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02-25-2003

OMB No. 0651-0027 (exp. 6/30/2005)



Tab Settings

To the Honorable Commissioner of Patents and Trademarks

102372409

Final documents or copy thereof.

1. Name of conveying party(ies):

Lex M. Cowser, Jacqueline Wyatt, Susan Freier, Brett P. Monia, Madeline M. Butler, and Robert McKay

Additional name(s) of conveying party(ies) attached? yes no

2. Name and address of receiving party(ies)

Name: ISIS Pharmaceuticals, Inc.

Internal Address: _____

Street Address: 2292 Faraday Avenue

City: Carlsbad State: CA Zip: _____

Additional name(s) & address(es) attached? yes no

3. Nature of Conveyance:

- Assignment Merger
- Security Agreement Change of Name
- Other _____

Execution Date: October 30, 2000; November 1, 2000; October 30, 2000; November 15, 2000; October 30, 2000; and October 27, 2000

4. Application number(s) or patent number(s):

If this document is being filed together with a new application, the execution date of the application is _____

A. Patent Application No.(s)

09/629,644

B. Patent No.(s)

Additional numbers attached? yes no

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

Name: Mary E. Bak, Esquire

Howson and Howson

Internal Address: _____

Street Address: Spring House Corporate Center,
Box 457

City: Spring House State: PA Zip: 19477

6. Total number of applications and Patents involved: _____

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

Enclosed

Authorized to be charged to deposit account

8. Deposit account number:

08-3040

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

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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.

Mary E. Bak

Name of Person Signing

Mary E. Bak
Signature

Feb 13, 2003

Date

Total number of pages, including cover sheet, attachments, and document: 7

Mail documents to be recorded with required coversheet information to:
Commissioner of Patents and Trademarks, Box Assignments
Washington, DC 20231

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PATENT

REEL: 013769 FRAME: 0250

A S S I G N M E N T .

WHEREAS, We, Lex M. Cowsert, of Carlsbad, State of California; Jacqueline Wyatt, of Encinitas, State of California; Susan M. Freier, of San Diego, State of California; Brett P. Monia, of La Costa, State of California; Madeline M. Butler, of Rancho Santa Fe, State of California and Robert McKay, of San Diego, State of California, have invented a certain invention entitled Antisense Modulation of PTP1B Expression, for which we have made application for Letters Patent of the United States, said application having been filed July 31, 2000 and assigned Application No. 09/629,644; and

WHEREAS, ISIS Pharmaceuticals, Inc. hereinafter referred to as the assignee, of 2292 Faraday Avenue, Carlsbad CA, a corporation of Delaware, 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

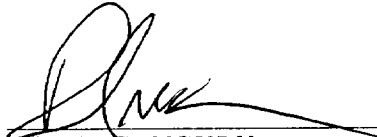
ASSIGNMENT - PAGE 1 OF 4

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.

ASSIGNMENT - PAGE 2 OF 4

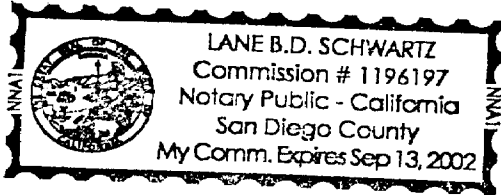
DATED: 10/27, 2000

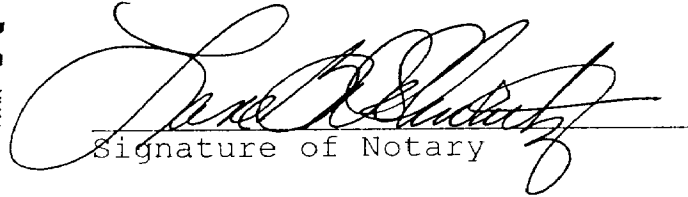

ROBERT MCKAY

State of California)
County of San Diego) ss.

On this 27th day of October, 2000, before me, the undersigned, a Notary Public for the state, personally appeared Robert McKay, personally known to me/~~proved to me on the basis of satisfactory evidence~~ to be the person who subscribed the foregoing Assignment and acknowledged to me that he executed the same in his authorized capacity, and that by his signature on the Assignment the person, or the entity on behalf of which the person acted, executed the Assignment.

WITNESS my hand and official seal.




Signature of Notary