FORM 3

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

Washington, D.C. 20549

## INITIAL STATEMENT OF BENEFICIAL OWNERSHIP OF SECURITIES

	OMB APPROVAL						
	OMB Number: 3235-0104 Estimated average burden						
	hours per response						

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*  DiMarchi Richard D  2. Date of Event Requiring Statement (Month/Day/Year) 12/14/2004			3. Issuer Name and Ticker or Trading Symbol  ISIS PHARMACEUTICALS INC [ ISIS ]							
ast) (First) (Middle) NDIANA UNIVERSITY, CHEMISTRY			Relationship of Reporting Person(s) to Issuer (Check all applicable)     X Director 10% Owner		(Mo	5. If Amendment, Date of Original Filed (Month/Day/Year)				
A508 800 EAST KIRKWOOD			Officer (give title below)	Other (spe below)	App	5. Individual or Joint/Group Filing (Check Applicable Line) X Form filed by One Reporting Person				
(Street) BLOOMINGTON IN 47405							Form filed b Reporting P	y More than One erson		
(City) (State) (Zip)										
Table I - Non-Derivative Securities Beneficially Owned										
1. Title of Security (Instr. 4)			Beneficially Owned (Instr. 4)				4. Nature of Indirect Beneficial Ownership (Instr. 5)			
No securities benefically owned				0	D					
Table II - Derivative Securities Beneficially Owned (e.g., puts, calls, warrants, options, convertible securities)										
1. Title of Derivative Security (Instr. 4)  2. Date Exercisable at Expiration Date (Month/Day/Year)			Underlying Derivative Security (Instr. 4) Conv		4. Conversion or Exercise Price of	5. Ownership Form: Direct (D)	6. Nature of Indirect Beneficial Ownership (Instr. 5)			
	Date Exercisable	Expiration Date	n Title		Amount or Number of Shares	Derivative Security	or Indirect (I) (Instr. 5)			

**Explanation of Responses:** 

Remarks:

Richard D DiMarchi 12/15/2004

\*\* Signature of Reporting Person Date

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

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.

 $<sup>^{\</sup>star}$  If the form is filed by more than one reporting person, see Instruction 5 (b)(v).

<sup>\*\*</sup> Intentional misstatements or omissions of facts constitute Federal Criminal Violations See 18 U.S.C. 1001 and 15 U.S.C. 78ff(a).