Deutsche Bank AG Taunusanlage 12, D-60325 Frankfurt am Main Federal Republic of Germany

Jeffrey A. Ruiz Vice President

Telephone: (212) 469-3667

February 13, 2003

Securities and Exchange Commission SEC Document Control 450 Fifth Street, N.W. Washington, DC 20549 Attn: Filing Desk

Dear Sir or Madame:

Re: Filing of Schedule 13G - Isis Pharmaceuticals Incorporated

Pursuant to Rule 13d-1 of the Securities Exchange Act of 1934, attached is one copy of Schedule 13G with respect to the common stock of the above referenced corporation.

Please acknowledge your receipt of the Schedule  ${\bf 13G}$  by return e-mail confirmation.

Sincerely,

Jeffrey A. Ruiz

**Enclosures** 

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

SCHEDULE 13G (Amendment No. )

Under the Securities Exchange Act of 1934

Isis Pharmaceuticals Incorporated

NAME OF ISSUER:

Common Stock (\$0.001 Par Value)

TITLE OF CLASS OF SECURITIES

464330109

CUSIP NUMBER

December 31, 2002

(Date of Event Which Requires Filing of this Statement)

Check the appropriate box to designate the rule pursuant to which this

Schedule is filed:

[X] Rule 13d-1(b)

[ ]Rule 13d-1(c)

[ ]Rule 13d-1(d)

- 1. NAME OF REPORTING PERSONS
  - S.S. OR I.R.S. IDENTIFICATION NOS. OF ABOVE PERSONS

Deutsche Bank AG\*

- CHECK THE APPROPRIATE BOX IF A MEMBER OF A GROUP 2.
  - (A) [ ] (B) [ ]
- SEC USE ONLY
  - CITIZENSHIP OR PLACE OF ORGANIZATION

Federal Republic of Germany

NUMBER OF 5. SOLE VOTING POWER

**SHARES** 2,671,747

**BENEFICIALLY** SHARED VOTING POWER

OWNED BY 9,999

**EACH** SOLE DISPOSITIVE POWER

REPORTING 2,909,918

PERSON WITH 8. SHARED DISPOSITIVE POWER

223,500

9. AGGREGATE AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON

3,133,418

- CHECK BOX IF THE AGGREGATE AMOUNT IN ROW 9 EXCLUDES CERTAIN 10. SHARES [ ]
- PERCENT OF CLASS REPRESENTED BY AMOUNT IN ROW 9 11.

5.7%

12. TYPE OF REPORTING PERSON

HC, CO, BK

\* In accordance with Securities Exchange Act Release No. 39538 (January 12, 1998), this filing reflects the securities beneficially owned by the Private Clients and Asset Management business group ("PCAM") of Deutsche Bank AG and its subsidiaries and affiliates (collectively, "DBAG"). This filing does not reflect securities, if any, beneficially owned by any other business group of DBAG. Consistent with Rule 13d-4 under the Securities Exchange Act of 1934 ("Act"), this filing shall not be construed as an admission that PCAM is, for purposes of Section 13(d) under the Act, the beneficial owner of any securities covered by the filing.

	AME OF REPORTING PERSONS S.S. OR I.R.S. IDENTIFICATION NOS. OF ABOVE PERSONS				
	Deutsche	Asset Ma	nagement Europe GmbH		
2. CH	ECK THE APPR (A) [ ] (B) [ ]	OPRIATE B	OX IF A MEMBER OF A GROUP		
3. SE	3. SEC USE ONLY				
4. CITIZENSHIP OR PLACE OF ORGANIZATION					
F	ederal Repub	lic of Ge	rmany		
OWNED EACH REPOR	S ICIALLY BY	<ul><li>5.</li><li>6.</li><li>7.</li><li>8.</li></ul>	SOLE VOTING POWER  1,362,400 SHARED VOTING POWER  0 SOLE DISPOSITIVE POWER  1,362,400 SHARED DISPOSITIVE POWER  189,000		
9. AGGREGATE AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON					
	1,362,40	9			
10.	CHECK BOX IF THE AGGREGATE AMOUNT IN ROW 9 EXCLUDES CERTAI SHARES $[\ \ ]$				
11.	PERCENT OF	CLASS REP	RESENTED BY AMOUNT IN ROW 9		
	2.5%				

