	The Tessier-Lavigne/Hynes Irrevocable Trust 2 ⁽⁸⁾
Common Stock	12/29/2021		S		10,700	D	\$43.87 ⁽¹⁰⁾	85,698	I	The Tessier-Lavigne/Hynes Irrevocable Trust 2 ⁽⁸⁾
Common Stock	12/30/2021		S		5,625	D	\$45.31 ⁽¹¹⁾	80,073	I	The Tessier-Lavigne/Hynes Irrevocable Trust 2 ⁽⁸⁾

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	12/30/2021		S		900	D	\$45.71 ⁽¹²⁾	79,173	I	The Tessier-Lavigne/Hynes Irrevocable Trust 2 ⁽⁸⁾
Common Stock	10/04/2021		G	V	100,000	A	\$0 ⁽¹⁾	100,000	I	The Tessier-Lavigne/Hynes Irrevocable Trust 3 ⁽¹³⁾
Common Stock	12/28/2021		S		3,646	D	\$44.37 ⁽¹⁴⁾	96,354	I	The Tessier-Lavigne/Hynes Irrevocable Trust 3 ⁽¹³⁾
Common Stock	12/29/2021		S		10,704	D	\$43.88 ⁽¹⁵⁾	85,650	I	The Tessier-Lavigne/Hynes Irrevocable Trust 3 ⁽¹³⁾
Common Stock	12/30/2021		S		6,002	D	\$45.31 ⁽¹⁶⁾	79,648	I	The Tessier-Lavigne/Hynes Irrevocable Trust 3 ⁽¹³⁾
Common Stock	12/30/2021		S		800	D	\$45.72 ⁽¹⁷⁾	78,848	I	The Tessier-Lavigne/Hynes Irrevocable Trust 3 ⁽¹³⁾

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:

- This transaction involved a gift of securities by the Reporting Person. Disposition price is not applicable as this transaction was a bona fide gift.
- Includes 2,074 restricted stock units.
- The shares are held of record by The Tessier-Lavigne/Hynes Irrevocable Trust 1, 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 purchased ranging from \$44.19 to \$44.74 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 purchased ranging from \$43.50 to \$44.07 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 purchased ranging from \$44.42 to \$45.41 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 purchased ranging from \$45.42 to \$45.81 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 Tessier-Lavigne/Hynes Irrevocable Trust 2, 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 purchased ranging from \$44.19 to \$44.64 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 purchased ranging from \$43.51 to \$44.07 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 purchased ranging from \$44.65 to \$45.63 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 purchased ranging from \$45.66 to \$45.77 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 Tessier-Lavigne/Hynes Irrevocable Trust 3, 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 purchased ranging from \$44.18 to \$44.63 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 purchased ranging from \$43.50 to \$44.09 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 purchased ranging from \$44.65 to \$45.62 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 purchased ranging from \$45.67 to \$45.80 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.

Remarks:

/s/ Tyler Nielsen, by power of attorney 12/30/2021

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.