FORM 4

ISALY SAMUEL D

(First)

601 LEXINGTON AVENUE, 54TH FLOOR

(Middle)

(Last)

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

subject to	STATEMENT
11 5	

Γ OF CHANGES IN BENEFICIAL OWNERSHIP

OMB APPROVAL OMB Number: Estimated average burden hours per response: 0.5

 $Footnotes^{(2)(4)}\\$

Footnotes(3)(4)

11. Nature of Indirect Beneficial Ownership (Instr. 4)

Section obligat	this box if no long 16. Form 4 or ions may continuition 1(b).	onger subject to Form 5 nue. See	S		ed pur	suant to	o Section	16(a	a) of th	he Se	Curities Exchan	ge Act o		RSHIP		OMB Nun Estimated hours per	l average	
1. Name and Address of Reporting Person* ORBIMED ADVISORS LLC (Last) (First) (Middle) 601 LEXINGTON AVENUE, 54TH FLOOR				PI PI 3.	2. Issuer Name and Ticker or Trading Symbol PIERIS PHARMACEUTICALS, INC. [PIRS] 3. Date of Earliest Transaction (Month/Day/Year) 01/03/2018 5. Relationship of (Check all applicate Director Officer (g below)									plicable) ctor cer (give	X 10% Owner			
(Street) NEW YORK NY 10022 (City) (State) (Zip)			4.	If Amer	ndment,	Date	of Ori	iginal	Filed (Month/Da	ay/Year)		Line) Forr	n filed b	y One R	eporting	eck Applicable Person Reporting		
		Tab	le I -	Non-Deriv	vativ	e Sec	urities	Ac	quir	red,	Disposed o	of, or E	Benefic	cially Own	ed			
1. Title of Security (Instr. 3) 2. Transaction Date (Month/Day/Ye			- 1	Execut if any	Deemed cution Date, y nth/Day/Year)		3. Transaction Code (Instr. 8)		4. Securities Acquired (A) or Disposed Of (D) (Instr. 3, 4 a			5. Amount of Securities Beneficially Owned Followin Reported		6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4)		7. Nature of Indirect Beneficial Ownership		
								c	Code	v	Amount	(A) or (D)	Price	Transactio		(Instr.		
Common	Stock			01/03/20	18				s		1,693,492	D	\$7.35	(1) 5,500,	,730	I		See Footnotes ⁽²⁾
Common	Common Stock 0:		01/03/20	18				S		16,128	D	\$7.35	(1) 49,2	70 I		I	See Footnotes ⁽³⁾	
		Ta	able								sposed of, s, convertib							
1. Title of Derivative Security (Instr. 3) 2. Conversion or Exercise Price of Derivative Security		Date Exec (Month/Day/Year) if an		Deemed ution Date, y hth/Day/Year)		saction (Instr.			Expiration e (Month/Da			7. Title and Amount of Securities Underlying Derivative Security (Instr. and 4)			9. Num derivat Securi Benefi Owned Follow Report Transa (Instr.	tive ties cially I ing ted action(s)	10. Owner Form: Direct or Indi (I) (Ins	(D) Benefici Ownersl rect (Instr. 4)
					Code	· V	(A)	(D)	Date Exe	e rcisab	Expiration Date	Title	Amoun or Numbe of Shares					
1		Reporting Person*	1								,			,	,			
(Last)	INGTON A	(First) AVENUE, 54TH		(Middle)														
(Street) NEW YO	ORK	NY		10022														
(City)		(State)		(Zip)		-												
		Reporting Person*																
(Last)	KINGTON A	(First) AVENUE, 54TH		(Middle)														
(Street) NEW YO	ORK	NY		10022														
(City)		(State)		(Zip)														
1. Name ar	nd Address of	Reporting Person*																

(Street) NEW YORK	NY	10022
(City)	(State)	(Zip)

Explanation of Responses:

- 1. These shares of the Issuer's common stock ("Shares") were sold in a block order at a price of \$7.35.
- 2. These Shares are held of record by OrbiMed Private Investments III, LP ("OPI III"). OrbiMed Capital GP III LLC ("GP III") is the general partner of OPI III, and OrbiMed Advisors LLC ("Advisors") is the managing member of GP III. Samuel D. Isaly ("Isaly"), a natural person, is the managing member of and owner of a controlling interest in Advisors. By virtue of such relationships, GP III, Advisors and Isaly may be deemed to have voting and investment power over the securities held by OPI III and as a result may be deemed to have beneficial ownership over such securities.
- 3. These Shares are held of record by OrbiMed Associates III, LP ("Associates III"). Advisors is the general partner of Associates III. Isaly is the managing member of and owner of a controlling interest in Advisors. By virtue of such relationships, Advisors and Isaly may be deemed to have voting and investment power over the securities held by Associates III and as a result may be deemed to have beneficial
- 4. This report on Form 4 is jointly filed by GP III, Advisors, and Isaly. Each of the Reporting Persons disclaims beneficial ownership of the securities reported herein for purposes of Rule 16a-1(a) under the Securities Exchange Act of 1934, as amended (the "Exchange Act"), except to the extent of its or his pecuniary interest therein, if any. This report shall not be deemed an admission that any of the Reporting Persons is a beneficial owner for the purpose of Section 16 of the Exchange Act, or for any other purpose.

01/04/2018 /s/ Samuel D. Isaly

** 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.