12. TYPE OF REPORTING PERSON

IC, CO

1. NAME OF REPORTING PERSONS S.S. OR I.R.S. IDENTIFICATION NOS. OF ABOVE PERSONS					
Deutsche	Bank Trus	st Company Americas			
2. CHECK THE APPROPRIATE BOX IF A MEMBER OF A GROUP  (A) [ ]  (B) [ ]					
3. SEC USE ONLY					
4. CITIZENSHIP OR PLACE OF ORGANIZATION					
Delaware					
SHARES BENEFICIALLY OWNED BY EACH REPORTING PERSON WITH  9. AGGREGATE AMOUNT	6. 7. 8. NT BENEFIC	SOLE VOTING POWER 857,847 SHARED VOTING POWER 9,999 SOLE DISPOSITIVE POWER 1,319,518 SHARED DISPOSITIVE POWER 0 CIALLY OWNED BY EACH REPORTING PERSON			
1,319,51	1,319,518				
10. CHECK BOX IF THE AGGREGATE AMOUNT IN ROW 9 EXCLUDES CERTAIN SHARES $[\ ]$					
11. PERCENT OF CLASS REPRESENTED BY AMOUNT IN ROW 9					
2.4%					
12. TYPE OF REPORTING PERSON					
BK, CO					

- 1. NAME OF REPORTING PERSONS
  - S.S. OR I.R.S. IDENTIFICATION NOS. OF ABOVE PERSONS

Deutsche Investment Management Americas Inc.

- 2. CHECK THE APPROPRIATE BOX IF A MEMBER OF A GROUP
  - (A) [ ] (B) [ ]
- 3. SEC USE ONLY
- 4. CITIZENSHIP OR PLACE OF ORGANIZATION

Delaware

NUMBER OF SOLE VOTING POWER 5.

**SHARES** 271,700 BENEFICIALLY

SHARED VOTING POWER 6.

OWNED BY

SOLE DISPOSITIVE POWER **EACH** 

**REPORTING** 48,200

PERSON WITH SHARED DISPOSITIVE POWER

223,500

9. AGGREGATE AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON

271,700

- 10. CHECK BOX IF THE AGGREGATE AMOUNT IN ROW 9 EXCLUDES CERTAIN SHARES [ ]
- 11. PERCENT OF CLASS REPRESENTED BY AMOUNT IN ROW 9

0.5%

12. TYPE OF REPORTING PERSON

IA, CO

- NAME OF REPORTING PERSONS
   S.S. OR I.R.S. IDENTIFICATION NOS. OF ABOVE PERSONS
   Deutsche Asset Management Group Ltd, London
- 2. CHECK THE APPROPRIATE BOX IF A MEMBER OF A GROUP
  - (A) [ ] (B) [ ]
- 3. SEC USE ONLY

BENEFICIALLY

4. CITIZENSHIP OR PLACE OF ORGANIZATION

United Kingdom

NUMBER OF 5. SOLE VOTING POWER SHARES 179,800

6. SHARED VOTING POWER

OWNED BY

EACH 7. SOLE DISPOSITIVE POWER

REPORTING 179,800

PERSON WITH 8. SHARED DISPOSITIVE POWER

0

9. AGGREGATE AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON

179,800

- 10. CHECK BOX IF THE AGGREGATE AMOUNT IN ROW 9 EXCLUDES CERTAIN SHARES  $[\ ]$
- 11. PERCENT OF CLASS REPRESENTED BY AMOUNT IN ROW 9

0.3%

12. TYPE OF REPORTING PERSON

IA, CO

Item 1(a). Name of Issuer:

Isis Pharmaceuticals, Inc. ("the Issuer")

Item 1(b). Address of Issuer's Principal Executive Offices:

The address of the Issuer's principal executive offices is:

2292 Faraday Avenue Carlsbad, CA 92008

Item 2(a). Name of Person Filing:

This statement is filed on behalf of Deutsche Bank AG, ("Reporting Person").

