FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

	омв г
STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP	Exnire

OMB Number:	3235-0287									
Expires:	December 31, 2014									
Estimated average burden										
hours per response:	0.5									

OMB APPROVAL

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*					2. Issuer Name and Ticker or Trading Symbol ISIS PHARMACEUTICALS INC [ISIS]									5. Relationship of Reporting Person(s) to Issuer (Check all applicable)				
HOLMLUD JON T				— <u>;</u>	3. Date of Earliest Transaction (Month/Day/Year)							- `	Dire	ctor er (give title	10% (Other below	(specify		
(Last)	(Fir	rst) (Middle)		07/01/2003										Vice P	vice President		
(Street)					4. If Amendment, Date of Original Filed (Month/Day/Year)									6. Individual or Joint/Group Filing (Check Applicable Line)				
(City)	(St	ate) (.	Zip)										X Form filed by One Reporting Person Form filed by More than One Reporting Person					
		Tabl	e I - Non-De	erivati	ive S	ecuri	ies Ac	quired	, Dis	posed o	f, or	Bene	eficia	ally Own	ed			
1. Title of Security (Instr. 3) 2. Transac Date (Month/Date)				•	Execution Date,		Code	Transaction Disposed Code (Instr. 5)		rities Acquired (A) d Of (D) (Instr. 3, 4			nd Secur Benet	icially d Following	6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4)	7. Nature of Indirect Beneficial Ownership		
								Code	v	Amount	(A (D) or)	Price	Trans	rted action(s) . 3 and 4)		(Instr. 4)	
Common Stock			07	07/01/2003		07/0	1/2003	J	V	983(1)	() A		4.39	945	6,576	D		
		Та	ble II - Deriv (e.g.							sed of, onvertib				y Owned				
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)	Co	Transaction Code (Instr.		n of E		6. Date Exercisable and Expiration Date (Month/Day/Year)			7. Title and Amount of Securities Underlying Derivative Security (Instr. and 4)		8. Price of Derivative Security (Instr. 5)		Ownership Form: Direct (D) or Indirect (I) (Instr. 4)	11. Nature of Indirect Beneficial Ownership (Instr. 4)	
				Cod	ode V	(A) (D)	Date Exercisa		Expiration Date	Title	Amo or Num of Sha	ber					

Explanation of Responses:

1. Reporting 983 shares acquired under the Isis Pharmaceuticals, Inc. 2000 Employee Stock Purchase Plan on July 1, 2003.

07/02/2003 Jon T. Holmlund

** 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.