

# iGEM and Synthetic Biology

*Opportunities for Canada*

Andrew Hessel

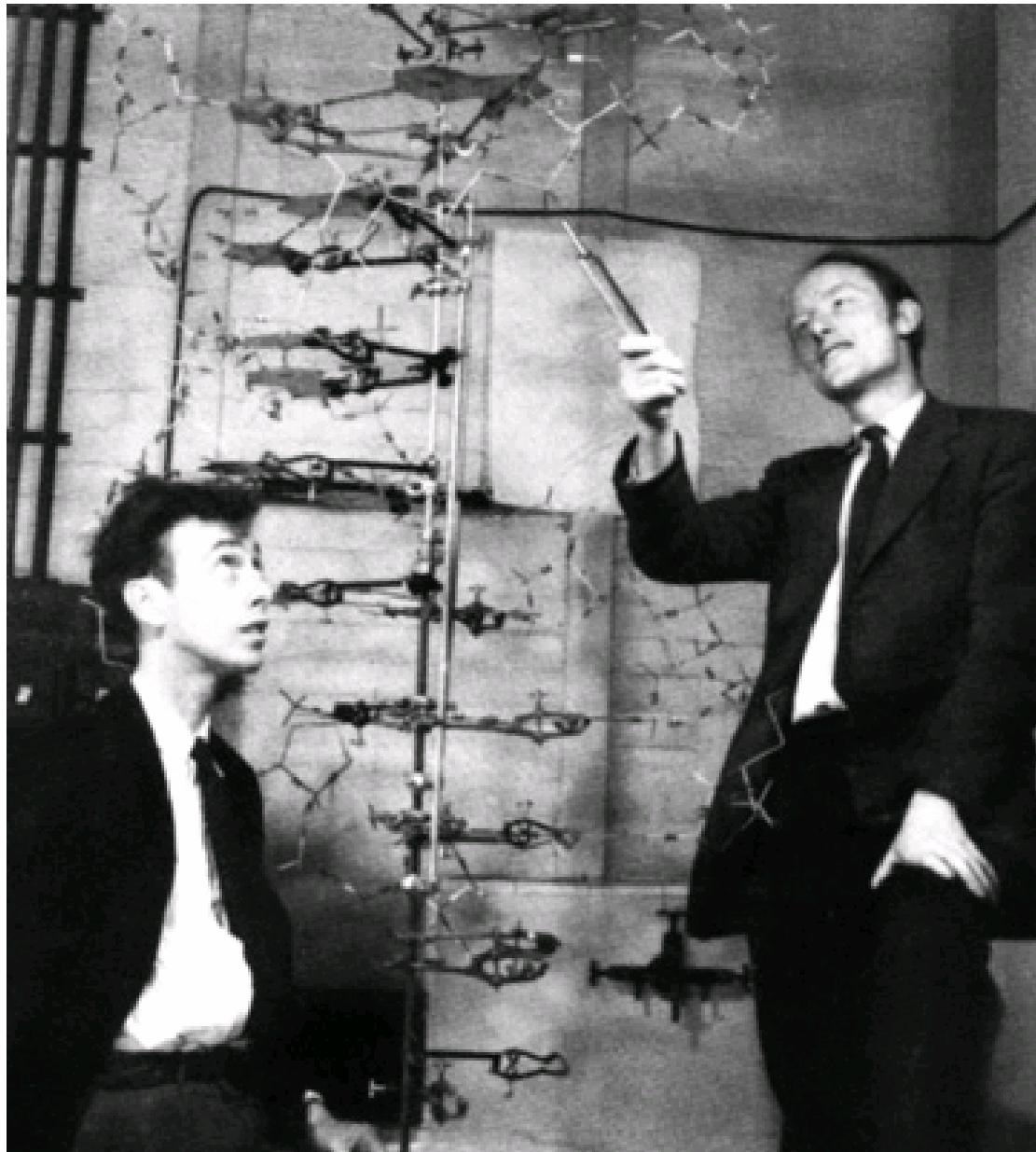
[ahessel@gmail.com](mailto:ahessel@gmail.com)

