(Last)

(First)

(Middle)

FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

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

Check this box if no longer subject to Section 16. Form 4 or Form 5 obligations may continue. See

Instruc	ction 1(b).			Fil					a) of the Secur Investment C			934					
		f Reporting Person							ker or Trading utics Inc			(Ch	Relationship leck all applic	cable)	Pers	. ,	
(Last)	,	=irst) N STREET, 3RE	(Middle)		06/	/16/2	2023		action (Month		. 04		Officer below)	(give title		Other (s below)	specify
(Street)		JY	10014		- 4. II	rAme	enament, L	ate c	of Original File	a (Montn/Da	ıy/ Year)	Lin	Form f	iled by One iled by More	Repo	rting Perso	n
, THE WIT	JKK I	V I	10014		-	مار	10h5-1	(c)	Transac	tion Indi	ication		Persor	1			
(City)	?)	State)	(Zip)			Che	ck this box t	o indi	cate that a trans	saction was m	nade pursuar			n or written pl	lan tha	at is intended	l to
		Ta	ble I - Nor	n-Deriv	vative	e Se	curities	Ac	quired, Di	sposed o	f, or Ber	neficial	ly Owned				
1. Title of Security (Instr. 3)		Date	Transaction ate Month/Day/Year)		2A. Deemed Execution Date, if any (Month/Day/Year)		3. Transaction Code (Instr 8)	n Dispose	rities Acquired (A) or ed Of (D) (Instr. 3, 4 and		d Securities Form Beneficially Owned Following Reported Transaction(s)		Form	: Direct Indirect str. 4)	7. Nature of Indirect Beneficial Ownership (Instr. 4)		
									uired, Disp	osed of,	or Bene	eficially	Owned	and 4)			
1. Title of Derivative Security (Instr. 3) 2. Conversion or Exercise (Instr. 3) Price of Derivative Security 3. Transaction Date (Month/Day/Year)		3A. Deemed Execution D if any (Month/Day)	l 4 Pate, 1	4. Transaction Code (Instr.		5. Number of Derivative		options, convertible 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		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 (Instr. 4)	n(s)		
Non- Qualified Stock Option (right to buy)	\$5.56	06/16/2023			A		23,500 ⁽¹⁾		(1)	06/15/2033	Common Stock	23,500	\$0.00	23,500		I	See Footnotes ⁽¹⁾ (2)(3)(4)(5)(6) (7)
Non- Qualified Stock Option (right to buy)	\$5.56	06/16/2023			A		23,500 ⁽¹⁾		(1)	06/15/2033	Common Stock	23,500	\$0.00	23,500		I	See Footnotes ⁽¹⁾ (3)(4)(5)(6)(7) (8)
		f Reporting Person											•				
(Last) 860 WAS	SHINGTO	(First) N STREET, 3RE	(Midd	le)													
(Street) NEW Y	ORK	NY	1001	4													
(City)		(State)	(Zip)														
1. Name ai <u>667, L.</u>		f Reporting Person	*														
(Last) 860 WAS	SHINGTO	(First) N STREET, 3RI	(Midd	le)													
(Street) NEW Y	ORK	NY	1001	4													
(City)		(State)	(Zip)														
	nd Address o	f Reporting Person $\overset{\circ}{ ext{N}}$	*														

860 WASHINGTON STREET, 3RD FLOOR							
(Street) NEW YORK	NY	10014					
(City)	(State)	(Zip)					
1. Name and Address of Reporting Person* BAKER FELIX							
(Last) 860 WASHINGTO	(First) N STREET, 3RD FL	(Middle)					
(0)							
(Street) NEW YORK	NY	10014					
(City)	(State)	(Zip)					
1. Name and Address of Reporting Person* Baker Bros. Advisors (GP) LLC							
(Last) 860 WASHINGTO	(First) N STREET, 3RD FL	(Middle) OOR					
(Street) NEW YORK	NY	10014					
(City)	(State)	(Zip)					
1. Name and Address of Reporting Person* <u>Baker Brothers Life Sciences LP</u>							
(Last) 860 WASHINGTO	ast) (First) (Middle) 0 WASHINGTON STREET, 3RD FLOOR						
(Street) NEW YORK	NY	10014					
(City)	(State)	(Zip)					

Explanation of Responses:

1. Includes 23,500 non-qualified stock options ("Stock Options") exercisable solely into common stock ("Common Stock") of Prelude Therapeutics Incorporated (the "Issuer") granted by the Issuer to Julian C. Baker, a managing member of Baker Bros. Advisors (GP) LLC (the "Advisor GP"), on June 16, 2023, in his capacity as a director of the Issuer, pursuant to the Issuer's 2020 Stock Incentive Plan, as Amended and Restated ("Incentive Plan") The Stock Options have a strike price of \$5.56 per share, and vest on the earlier of the first anniversary of the grant date or the date of the next annual general shareholders' meeting of the Issuer, subject to Julian C. Baker's continued service on the board of directors of the Issuer (the "Board") or an earlier change in control of the Issuer. The Stock Options expire on June 15, 2033. Julian C. Baker serves on the Board as a representative of the Funds (as defined below) and their affiliates.

