FORM 4

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

Washington, D.C. 20549

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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).

Check this box to indicate that a Check this box to indicate that a transaction was made pursuant to a contract, instruction or written plan for the purchase or sale of equity securities of the issuer that is intended to satisfy the affirmative defense conditions of Rule 10b5-1(c). See Instruction 10.

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

Instruc	uon 10.																		
1. Name and Address of Reporting Person* HOUGEN ELIZABETH L					2. Issuer Name and Ticker or Trading Symbol IONIS PHARMACEUTICALS INC [IONS]									Relationship of Reporting Person(s) to Issuer (Check all applicable) Director 10% Owner					
(Last) (First) (Middle) 2855 GAZELLE COURT							of Earliest	Trans	saction (Month	ı/Day	y/Year)		Officer (give title Other (specify below) below) EVP, Finance & CFO						
(Street)	BAD C	A	92010		4. If Amendment, Date of Original Filed (Month/Day/Year)							Line	Individual or Joint/Group Filing (Check Applicable ne) Form filed by One Reporting Person Form filed by More than One Reporting Person						
(City)	(S	tate)	(Zip)											1 010011					
		Tal	ole I - Noi	า-Deriv	ative	Se	curitie	s Ac	quired, Di	spo	osed of	f, or Ber	eficially	y Owned					
1. Title of Security (Instr. 3) 2. Trans Date (Month/I					Execution Date,			Code (Inst	Transaction Disposed Of (D) (Instance Code (Instr. 5)				Securities Fe Beneficially (E		6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4)		7. Nature of Indirect Beneficial Ownership		
								Code V	4	Amount	(A) or (D) Price						Instr. 4)		
	Table II - Derivative Securities Acquired, Disposed of, or Beneficially Owned (e.g., puts, calls, warrants, options, convertible securities)																		
				ansact						es G Security	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)	Beneficial Ownership (Instr. 4)				
		Code V (A) (D) Exercisable Date Title Shares																	
Non- Qualified Stock Option (right to buy)	\$34.69	01/02/2025			A		31,000		01/02/2026 ⁽¹⁾	01/	/01/2035	Common Stock	31,000	\$0.0	31,00	0	D		

Explanation of Responses:

1. Grant on 01/02/2025 to reporting person of stock options under the Ionis Pharmaceuticals, Inc. Amended and Restated 2011 Equity Incentive Plan. Following this transaction, the option was exercisable as to 0 shares on 01/02/2025. 25% of the shares subject to the option will vest and become exercisable on 01/02/2026. Thereafter, the remaining shares subject to the option will vest and become exercisable in 36 equal monthly installments over the next 3 years.

> By: Patrick R. O'Neil, attorney-01/03/2025

in-fact For: Elizabeth L.

<u>Hougen</u>

** 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.