UNITED STATES SECURITIES AND EXCHANGE COMMISSION WASHINGTON, D.C. 20549

SCHEDULE 13G

UNDER THE SECURITIES EXCHANGE ACT OF 1934 (AMENDMENT NO. 6)*

	Isis Pharmaceutio	cals, Inc.	
	(Name of Iss	suer)	
	Common Sto	ock	
	(Title of Class of	Securities)	
	464330-10	-9	
•	(CUSIP Numl		
	December 31,	2003	
(Date of Event Which Requires I	Filing of this Statement)	
Check the app is filed:	propriate box to designate the	rule pursuant to which this Schedule	;
/x/ / / / /	Rule 13d-1(b) Rule 13d-1(c) Rule 13d-1(d)		
person's init securities, a		respect to the subject class of nt containing information which	
deemed to be Act of 1934	"filed" for the purpose of Sec "Act") or otherwise subject to it shall be subject to all other	of this cover page shall not be ction 18 of the Securities Exchange o the liabilities of that section er provisions of the Act (however,	
	Page of 9 Pa		
	SCHEDU	ULE 13G	
CUSIP NO. 464	1330-10-9	Page 2 of 9 Pages	;
• •	REPORTING PERSONS DENTIFICATION NOS. OF ABOVE PR	ERSONS (ENTITIES ONLY)	
	ıp Global Markets Inc. Salomon Smith Barney Inc."		
(2) CHECK TH		OF A GROUP (SEE INSTRUCTIONS)	
		(a) / / (b) / /	
(3) SEC USE			

New York

(4) CITIZENSHIP OR PLACE OF ORGANIZATION

	NUMBER OF	(5) SOLE VOTING POWER	0
	SHARES		
	BENEFICIALLY	(6) SHARED VOTING POWER	4,118,112
	OWNED BY		
	EACH	(7) SOLE DISPOSITIVE POWER	0
	REPORTING		
	PERSON	(8) SHARED DISPOSITIVE POWER	4,118,112
	WITH:		
(9)	AGGREGATE AMOUNT BE	NEFICIALLY OWNED BY EACH REPORTING PERSON	
(10)	CHECK IF THE AGGREINSTRUCTIONS) //	GATE AMOUNT IN ROW (9) EXCLUDES CERTAIN SHA	ARES (SEE
		EPRESENTED BY AMOUNT IN ROW (9)	7.4%
(12)		PERSON (SEE INSTRUCTIONS)	BD
CUSI	P NO. 464330-10-9		e 3 of 9 Pages
(1)	NAMES OF REPORTING I.R.S. IDENTIFICAT	ION NOS. OF ABOVE PERSONS (ENTITIES ONLY)	
	Citigroup Financia f/k/a "Salomon Bro	thers Holding Company Inc"	
(2)	CHECK THE APPROPRI	ATE BOX IF A MEMBER OF A GROUP (SEE INSTRUC	CTIONS)
			(a) // (b) //
(3)	SEC USE ONLY		
(4)	CITIZENSHIP OR PLA	CE OF ORGANIZATION	Delaware
	NUMBER OF	(5) SOLE VOTING DOWER	
		(3) SOLE VOTING FOWER	
	SHARES	(3) SOLE VOTING FOWER	
	SHARES	(6) SHARED VOTING POWER	4,118,112
	SHARES BENEFICIALLY		4,118,112
	SHARES BENEFICIALLY OWNED BY	(6) SHARED VOTING POWER (7) SOLE DISPOSITIVE POWER	4,118,112
	SHARES BENEFICIALLY OWNED BY EACH	(6) SHARED VOTING POWER	4,118,112

