FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

OMB APPRO	DVAL					
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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 Address of Reporting Person* HOUGEN ELIZABETH L (Last) (First) (Middle) C/O IONIS PHARMACEUTICALS, INC.							2. Issuer Name and Ticker or Trading Symbol IONIS PHARMACEUTICALS INC] 3. Date of Earliest Transaction (Month/Day/Year) 11/14/2016										5. Relationship of Reporting Person(s) to Issuer (Check all applicable) Director 10% Owner X Officer (give title Other (specify below) SVP, Finance & CFO					
2855 GAZELLE COURT (Street) CARLSBAD CA 92010 (City) (State) (Zip)							4. If Amendment, Date of Original Filed (Month/Day/Year) tive Securities Acquired, Disposed of, or Benefic										S. Individual or Joint/Group Filing (Check Applicable Line) X Form filed by One Reporting Person Form filed by More than One Reporting Person					
1. Title of Security (Instr. 3) 2. Transac Date (Month/Da						ar) i	2A. Deemed Execution Date, if any (Month/Day/Year)			3. Transact Code (In: 8)	ion	4. Securities Acquired (A) Disposed Of (D) (Instr. 3, 4 5)			or 5. Amou Securiti Benefici Owned		nt of es ally Following	6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4)		7. Nature of Indirect Beneficial Ownership (Instr. 4)		
						-	Code	,	Amount	(A) o (D)	r Pri	се	Reporte Transac (Instr. 3	tion(s)			insu. 4)					
Common	Stock		1/2016	2016 11/14/2016				M ⁽¹⁾		2,500(1) A	\$1	11.27	9,	048		D					
Common	n Stock 11/14/2016 11/14/2016 s ⁽¹⁾ 2,500 ⁽¹⁾ D \$47.5 6,548 D																					
		Т	able II -										, or Ben ble sec			Owned						
1. Title of Derivative Security (Instr. 3)	2. Conversion or Exercise Price of Derivative Security	3. Transaction Date (Month/Day/Year)	3A. Deemd Execution if any (Month/Da	Date,	4. Transactioi Code (Instr 8)					Date Exer piration E onth/Day/	ate		7. Title and Amount of Securities Underlying Derivative Security (Instr. 3 and 4)		D S (I	. Price of erivative ecurity nstr. 5)	9. Number derivative Securities Beneficial Owned Following Reported Transactio (Instr. 4)	e s lly	10. Ownership Form: Direct (D) or Indirect (I) (Instr. 4)	11. Nature of Indirect Beneficial Ownership (Instr. 4)		
					Code	v	(A)	(D)	Dat Exe	te ercisable		expiration Date	Title	Amou or Numl of Share	ber							
Employee Stock Option (right to buy)	\$11.27	11/14/2016	11/14/2	2016	М			2,500	01/	./04/2014	0:	1/03/2017	Common Stock	2,50	00	\$0	2,500		D			

Explanation of Responses:

1. Acquired as a result of exercising a stock option that was scheduled to expire on 1/3/2017. The purchase and sale reported on this Form 4 was effective pursuant to a Rule 10b5-1 Trading Plan adopted by the reporting person on 2/3/2015.

Remarks:

/s/Elizabeth L. Hougen 11/16/2016

** 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.