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FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

Check this box if no longer subject to
Section 16. Form 4 or Form 5
obligations may continue. See
Instruction 1(b).

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

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

OMB APPROVAL 0005 5

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	hours per response:	0.5
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1. Name and Addre	1 0		2. Issuer Name and Ticker or Trading Symbol <u>IONIS PHARMACEUTICALS INC</u> [IONS]		tionship of Reporting Person(s) to Issuer (all applicable) Director 10% Owner Officer (give title Other (specify		
(Lact)	(Firct)	(Middle)		<u> </u>	below)	below)	
(Last) (First) (Middle) C/O IONIS PHARMACEUTICALS, INC. 2855 GAZELLE COURT			3. Date of Earliest Transaction (Month/Day/Year) 12/05/2016		SVP, Finance & CFO		
<u>.</u>			4. If Amendment, Date of Original Filed (Month/Day/Year)		ividual or Joint/Group Filing (Check Applicable		
(Street)				Line)			
CARLSBAD	СА	92010			Form filed by One Re	porting Person	
					Form filed by More the Person	an One Reporting	
(City)	(State)	(Zip)					

Table I - Non-Derivative Securities Acquired, Disposed of, or Beneficially Owned

1. Title of Security (Instr. 3)	2. Transaction Date (Month/Day/Year)	Execution Date,		ction Instr.				5. Amount of Securities Beneficially Owned Following Reported	Form: Direct (D) or Indirect (I) (Instr. 4)	7. Nature of Indirect Beneficial Ownership (Instr. 4)
			Code	v	Amount	(A) or (D)	Price	Transaction(s) (Instr. 3 and 4)		(1130.4)
Common Stock	12/05/2016	12/05/2016	M ⁽¹⁾		2,500 ⁽¹⁾	Α	\$11.27	9,048	D	
Common Stock	12/05/2016	12/05/2016	S ⁽¹⁾		2,500 ⁽¹⁾	D	\$46.05	6,548	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. Transa Code (8)		5. Number of Derivative Securities Acquired (A) or Disposed of (D) (Instr. 3, 4 and 5)		ate Amount of		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	Title	Amount or Number of Shares	4			
Employee Stock Option (right to buy)	\$11.27	12/05/2016	12/05/2016	М			2,500	01/04/2014	01/03/2017	Common Stock	2,500	\$0	0	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 Hougen

12/07/2016

** 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.