(9) AGGREGATE AMOUNT B	SENEFICIALLY OWNED BY EACH REPORTING PER	(SUN 4, 116, 112
(10) CHECK IF THE AGGR INSTRUCTIONS) /	REGATE AMOUNT IN ROW (9) EXCLUDES CERTA:	IN SHARES (SEE
(11) PERCENT OF CLASS	REPRESENTED BY AMOUNT IN ROW (9)	7.4%
(12) TYPE OF REPORTING	G PERSON (SEE INSTRUCTIONS)	CO
	SCHEDULE 13G	
CUSIP NO. 464330-10-9		Page 4 of 9 Pages
(1) NAMES OF REPORTIN	IG PERSONS ATION NOS. OF ABOVE PERSONS (ENTITIES OF	NLY)
Citigroup Global f/k/a "Salomon Sm	Markets Holdings Inc. nith Barney Holdings Inc."	
(2) CHECK THE APPROPR	RIATE BOX IF A MEMBER OF A GROUP (SEE IN	NSTRUCTIONS)
		(a) // (b) //
(3) SEC USE ONLY (4) CITIZENSHIP OR PL	ACE OF ORGANIZATION	New York
NUMBER OF	(5) SOLE VOTING POWER	0
SHARES		
BENEFICIALLY OWNED BY	(6) SHARED VOTING POWER	5,626,062
EACH	(7) SOLE DISPOSITIVE POWER	
REPORTING		
PERSON	(8) SHARED DISPOSITIVE POWER	5,626,062
WITH:		
(9) AGGREGATE AMOUNT B	BENEFICIALLY OWNED BY EACH REPORTING PER	RSON 5,626,062
(10) CHECK IF THE AGGR INSTRUCTIONS) /	REGATE AMOUNT IN ROW (9) EXCLUDES CERTAI	IN SHARES (SEE
	REPRESENTED BY AMOUNT IN ROW (9)	10.1%
	G PERSON (SEE INSTRUCTIONS)	HC

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SCHEDULE 13G

CUSIP NO. 4643	330-10-9		Page 5 of 9 Pages		
(1) NAMES OF I.R.S. ID	REPORTING PE	RSONS NOS. OF ABOVE PERSONS (ENTITIES ON	NLY)		
Citigroup	Inc.				
(2) CHECK THE	APPROPRIATE	BOX IF A MEMBER OF A GROUP (SEE IN			
			(a) // (b) //		
(3) SEC USE C					
(4) CITIZENSH	HIP OR PLACE (OF ORGANIZATION	Delaware		
NUMBER	0F	(5) SOLE VOTING POWER	0		
SHARES	5				
BENEFICIA	ALLY	(6) SHARED VOTING POWER	5,679,982*		
OWNED B	BY				
EACH		(7) SOLE DISPOSITIVE POWER	0		
REPORTIN	IG				
PERSON		(8) SHARED DISPOSITIVE POWER	5,679,982*		
WITH:					
		ICIALLY OWNED BY EACH REPORTING PER			
(10) CHECK IF		E AMOUNT IN ROW (9) EXCLUDES CERTAI			
		ESENTED BY AMOUNT IN ROW (9)	10.2%*		
(12) TYPE OF R	REPORTING PER	SON (SEE INSTRUCTIONS)	НС		
		the other reporting persons.			
Item 1(a).	Name of Iss	uer:			
	Isis Pharmaceuticals, Inc.				
Item 1(b).	Address of	Issuer's Principal Executive Office	es:		
	2292 Farada Carlsbad, C				

Citigroup Global Markets Inc. ("CGM") Citigroup Financial Products Inc. ("CFP") Citigroup Global Markets Holdings Inc. ("CGM Holdings") Citigroup Inc. ("Citigroup") Item 2(b). Address or Principal Office or, if none, Residence: The address of the principal office of each of CGM, CFP and CGM Holdings is: 388 Greenwich Street New York, NY 10013 The address of the principal office of Citigroup is: 399 Park Avenue New York, NY 10043 Item 2(c). Citizenship or Place of Organization: CGM and CGM Holdings are New York corporations. CFP and Citigroup are Delaware corporations. Item 2(d). Title of Class of Securities: Common Stock Item 2(e). Cusip Number: 464330-10-9 Page 6 of 9 Pages Item 3. If this Statement is Filed Pursuant to Sections 240.13d-1(b) or 240.13d-2(b) or (c), Check Whether the Person Filing is a(n): [X] Broker or dealer registered under section 15 of the Act (15 U.S.C. 780); (b) [] Bank as defined in section 3(a)(6) of the Act (15 U.S.C. 78c); (c) [] Insurance company as defined in section 3(a)(19) of the Act (15 U.S.C. 78c); (d) [] Investment company registered under Section 8 of the Investment Company Act of 1940 (15 U.S.C. 80a-8); (e) [] Investment adviser in accordance with Section 240.13d-1(b)(1)(ii)(E); (f) [] Employee benefit plan or endowment fund in accordance with Section 240.13d-1(b)(1)(ii)(F); [X] Parent holding company or control person in accordance (g) with Section 240.13d-1(b)(1)(ii)(G); [] Savings association as defined in Section 3(b) of the (h) Federal Deposit Insurance Act (12 U.S.C. 1813); [] Church plan that is excluded from the definition of an (i) investment company under Section 3(c)(14) of the Investment Company Act of 1940 (15 U.S.C. 80a-3) (j) [] Group, in accordance with Section 240.13d-1(b)(1)(ii)(J). Ownership. (as of December 31, 2003) Item 4. (a) Amount beneficially owned: See item 9 of cover pages

