FORM 4

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

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

Washington, D.C. 20549

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

OMB APPROVAL

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Instruction 1(b).	oritinae. Gee	Filed	pursuant to Section 16(a) of the Securities Exchange Act of 1934		<u>L'</u>	nours per respon	ise. 0.5
			or Section 30(h) of the Investment Company Act of 1940				
1. Name and Addres	ss of Reporting Perso	n <sup>*</sup>	2. Issuer Name and Ticker or Trading Symbol ISIS PHARMACEUTICALS INC [ ISIS ]		ionship of Rep all applicable)	oorting Person	(s) to Issuer
PARSHALL	<u>D LIININE</u>			X	Director		10% Owner
				X	Officer (give	title	Other (specify
(Last)	ast) (First) (Middle)		3. Date of Earliest Transaction (Month/Day/Year)		below)		below)
C/O ISIS PHAR	MACEUTICALS	, INC.	11/09/2006		Executi	ive Vice Pre	sident
1896 RUTHERF	ORD ROAD						
			4. If Amendment, Date of Original Filed (Month/Day/Year)		dual or Joint/G	Group Filing (C	heck Applicable
(Street)				Line)			
CARLSBAD	CA	92008		X	Form filed by	y One Reportir	ng Person
						y More than O	ne Reporting
(City)	(State)	(Zip)			Person		

(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)	2A. Deemed Execution Date, if any (Month/Day/Year)	3. Transaction Code (Instr. 8)		4. Securities A Disposed Of (	Acquired (D) (Instr.	(A) or 3, 4 and 5)	5. Amount of Securities Beneficially Owned 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 Transaction(s) (Instr. 3 and 4)		(Instr. 4)			
Common Stock	11/09/2006	11/09/2006	M <sup>(1)</sup>		5,000(1)	A	\$5.15	5,000	I	By daughter			
Common Stock	11/09/2006	11/09/2006	S <sup>(2)</sup>		5,000(2)	D	\$10.13	0	I	By daughter			
Common Stock	11/09/2006	11/09/2006	M <sup>(1)</sup>		5,000(1)	A	\$5.15	5,000	I	By daughter			
Common Stock	11/09/2006	11/09/2006	S <sup>(2)</sup>		5,000 <sup>(2)</sup>	D	\$10.13	0	I	By daughter			
Common Stock	11/09/2006	11/09/2006	M <sup>(1)</sup>		10,000(1)	A	\$5.15	13,241	D				
Common Stock	11/09/2006	11/09/2006	S <sup>(2)</sup>		10,000(2)	D	\$10.13	3,241	D				
Common Stock	11/13/2006	11/13/2006	M <sup>(1)</sup>		10,000(1)	A	\$5.15	10,000	I	By daughter			
Common Stock	11/13/2006	11/13/2006	S <sup>(2)</sup>		10,000(2)	D	\$13.36	0	I	By daughter			
Common Stock	11/13/2006	11/13/2006	M <sup>(1)</sup>		10,000(1)	A	\$5.15	10,000	I	By daughter			
Common Stock	11/13/2006	11/13/2006	S <sup>(2)</sup>		10,000(2)	D	\$13.36	0	I	By daughter			
Common Stock	11/13/2006	11/13/2006	<b>M</b> <sup>(1)</sup>		10,000(1)	A	\$5.15	13,241	D				
Common Stock	11/13/2006	11/13/2006	S <sup>(2)</sup>		10,000(2)	D	\$13.36	3,241	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)		of		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 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				
Employee Stock Option (right to buy)	\$5.15	11/09/2006	11/09/2006	M			5,000	01/01/2006	12/31/2008	Common Stock	5,000	\$0	10,000	I	By daughter
Employee Stock Option (right to buy)	\$5.15	11/09/2006	11/09/2006	M			5,000	01/01/2006	12/31/2008	Common Stock	5,000	\$0	10,000	I	By daughter

	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)		of Deri Sec Acq (A) ( Disp of (I	ivative urities juired or posed D) (Instr. and 5)	Expiration Date of Si (Month/Day/Year) Und Deri (Inst		Expiration Date		Expiration Date (Month/Day/Year)		Expiration Date		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 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										
Employee Stock Option (right to buy)	\$5.15	11/09/2006	11/09/2006	M			10,000	01/01/2006	12/31/2008	Common Stock	10,000	\$0	16,466	D							
Employee Stock Option (right to buy)	\$5.15	11/13/2006	11/13/2006	М			10,000	01/01/2006	12/31/2008	Common Stock	10,000	\$0	0	I	By daughter						
Employee Stock Option (right to buy)	\$5.15	11/13/2006	11/13/2006	M			10,000	01/01/2006	12/31/2008	Common Stock	10,000	\$0	0	I	By daughter						
Employee Stock Option (right to buy)	\$5.15	11/13/2006	11/13/2006	М			10,000	01/01/2006	12/31/2008	Common Stock	10,000	\$0	6,466	D							

## Explanation of Responses:

- $1. The purchase \ reported \ was \ effected \ pursuant \ to \ a \ Rule \ 10b5-1 \ trading \ plan \ adopted \ by \ the \ reporting \ person \ on \ 10-2-06.$
- 2. The sale reported was effected pursuant to a Rule 10b5-1 trading plan adopted by the reporting person on 10-2-06.

## Remarks:

<u>/s/B. Lynne Parshall</u> <u>11/13/2006</u>

\*\* Signature of Reporting Person

Date

Reminder: Report on a separate line for each class of securities beneficially owned directly or indirectly.

- $^{\star}$  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.