- 2. After giving effect to the transaction reported herein and as a result of their ownership interest in (i) Baker Biotech Capital, L.P. and (ii) 667, L.P. ("667" and together with Baker Brothers Life Sciences, L.P. ("Life Sciences"), the "Funds"), Julian C. Baker and Felix J. Baker each may be deemed to have an indirect pecuniary interest in the Stock Options reported in column 9 of Table II held for the benefit of 667, a limited partnership of which the sole general partner is Baker Biotech Capital, L.P., a limited partnership of which the sole general partner is Baker Biotech Capital (GP), LLC, due to their interest in 667 and Baker Biotech Capital, L.P.'s right to receive an allocation of a portion of the profits from 667.
- 3. Pursuant to the policies of Baker Bros. Advisors LP (the "Adviser"), Julian C. Baker does not have a right to any of the Issuer's securities issued as compensation for his service on the Board and the Funds are entitled to an indirect proportionate pecuniary interest in the securities. The Funds each own an indirect proportionate pecuniary interest in the Stock Options. Solely as a result of their ownership interest in (i) the general partners of the Funds and (ii) the Funds, Felix J. Baker and Julian C. Baker may be deemed to have an indirect pecuniary interest in the Stock Options and any Common Stock acquired upon the exercise of Stock Options (i.e. no direct pecuniary interest).
- 4. Pursuant to agreements between Julian C. Baker and the Adviser and the policies of the Adviser, the Adviser has voting and dispositive power over the Stock Options and any Common Stock received as a result of the exercise of Stock Options.
- 5. The acquisition of Stock Options reported on this form represents a single grant to Julian C. Baker of 23,500 Stock Options on Table II. This grant of 23,500 Stock Options for Julian C. Baker is reported for each of the Funds as each has an indirect pecuniary interest in such securities.
- 6. The Adviser serves as the investment adviser to the Funds. In connection with the services provided by the Adviser, the Adviser receives an asset-based management fee that does not confer any pecuniary interest in the securities held directly by the Funds or for the benefit of the Funds. The Adviser GP is the Adviser's sole general partner. Julian C. Baker and Felix J. Baker are managing members of the Adviser has complete and unlimited discretion and authority with respect to the investment and voting power of the securities held directly by the Funds or for the benefit of the Funds. The general partners of the Funds have relinquished to the Adviser all discretion and authority with respect to the investment and voting power of the securities held directly by the Funds of for the benefit of the Funds.
- 7. Julian C. Baker, Felix J. Baker, the Adviser GP and the Adviser disclaim beneficial ownership of the securities held directly by the Funds or for the benefit of the Funds except to the extent of their pecuniary interest therein, and this report shall not be deemed an admission that any of Julian C. Baker, Felix J. Baker, the Adviser GP or the Adviser is a beneficial owner of such securities for purposes of Section 16 or any other purpose.

8. After giving effect to the transaction reported herein and as a result of their ownership interest in (i) Baker Brothers Life Sciences Capital, L.P. and (ii) Life Sciences, Julian C. Baker and Felix J. Baker each may be deemed to have an indirect pecuniary interest in the Stock Options reported in column 9 of Table II held for the benefit of Life Sciences, a limited partnership of which the sole general partner is Baker Brothers Life Sciences Capital, L.P., a limited partnership of which the sole general partner is Baker Brothers Life Sciences Capital (GP), LLC, due to their interest in Life Sciences and Baker Brothers Life Sciences Capital, L.P.'s right to receive an allocation of a portion of the profits from Life Sciences.

Remarks

Julian C. Baker, a managing member of Baker Bros. Advisors (GP) LLC, the sole general partner of Baker Bros. Advisors LP, is a director of Prelude Therapeutics Incorporated (the "Issuer"). By virtue of their representation on the board of directors of the Issuer, for purposes of Section 16 of the Securities Exchange Act of 1934, as amended, the reporting persons other than Julian C. Baker are deemed directors by deputization of the Issuer.

By: Baker Bros. Advisors LP,
Name: Scott L. Lessing, Title:
President /s/ Scott L. Lessing
Baker Bros. Advisors LP,
Mgmt. Co. and Inv. Adviser to
667, L.P., pursuant to authority
granted by Baker Biotech
Capital, L.P., GP to 667, L.P.
Name: Scott L. Lessing, Title:
President /s/ Scott L. Lessing

/s/ Julian C. Baker 06/21/2023 /s/ Felix J. Baker 06/21/2023 By: Baker Bros. Advisors (GP)

06/21/2023

06/21/2023

LLC, Name: Scott L. Lessing,

Title: President /s/ Scott L.

Lessing

Baker Bros. Advisors LP, Mgmt. Co. and Inv. Adviser to BAKER BROTHERS LIFE SCIENCES, L.P., pursuant to

authority granted by Baker

Brothers Life Sciences Capital, L.P., GP to Baker Brothers Life Sciences, L.P., Name: Scott L. Lessing, Title: President /s/

** 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.