(b) Percent of Class: See item 11 of cover pages

(c) Number of shares as to which the person has:

Item 2(a).

Name of Person Filing:

- (i) sole power to vote or to direct the vote:
- (ii) shared power to vote or to direct the vote:
- (iii) sole power to dispose or to direct the disposition of:
- (iv) shared power to dispose or to direct the disposition of:

See Items 5-8 of cover pages

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Item 5. Ownership of Five Percent or Less of a Class.

> If this statement is being filed to report the fact that as of the date hereof the reporting person has ceased to be the beneficial owner of more than five percent of the class of securities, check the following [].

Item 6. Ownership of More Than Five Percent on Behalf of Another Person.

Not Applicable.

Identification and Classification of the Subsidiary which Acquired Item 7. the Security being Reported on by the Parent Holding Company.

> CFP is the sole stockholder of CGM. CGM Holdings is the sole stockholder of CFP. Citigroup is the sole stockholder of CGM Holdings.

Item 8. Identification and Classification of Members of the Group.

Not Applicable.

Item 9. Notice of Dissolution of Group.

Not Applicable.

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Item 10. Certification.

By signing below I certify that, to the best of my knowledge and belief, the securities referred to above were acquired and are held in the ordinary course of business and were not acquired and are not held for the purpose of or with the effect of changing or influencing the control of the issuer of the securities and were not acquired and are not held in connection with or as a participant in any transaction having that purpose or effect.

STGNATURE.

After reasonable inquiry and to the best of my knowledge and belief, I certify that the information set forth in this statement is true, complete and correct.

Dated: February 10, 2004

CITIGROUP GLOBAL MARKETS INC.

By: /s/ Joseph B. Wollard

Name: Joseph B. Wollard

Title: Assistant Secretary

By: /s/ Joseph B. Wollard . Name: Joseph B. Wollard

Title: Assistant Secretary

CITIGROUP GLOBAL MARKETS HOLDINGS INC.

By: /s/ Joseph B. Wollard

Name: Joseph B. Wollard Title: Assistant Secretary

CITIGROUP INC.

By: /s/ Joseph B. Wollard

Name: Joseph B. Wollard Title: Assistant Secretary

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EXHIBIT INDEX TO SCHEDULE 13G -----

EXHIBIT 1

Agreement among CGM, CFP, CGM Holdings and Citigroup as to joint filing of Schedule 13G

EXHIBIT 1

AGREEMENT AS TO JOINT FILING OF SCHEDULE 13G

Each of the undersigned hereby affirms that it is individually eligible to use Schedule 13G, and agrees that this Schedule 13G is filed on its behalf.

Date: February 10, 2004

CITIGROUP GLOBAL MARKETS INC.

By: /s/ Joseph B. Wollard

Name: Joseph B. Wollard Title: Assistant Secretary

CITIGROUP FINANCIAL PRODUCTS INC.

By: /s/ Joseph B. Wollard

Name: Joseph B. Wollard Title: Assistant Secretary

CITIGROUP GLOBAL MARKETS HOLDINGS INC.

By: /s/ Joseph B. Wollard

Name: Joseph B. Wollard Title: Assistant Secretary

CITIGROUP INC.

By: /s/ Joseph B. Wollard

Name: Joseph B. Wollard Title: Assistant Secretary