To The Honorable Commissioner of Patents and Trademarks. Please record the attached original documents or copy thereof.				
	18-02		ess of receiving party(ies):	
Douglas L. Cole	02/20/97	Name: ISIS Pharmad	ceuticals, Inc.	
First Assignor	Date of Execution	Street Address: 229	22 Faraday Ayanya	
	02/06/97		22 I diaday Aveilue	
Second Assignor	Date of Execution	City: Carlsbad		
Third Assistance		State: California	ZIP 92008	
Third Assignor	Date of Execution	Additional name(s)	& address attached? ( )YES (X)NO	
Fourth Assignor	Date of Execution	3. Nature of Conve	eyance (check only one):	
		() New Assignme		
Fifth Assignor	Date of Execution	( ) License Agree ( ) Change of Na		
Sixth Assignor	Date of Execution	Application N	ce of Assignment filed in United States Io.08/775,011 recorded on 04/28/00 at 90 and Frame No. 0221	
Additional name(s) of conveying party(ies) attached	d? () YES (X) NO			
4. Identification of Application number(s), patent number(s) OR date of execution of application to which assignment refers:				
Application Nos.: 09/756,354 filed 01/08/01	Patent Nos.:		Date application was signed by the first named executing inventor	
			(Fill-in this date only if assignment is filed together with a new application)	
5. Name and address of party to whom correspondence concerning documents should be mailed:		6. Total number of applications and patents involved: 1		
Name: Michael P. Straher, Esq. WOODCOCK WASHBURN LLP		7. Total fee (37 CFR 3.41):\$40.00 () Enclosed		
One Liberty Place - 46th Floor Philadelphia, Pennsylvania 19103-7301 215-568-3100		() Authorized to be charged to Deposit Account Number 23-3050		
		Please charge any deficiency or credit any overpayment to Deposit Account Number 23-3050		
DO NOT USE THIS SPACE				
9. Statement and signature.				
To the best of my knowledge and belief, the	foregoing information	on is true and correct	and any attached copy is a true copy of	
the original document.	11.1.	Gin	1.41.21.30	
Michael P. Straher/38,325  Name of Person Signing/Reg. No.	, c./ felli	ature	October 26, 2001 Date	
Total number of pages including cover sheet, assignment document and additional pages attached thereto: 5				
Mail documents to be recorded with required cover sheet information to:				
Commissioner of Patents and Trademarks  Box Assignments				
Washington, D.C. 20231				

(Pending Application; Serial No. Known)

DOCKET NO.: ISIS-2266

## ASSIGNMENT

WHEREAS, we, Douglas L. Cole and Vasulinga Ravikumar, hereinafter referred to as the assignors, residing respectively at 4992 Smith Canyon Court, San Diego, California 92130 and 6606 Vireo Court, Carlsbad, California 92009 are the joint inventors of certain inventions or improvements for which we have made application for Letters Patent to the United States, identified as Serial No. 08/775,011, filed December 27, 1996, entitled METHOD OF SYNTHESIZING PHOSPHOROTHIOATE OLIGONUCLEOTIDES; 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 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

PATENT REEL: 012518 FRAME: 0819

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.

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

DOUGLAS L. COLE

V. T. Raview 2/1/97 (L.S.)

VACIU THEA DAVIVIMAD

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[WHERE ALL INVENTORS SIGN BEFORE THE SAME NOTARY:]
SS COUNTY OF:
On this day of, 19, before measurements came the above named <u>Douglas L. Cole and Vasulings Ravikumar</u> to me personally known and known to me to be the same individual who executed the foregoing assignment, and who acknowledged to me that execution of the same was of that person's own free will for the use and purposes therein set forth.
Notary Public
[WHERE INVENTORS SIGN BEFORE SEPARATE NOTARIES: ]
State of California:  COUNTY OF San Diego:  SS
on this <u>lom</u> day of <u>february</u> , 19 <u>7</u> , before m personally came the above named <u>Douglas L. Cole</u> to me personall known and known to me to be the same individual who executed th foregoing assignment, and who acknowledged to me that execution o the same was of that person's own free will for the use an purposes therein set forth.
CHARMAINE G. WEBB  COMM. # 1008645  Notary Public — California SAN DIEGO COUNTY My Comm. Expires NOV 7, 1997  Notary Public

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	Joint Inventors
State of California	:
State of California	SS:
personally came the all personally known and known executed the foregoing as	day of <u>Yestery</u> , 19—7, before me cove named <u>Vasulinga Ravikumar</u> to me cown to me to be the same individual who signment, and who acknowledged to me that s of that person's own free will for the set forth.
CHARMAINE G. WEBB COMM. # 1008645  Notary Public — Catifornia SAN DIEGO COUNTY My Comm. Expires NOV 7, 1997	Motary Public
<u> </u>	
COUNTY OF	ss .:
personally came the above me personally known and be executed the foregoing as	known to me to be the same individual who ssignment, and who acknowledged to me that as of that person's own free will for the
	Notary Public

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**RECORDED: 01/18/2002** 

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PATENT REEL: 012518 FRAME: 0822