## 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).

Date

Year)

Common Stock

Common Stock

(Month/ Day

10/15/02

10/15/02

Date.

lif anv

Year)

(Month/Day

10/15/02

10/15/02

## STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

OMB Number: 3235-0287

OMB APPROVAL

Expires: January 31, 2005 Estimated average burden hours per response. . .0.5

Direct (D)

(Instr. 4)

3245

or Indirect (I)

D

(Instr. 4)

Filed pursuant to Section 16(a) of the Securities Exchange Act of 1934, Section 17(a) of the Public Utility Holding Company Act of 1935 or Section 30(h) of the Investment Company Act of 1940 Filed By Romeo & Dye's Instant Form 4 Filer

2. Issuer Name **and** Ticker or Trading Symbol 6. Relationship of Reporting Person(s) 1. Name and Address of Reporting Person\* Isis Pharmaceuticals, Inc. (ISIS) to Issuer (Check all applicable) Parshall B. Lynne X Director 10% Owner  $\mathbf{X}$  Officer (give title below) Other (specify below) (Last) (First) (Middle) 3. I.R.S. Identification Number 4. Statement for of Reporting Person, Month/Day/Year 2292 Faraday Avenue if an entity (voluntary) 10/15/02 Executive Vice President 5. If Amendment, 7. Individual or Joint/Group Filing (Check Applicable Line) (Street) Date of Original X Form filed by One Reporting Person Carlsbad, CA 92008 (Month/Day/Year) Form filed by More than One Reporting Person (City) (State) (Zip) Table I — Non-Derivative Securities Acquired, Disposed of, or Beneficially Owned 2A. Deemed 3. Trans-4. Securities Acquired (A) or Disposed of (D) 5. Amount of 1. Title of Security 2. Trans-6. Owner-7. Nature of Indirect action Code (Instr. 3, 4 & 5) ship Form: Beneficial Ownership (Instr. 3) action Execution Securities

Price

\$5,75

\$8,9662

Beneficially

Owned Follow-

(Instr. 3 & 4)

ing Reported Transactions(s)

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

(Instr. 8)

Code

M

 $S^{(1)}$ 

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(A)

or

(D)

D

5000

5000

## FORM 4 (continued) Table II - Derivative Securities Acquired, Disposed of, or Beneficially Owned (e.g., puts, calls, warrants, options, convertible securities)

Amount

			(8	,, Pu	o, cuiio, wa	runes, open	,,, сог	. , с. с. с.	ic occurr	110)	(eig., pate), variance, options, convertible securities)													
1. Title of	2. Conver-	3. Trans-	3A.	4.	5. Number of Derivative		6. Date		7. Title and Amount		8. Price of	9. Number of	10.	11. Nature										
Derivative	sion or	action	Deemed	Trans-	Securities Acquired (A) or		Exercisable		of Underlying		Derivative	Derivative	Owner-	of Indirect										
Security	Excercise	Date	Execution	action	Disposed of (D)		and Expiration Secu		Securities		Security	Securities	ship	Beneficial										
	Price of		Date,	Code			Date		(Instr. 3 &	4)	(Instr. 5)	Beneficially	Form	Ownership										
(Instr. 3)	Derivative	(Month/	if any		(Instr. 3, 4 & 5)		(Month/Day/ Year)				1	Owned	of	(Instr. 4)										
	Security	Day/ Year)	(Month/	(Instr.								Following	Deriv-											
		′	Day/ Year)	8)								Reported	ative											
			<b>_</b>									Transaction(s)	Security:											
				Code	(A)	(D)	Date	Expira-	Title	Amount	1	(Instr. 4)	Direct											
					` ′	` ′	Exer-	tion		or			(D)											
							cisable	Date		Number			or											
										of			Indirect											
							1			Shares			(I)											
													(Instr. 4)											
Common	\$5.75	10/15/02	10/15/02	M		500	0 11/1/96	10/17/02	Common	5000		3628	D											
Stock				1			1		Stock					1										

Explanation of Responses:

(1) The sale reported in this Form 4 was effected pursuant to a Rule 10b5-1 trading plan adopted by the reporting person on February 28, 2002.

By: /s/ B. Lynne Parshall **B. Lynne Parshall** \*\*Signature of Reporting Person

10/15/02 Date

\*\*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.

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<sup>\*</sup> If the form is filed by more than one reporting person, see Instruction 4(b)(v).