## Research Portfolio Online Reporting Tools (RePORT)

# Project Information @

1R21HG003977-01

### DESCRIPTION DETAILS RESULTS HISTORY SUBPROJECTS

Project Number: 1R21HG003977-01

EVOLVING NOVEL POLYMERASES FOR GENOME

 Contact PI / Project Leader:
 ROMESBERG, FLOYD E.

 Awardee Organization:
 SCRIPPS RESEARCH INSTITUTE

### Abstract Text:

Title:

DESCRIPTION (provided by applicant): There is great incentive for developing low cost technologies for sequencing genomes. While significant advances have been made in conventional sequencing methods, the reductions in cost and time required to make genome sequencing more feasable require the development of new methodologies. Indeed many new and promising methods are under development. However, these new approaches are generally limited by the inability of DNA polymerases to synthesize DNA with modified deoxynucleotide triphosphates (dNTPs). We propose to use our recently developed activity-based selection system to evolve DNA polymerases that efficiently recognize these modified dNTPs, and thus overcome one of the most significant barriers to the development of the novel sequencing methods. The selection system is based on the co-display on phage of DNA polymerase libraries and their DNA substrates, such that mutants that recognize the modified dNTPs covalently attach a biotin-dUTP, allowing for their selective isolation. In addition to the selections, various modern gene shuffling approaches are proposed to generate polymerase diversity. The Specific Aims of the proposal are: Specific Aim 1: Evolve DNA polymerases that efficiently synthesize DNA using fluorophore-labeled dNTPs to enable novel synthesis-by-sequencing methodologies. Specific Aim 2: Evolve polymerases that efficiently incorporate reversibly fluorophore tagged dNTPs with 3'-O-allyl protecting groups and then continue synthesis after double deprotection. Specific Aim 3: Collaborate to evaluate evolved polymerases in actual sequencing applications, rho evolution of polymerases better suited for sequencing applications, and their evaluation in the actual sequencing applications, promises to dramatically facilitate the development of the methods and help make genome sequencing more commonplace.

#### Project Terms:

bacterial virus; biotin; directed evolution; DNA; DNA directed DNA polymerase; fluorescent dye /probe; genome; ionophores

Contact PI Infor	mation:	Program Officia Information:		Other PI Informati	on:	
Name: ROMESB Email: <u>Click to vir</u> address Title: PROFESS(	ew contact PI email	Name: SCHLOS Email: <u>Click to vi</u> address		Not Applicable		
Organization:				artment / Educationa itution Type:	al Coi	ngressional District:
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Other Informatio	on:					
(10)B)	<u>4-002</u> Special Emphasis Par 6 <b>Award Notice Date</b>		Project Star	ber: 781613492 rt Date: 19-MAY-2006 rt Date: 19-MAY-2006		a: 172 d Date: 30-APR-2008 d Date: 30-APR-2007
Administering In	nstitutes or Centers:					
NATIONAL HUM	AN GENOME RESEA	RCH INSTITUTE				
Project Funding	Information for 200	6:				
Total Funding: \$	278,850					
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2006	NATIONA INSTITUT	L HUMAN GENOM E	E RESEARCH	l \$278,850		
History:						
Project	Sub # Project Title	Prir	ntact ncipal estigator	Organization	FY Admin IC	Funding FY Total Cost IC by IC
5R21HG003977- 02	EVOLVING NO POLYMERASE GENOME SEQ	S FOR FUC	MESBERG, DYD E.	SCRIPPS RESEARCH INSTITUTE	2007 NHGRI	NHGRI \$225,636
1R21HG003977- 01	EVOLVING NO POLYMERASE GENOME SEQ	SFOR FLC	MESBERG, DYD E.	SCRIPPS RESEARCH INSTITUTE	2006 NHGRI	Columbia Exh

Columbia Exhibit 2103 Illumina, Inc. v. The Trustees of Columbia University in the City of New York IPR2018-00291, 318, 322, 385 & 797

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 Contact PI / Project Leader:
 ROMESBERG, FLOYD E.

 Awardee Organization:
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Contact PI Info	rmation:	Program Officia Information:	1	Other PI Informati	on:		~	
	BERG, FLOYD E. <u>view contact PI email</u> SOR	Name: SCHLOS Email: <u>Click to vi</u> address		Not Applicable				
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Other Informati	on:							
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Project Number	Sub # Project Title	Pri	ntact ncipal estigator	Organization	124	Admin IC	Funding IC	FY Total Cost by IC
5R21HG003977- 02	EVOLVING NO POLYMERASE GENOME SEC	S FOR	MESBERG, DYD E.	SCRIPPS RESEARCH INSTITUTE	2007	NHGRI	NHGRI	\$225,636
1R21HG003977- 01	EVOLVING NO POLYMERASE GENOME SEC	SFOR FOR	MESBERG, DYD E.	SCRIPPS RESEARCH INSTITUTE	2006	NHGRI	NHGRI	\$278,850

Exhibit: <u>2 103</u> Witness: <u>Ronesberg</u> Date: <u>16-9-07</u>

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Project Number Sub # Project Title	Principal	Organization	FY Admin IC	Tota
	Investigator			by I

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