FORM 4

## **UNITED STATES SECURITIES AND EXCHANGE COMMISSION**

Washington, D.C. 20549

## STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

1	OIVID APPROVAL								
	OMB Number:	3235-0287							
	Estimated average burden								

0.5

hours per response

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*  Geary Richard S						2. Issuer Name and Ticker or Trading Symbol  ISIS PHARMACEUTICALS INC [ ISIS ]					(Che	ck all application	able)	Person(s) to Iss	wner
(Last) (First) (Middle) C/O ISIS PHARMACEUTICALS, INC. 2855 GAZELLE COURT						3. Date of Earliest Transaction (Month/Day/Year) 01/15/2014						below)	fficer (give title Other (spec below) SVP, Development		specily
(Street) CARLSBAD CA 92010 (City) (State) (Zip)					4. If Ai	4. If Amendment, Date of Original Filed (Month/Day/Year)					6. Inc Line)	′			
		Та	ble I - No	on-Deri	vative \$	Securities Acc	quired	l, Di	sposed of	or Ber	neficially	Owned			
1. Title of Security (Instr. 3)  2. Transact Date (Month/Day				n/Day/Year)   E	2A. Deemed Execution Date, if any (Month/Day/Year)	3. Transaction Code (Instr. 8)					es ally Following	6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4)	7. Nature of Indirect Beneficial Ownership		
							Code	v	Amount	(A) or (D)	Price	Reported Transact (Instr. 3 a	ion(s)		(Instr. 4)
Common Stock 01/15/3					/2014	01/15/2014	M <sup>(1)</sup>		776(1)	A	\$0	4,386		D	
Common Stock 01/1					/2014	01/16/2014	<b>S</b> <sup>(2)</sup>		296(2)	D	\$47.081	4,090		D	
Common Stock 01/15/2					/2014	01/15/2014	M <sup>(3)</sup>		1,398(3)	A	\$0	5,488		D	
Common Stock 01/16/2						01/16/2014	S <sup>(4)</sup>		533(4)	D	\$47.081	3 4,9	955	D	
			Table II			ecurities Acqualls, warrants						Owned			
1. Title of Derivative Security (Instr. 3)	rative   Conversion   Date rity   or Exercise   (Month/Day/Year)   if any   Conversion   Date		I. Fransaction Code (Instr 3)		6. Date Exercisable and Expiration Date (Month/Day/Year)		Date	7. Title and Amount of Securities Underlying Derivative Security (Instr. 3 and 4)		8. Price of Derivative Security (Instr. 5)	9. Numbe derivative Securities Beneficia Owned Following Reported	Ownershi Form: Direct (D) or Indirect (I) (Instr. 4	Beneficial Ownership (Instr. 4)		

## **Explanation of Responses:**

(5)

(5)

(5)

01/15/2014

01/15/2014

01/15/2014

Restricted

Stock

Units Restricted

Units Restricted

Stock

Units

1. Acquired pursuant to vesting and release of shares pursuant to an RSU award granted on January 15, 2012.

01/15/2014

01/15/2014

01/15/2014

2. The sale was made pursuant to an automatic sale to cover the required tax withholding obligations pursuant to the 2011 Equity Incentive Plan Restricted Stock Unit Agreement and Grant Notice, dated January 15, 2012

(A)

7,500

Code

Α

(D)

776<sup>(2)</sup>

1,398<sup>(4)</sup>

- $3. \ Acquired \ pursuant \ to \ vesting \ and \ release \ of \ shares \ pursuant \ to \ an \ RSU \ award \ granted \ on \ January \ 30, \ 2013.$
- 4. The sale was made pursuant to an automatic sale to cover the required tax withholding obligations pursuant to the 2011 Equity Incentive Plan Restricted Stock Unit Agreement and Grant Notice, dated January 30, 2013.
- 5. Each Restricted Stock Unit represents a contingent right to receive one share of Isis common stock, or its equivalent cash value.
- 6. Grant 1/15/2014 to reporting person of Restricted Stock Units under the Isis Pharmaceuticals, Inc. 2011 Equity Incentive Plan. Restricted Stock Units vest in four equal annual installments, the first installment vesting on January 15, 2015. Upon vesting, the restricted stock units will be paid out in whole shares of Isis common stock or cash as may be determined by the Company. The RSU shall be fully vested on January 15, 2018. The RSU is vested as to 0 shares on January 15, 2014.
- 7. Restricted Stock Unit Award granted on January 15, 2012. Restricted Stock Units vest in equal annual installments, the first installment vested on January 15, 2013. Upon vesting, the restricted stock units will be paid out in whole shares of Isis common stock or cash as may be determined by the Company. The RSU shall be fully vested on January 15, 2016.
- 8. Restricted Stock Unit Award granted on January 30, 2013. Restricted Stock Units vest in equal annual installments, the first installment vested on January 15, 2014. Upon vesting, the restricted stock units will be paid out in whole shares of Isis common stock or cash as may be determined by the Company. The RSU shall be fully vested on January 15, 2017.

## Remarks:

/s/B. Lynne Parshall, attorneyin-fact

Amount

or Number of Shares

7,500

776

1,398

\$0

\$0

Expiration

(6)

(7)

(8)

Date

Title

Common

Stock

Commor

Stock

Commor

Stock

Exercisable

(6)

(7)

(8)

01/17/2014

Transaction(s)

7,500

1,551

4,192

D

D

D

(Instr. 4)

III-Iact

\*\* 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.