Item 2(b). Address of Principal Business Office or, if none, Residence:

The principal place of business of the Reporting Person is:

Taunusanlage 12, D-60325 Frankfurt am Main Federal Republic of Germany

Item 2(c). Citizenship:

The citizenship of the Reporting Person is set forth on the cover page.

Item 2(d). Title of Class of Securities:

The title of the securities is common stock, \$0.001 par value ("Common Stock").

Item 2(e). CUSIP Number:

The CUSIP number of the Common Stock is set forth on the cover page.

Item 3. If this statement is filed pursuant to Rules 13d-1(b), or

13d-2(b) or (c), check whether the person filing is a:

- (a) [] Broker or dealer registered under section 15 of the Act;
- (b) [X ] Bank as defined in section 3(a)(6) of the Act;

Deutsche Bank Trust Company Americas

- (c) []Insurance Company as defined in section 3(a)(19) of the Act;
- (d) [ X ] Investment Company registered under section 8 of the Investment Company Act of 1940;

Deutsche Asset Management Group Ltd, London

(e) [ X ] An investment adviser in accordance with Rule 13d-1(b)(1)(ii)(E);

Deutsche Investment Management Americas Inc.

Deutsche Asset Management Europe GmbH

- (f) [ ] An employee benefit plan, or endowment fund in accordance with Rule 13d-1 (b)(1)(ii)(F);
- (g) [X ] parent holding company or control person in accordance with Rule 13d-1 (b)(1)(ii)(G);

Deutsche Bank AG

(h) [ ] A savings association as defined in section 3(b) of the Federal Deposit Insurance Act;

- (i) [ ] A church plan that is excluded from the definition of an investment company under section 3(c)(14) of the Investment Company Act of 1940;
- (j) [] Group, in accordance with Rule 13d-1 (b)(1)(ii)(J).

#### Item 4. Ownership.

(a) Amount beneficially owned:

The Reporting Person owns the amount of the Common Stock as set forth on the cover page.

(b) Percent of class:

The Reporting Person owns the percentage of the Common Stock as set forth on the cover page.

- (c) Number of shares as to which such person has:
  - (i) sole power to vote or to direct the vote:

The Reporting Person has the sole power to vote or direct the vote of the Common Stock as set forth on the cover page.

(ii) shared power to vote or to direct the vote:

The Reporting Person has the shared power to vote or direct the vote of the Common Stock as set forth on the cover page.

(iii) sole power to dispose or to direct the
disposition of:

The Reporting Person has the sole power to dispose or direct the disposition of the Common Stock as set forth on the cover page.

(iv) shared power to dispose or to direct the
disposition of:

The Reporting Person has the shared power to dispose or direct the disposition of the Common Stock as set forth on the cover page.

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

Not applicable.

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

Not applicable.

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

Subsidiary Item 3 Classification

Deutsche Bank Trust Company Americas Bank

Deutsche Investment Management Americas Inc. Investment Advisor

Deutsche Asset Management Europe GmbH Investment Company

Deutsche Asset Management Group Ltd, London Investment Company

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

Not applicable.

Item 9. Notice of Dissolution of Group.

Not applicable.

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.

### SIGNATURE

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

DEUTSCHE BANK AG

By: /s/ Jeffrey A. Ruiz Name: Jeffrey A. Ruiz Title: Vice President

By: /s/ Margaret M. Adams Name: Margaret M. Adams

Title: Director

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.

Deutsche Asset Management Europe Gmbh

By: /s/ Michaela Bundschuh Name: Michaela Bundschuh

Title: Head of Global Position Monitoring

By: /s/ Susan Seidel Name: Susan Seidel Title: Vice President

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

Deutsche Bank Trust Company Americas

By: /s/ James T. Byrne, Jr. Name: James T. Byrne, Jr. Title: Secretary

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

Deutsche Investment Management Americas Inc.

By: /s/ William G. Butterly III Name: William G. Butterly III Title: Secretary

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

Deutsche Asset Management Group Ltd, London

By: /s/ Patrick Galligan Name: Patrick Galligan

Title: Director

By: /s/ Hugh Male Name: Hugh Male Title: Vice President