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To The Honorable Commissioner of Patents and Trademarks. Please record the attached documents or copy thereof.

## 1. Name of conveying party(ies):

Ian Popoff, Vickie L. Brown-Driver and Lex M. Cowser

Additional name(s) of conveying party(ies)  
attached? ( ) YES (X) NO

## 3. Nature of Conveyance:

- (X) Assignment ( ) Merger  
 ( ) Security Agreement  
 ( ) Change of Name  
 ( ) Other:

Execution Date: Feb. 24, 2000

## 2. Name and address of receiving party(ies):

Name: Isis Pharmaceuticals Inc.

Internal Address:

Street Address: 2922 Faraday Ave

City Carlsbad

State California

Zip 92008

Additional name(s) &amp; address attached?

( ) YES (X) NO



## 4. Application number(s) or patent number (s): N/A

If this document is being filed together with a new application, the execution date of the application is: 2/24/2000

A. Patent Application No.(s) Not yet assigned

B. Patent No.(s) N/A

Additional numbers attached? ( ) YES (X) NO

5. Name and address of party to whom  
correspondence concerning documents should be  
mailed:

Name: Jane Massey Licata

Address: Law Offices of Jane Massey Licata

66 East Main Street

Marlton NJ 08053

6. Total number of applications and patents  
involved: 1

7. Total fee (37 CFR 3.41):.....\$40.00

( ) Enclosed

(X) Authorized to be charged to deposit account

8. Deposit account number: 500252

(Attach duplicate copy of this page if paying by  
deposit account)

03/28/2000 JSHABAZZ 00000160 500252 09517584

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## 9. Statement and signature.

To the best of my knowledge and belief, the foregoing information is true and correct and any attached copy is  
a true copy of the original document.

Laurel Spear Bernstein /

Name of Person Signing/Reg.No. 37,280

Signature

March 2, 2000  
Date

Total number of pages including cover sheet, attachments, and document:: 4

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 Management and Budget, Paperwork Reduction Project. (0651-0011).

PATENT

REEL: 010636 FRAME: 0586

A S S I G N M E N T

WHEREAS, we Ian Popoff, residing at 904 Hygeia Ave Encinitas CA 92024, Vickie L. Brown-Driver, residing at 1015 Santa Florencia Solana Beach CA 92075 and Lex M. Cowsert, residing at 3008 Newshire Street Carlsbad CA 92008 hereinafter referred to as the assignors, are the joint inventors of certain inventions or improvements for which we have made application for Letters Patent to the United States, identified as Case No. RTS-0121 entitled **Antisense Modulation of E2F transcription factor 1 Expression**; filed concurrently herewith; and

WHEREAS, **Isis Pharmaceuticals Inc.** hereinafter referred to as the assignee, of **2922 Faraday Ave, Carlsbad, California 92008** is desirous of acquiring the entire right, title and interest in and to the said inventions or improvements and in and to the said application, and in, to and under any and all Letters Patent which may be granted on or as a result thereof in any and all countries:

NOW, THEREFORE, for and in consideration of the sum of One Dollar (\$1.00) to each of us in hand paid by said assignee, and other good and valuable consideration, the receipt of which is hereby acknowledged, we, the said assignors, have sold, assigned, transferred and set over, and by these presents do hereby sell, assign, transfer and set over to said assignee, the entire right, title and interest in and to said inventions or improvements and said application and any and all continuations, divisions and renewals of and substitutes for said application, and in, to and under any and all Letters Patent which may be granted on or as a result thereof in the United States and any and all other countries, and any reissue or reissues or extension or extensions of said Letters Patent, and assign to and authorize said assignee, to file in our names applications for Letters Patent in all countries, the same to be held and enjoyed by said assignee, its successors, assigns, nominees or legal representatives, to the full end of the term or terms for which said Letters Patent respectively may be granted, reissued or extended, as fully and entirely as the same would have been held and enjoyed by us had this assignment, sale and transfer not been made.

