FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

hours per response:

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0.5

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

1. Name and Address of Reporting Person* PARSHALL B LYNNE						2. Issuer Name and Ticker or Trading Symbol ISIS PHARMACEUTICALS INC [ISIS]							5. Relationship of Reporting Person(s) to Issuer (Check all applicable)					
												X				10% Ow		
(Last) (First) (Middle) C/O ISIS PHARMACEUTICALS, INC.													Officer (g below)	ive title		Other (sp below)	pecify	
					3. Date of Earliest Transaction (Month/Day/Year) 01/13/2012								Director, COO					
															- ,			
2855 GAZELLE COURT																		
(Street)					4. If Amendment, Date of Original Filed (Month/Day/Year)							6. Indi	6. Individual or Joint/Group Filing (Check Applicable Line)					
CARLSBAD		CA 92010								X	X Form filed by One Reporting Person							
					Form file:								d by More than One Reporting Person					
(City)	()	State)	(Zip)															
			Table I - Non-I	Deriva	tive :	Securiti	es Acqu	iired,	Disp	osed of, c	r Benef	icially C	Owned					
1. Title of Security (Instr. 3)								3. 4. Securities Acquired (A			A) or	or 5. Amount of		6. Own		. Nature of		
Date (Month/					Execution Dat Day/Year) if any		·	Code (Instr.		(D) (Instr. 3, 4 and 5)		Securities Beneficially Owned		(D) or Indirect		Indirect Beneficial		
					(Month)		Day/Year)	8)					Following Reported		(I) (Ins		Ownership (Instr. 4)	
							Code	v	Amount	it (A) or Pri		Transaction (Instr. 3 and	n(s) d 4)					
			Table II - De	rivati	ivo S	ocuritios	. Acqui	rod D	ieno	end of or	Ponofic	ially Ov	vned					
										onvertible			viieu					
1. Title of Derivative Security (Instr. 3)	2. Conversion or Exercise Price of Derivative	3. Transaction Date (Month/Day/Year)	3A. Deemed Execution Date, if any (Month/Day/Year)	4. Transaction Code (Instr. 8)		5. Number of Derivative Securities Acquired (A) or Disposed of (D) (Instr. 3, 4 and 5)		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		10. Ownership Form: Direct (D) or Indirect	Beneficial Ownership (Instr. 4)		
	Security						(D)					Amount	1	Followin Reporte	ď	(I) (Instr. 4)		
										Expiration Date	Title	or Number of Shares		Transac (Instr. 4)				
				Code	v	(A)		Date Exerc	Date Exercisable									
Employee Stock Option (right to buy)	\$5.25	01/13/2012	01/13/2012	G ⁽¹⁾			20,000(1)	01/03/2010		01/02/2013	Common Stock	20,000	\$0	19,5	06	D		
Employee																	İ	
Stock Option (right to buy)	\$5.25	01/13/2012	01/13/2012	G ⁽¹⁾		10,000 ⁽¹⁾		01/03	3/2008	01/02/2013	Common Stock	10,000	\$0	15,0	00	I	By daughter	
Employee																		
Stock Option (right to	\$5.25	01/13/2012	01/13/2012	G ⁽¹⁾		10,000 ⁽¹⁾		01/03	3/2009	01/02/2013	Common Stock	10,000	\$0	15,0	00	I	By daughter	

Explanation of Responses:

1. The reporting person transferred options to her two daughters who share reporting person's household, divided equally.

Remarks:

/s/B. Lynne Parshall

01/13/2012

** 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.