AND we hereby covenant that we have full right to convey the entire interest herein assigned, and that we have not executed and will not execute any agreement in conflict herewith, and we further covenant and agree that we will each time request is made and without undue delay, execute and deliver all such papers as may be necessary or desirable to perfect the title to said inventions or improvements, said application and said Letters Patent to said assignee, its successors, assigns, nominees, or legal representatives, and each of us agrees to communicate to said assignee or to its nominee all known facts respecting said inventions or improvements, said application and said Letters Patent, to testify in any legal proceedings, to sign all lawful papers, to execute all disclaimers and divisional, continuing, reissue and foreign applications, to make all rightful oaths, and generally to do everything possible to aid said assignee, its successors, assigns, nominees and legal representatives to obtain and enforce for its or

their own benefit proper patent protection for said inventions or improvements in any and all countries.

AND we hereby authorize and request the Commissioner of Patents and Trademarks of the United States and any official of any country or countries foreign to the United States whose duty it is to issue patents on applications as aforesaid, to issue to said assignee, as assignee of the entire right, title and interest, any and all Letters Patent for said inventions or improvements, including any and all Letters Patent of the United States which may be issued and granted on or as a result of the application aforesaid, in accordance with the terms of this assignment.

We further authorize and direct our attorneys to insert below\* the serial number and filing date of said application now identified as Case No. RTS-0121 as soon as the same shall have been made known to them by the United States Patent Office.

IN WITNESS WHEREOF, we have hereunto set our hands and seals.

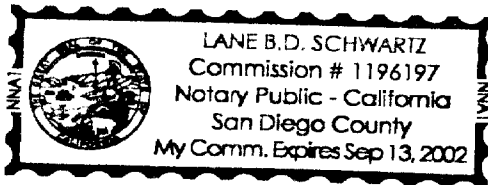
Dated:

1/28/00

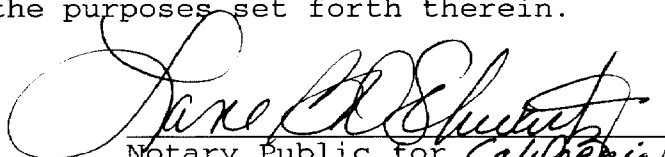
  
Ian Popoff

STATE OF California )  
COUNTY OF San Diego ) ss.

This 28<sup>th</sup> day of January, 2000, before me personally came the above-named Ian Popoff, who executed the foregoing Assignment in my presence, and who acknowledged to me that he/she executed the same of his/her own free will for the purposes set forth therein.



[SEAL]

  
Notary Public for California  
My commission expires: 9-13-2002

Dated: 2-24-00

Vickie L. Brown-Driver  
Vickie L. Brown-Driver

STATE OF California,  
COUNTY OF San Diego )  
SS.

This 24 day of February 2000, before me personally came the above-named Vickie L. Brown-Driver, who executed the foregoing Assignment in my presence, and who acknowledged to me that he/she executed the same of his/her own free will for the purposes set forth therein.



Elana Guglielmo  
Notary Public for California  
My commission expires: 11/23/01

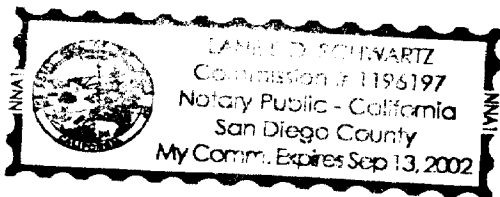
[SEAL]

Dated: 2/8<sup>th</sup>/00

Lex M. Cowser  
Lex M. Cowser

STATE OF California,  
COUNTY OF San Diego )  
SS.

This 9<sup>th</sup> day of February, 2000, before me personally came the above-named Lex M. Cowser, who executed the foregoing Assignment in my presence, and who acknowledged to me that he/she executed the same of his/her own free will for the purposes set forth therein.



Daniel D. Schwartz  
Notary Public for California  
My commission expires: 09-13-02

[SEAL]