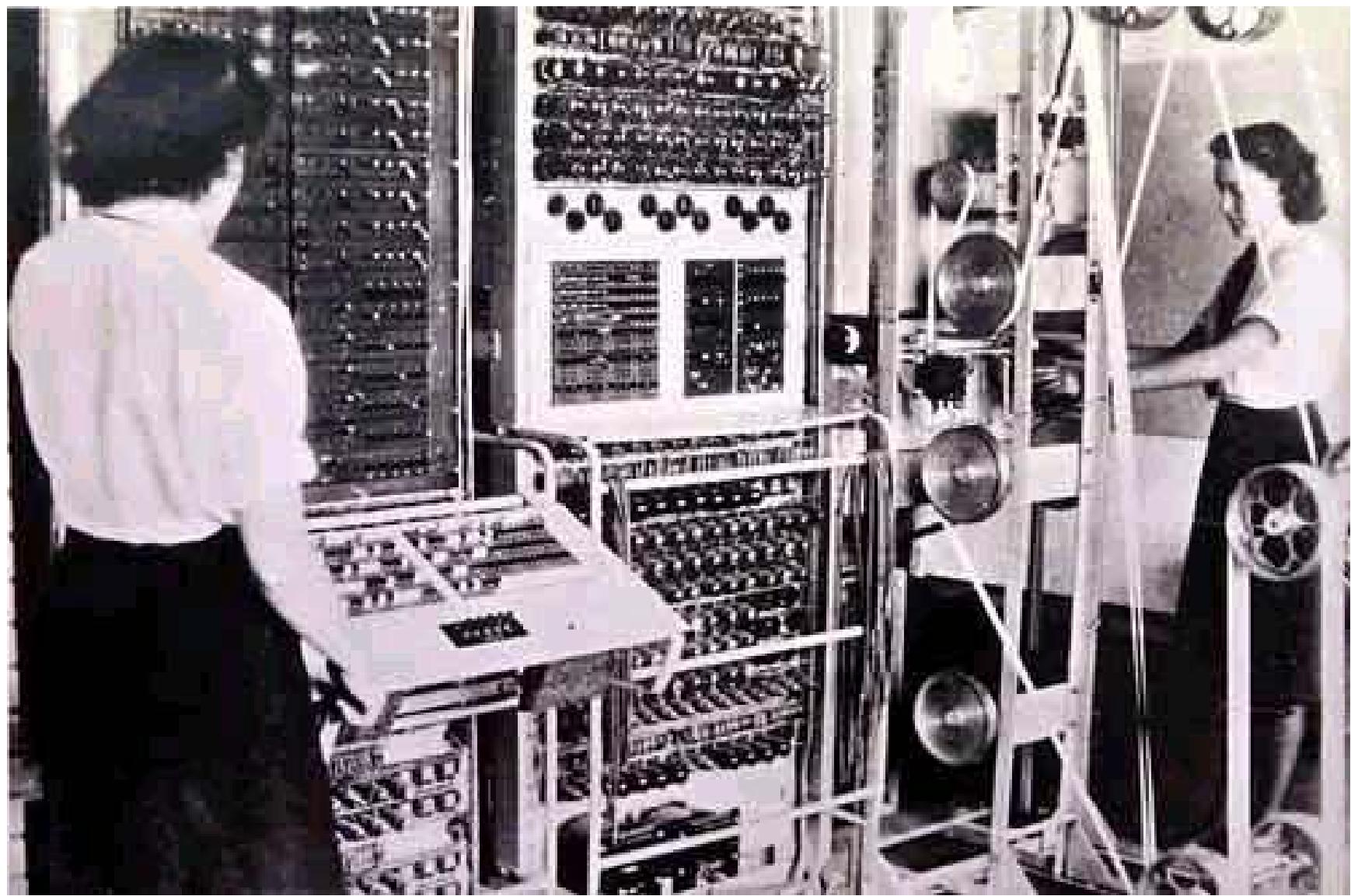
University of Ottawa. November 23 2007

**Synthetic biology is a basket of technologies that facilitate the engineering of living organisms**

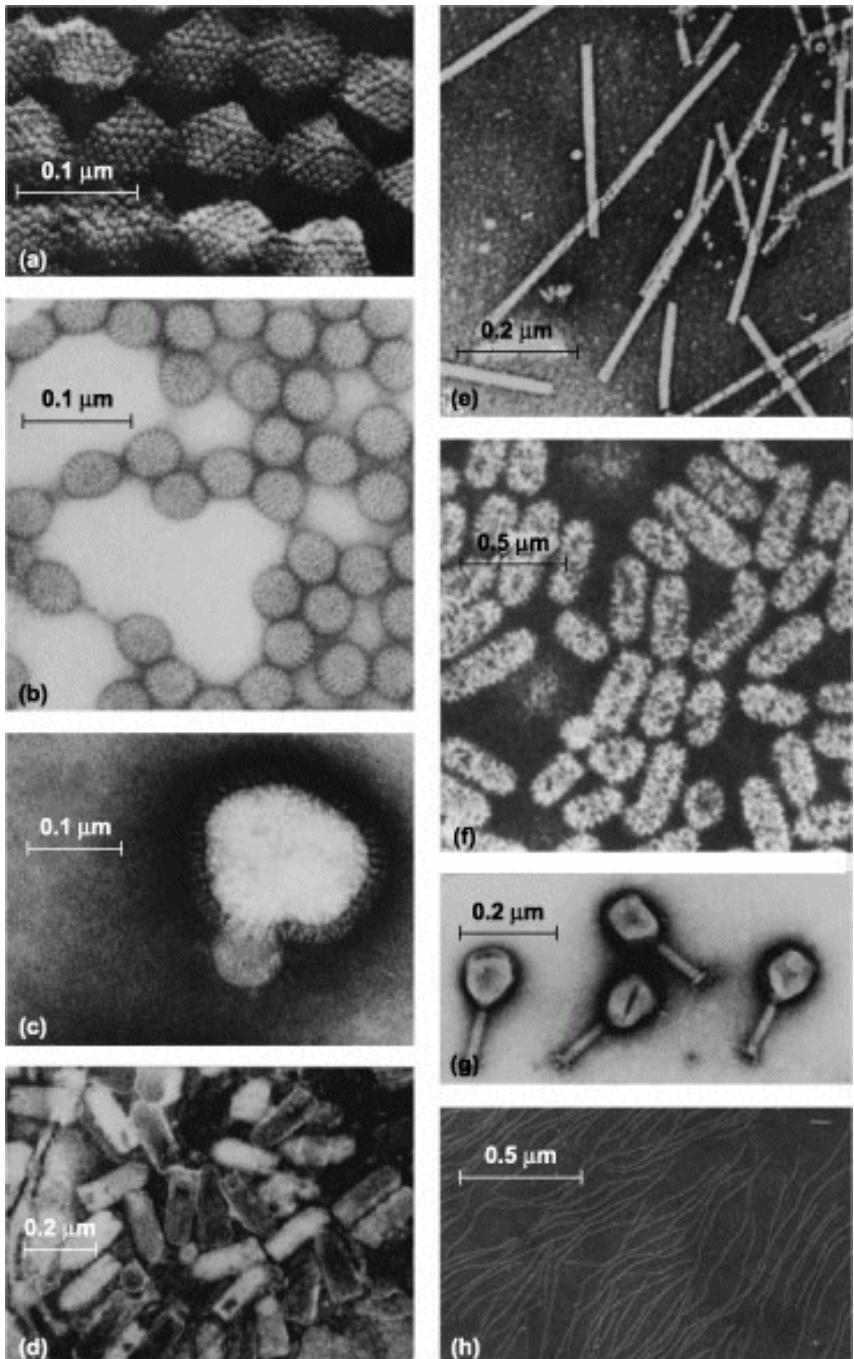
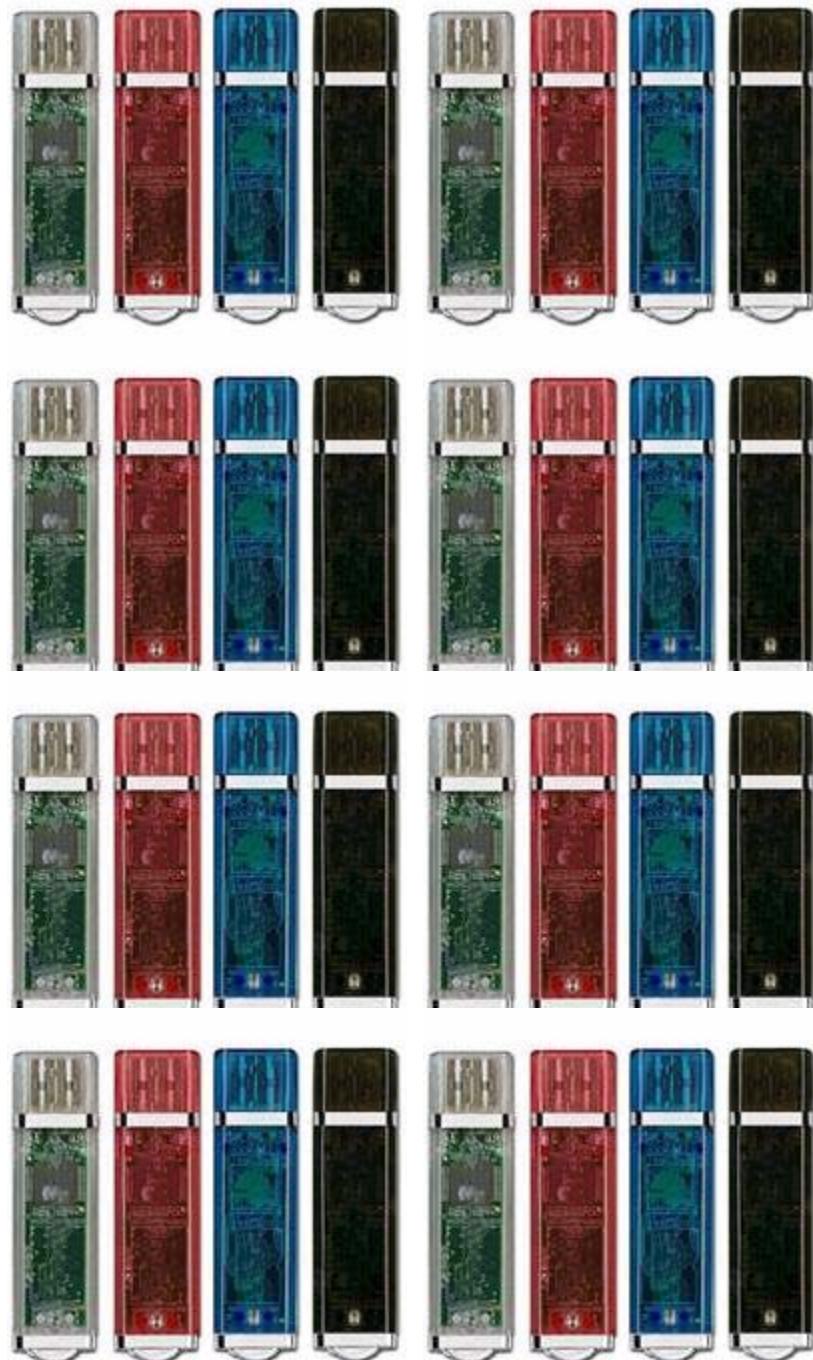
**Potentially the single most important technology of this century...**











Bits (0s,1s) encode information for digital electronic processors (computers)

DNA encodes information for analog biochemical processors (cells)

DNA is source code for biological systems

**Change the code, change metabolism**

# Genomes

are special programs that encode:

- Instructions for biochemical processors
- Machinery to duplicate the processor and install the program onto processors

*Equivalent in computing not to an operating system but to all the software it would take to run a computer production line.*



danielig 072006 - S3is

C:\Command.com

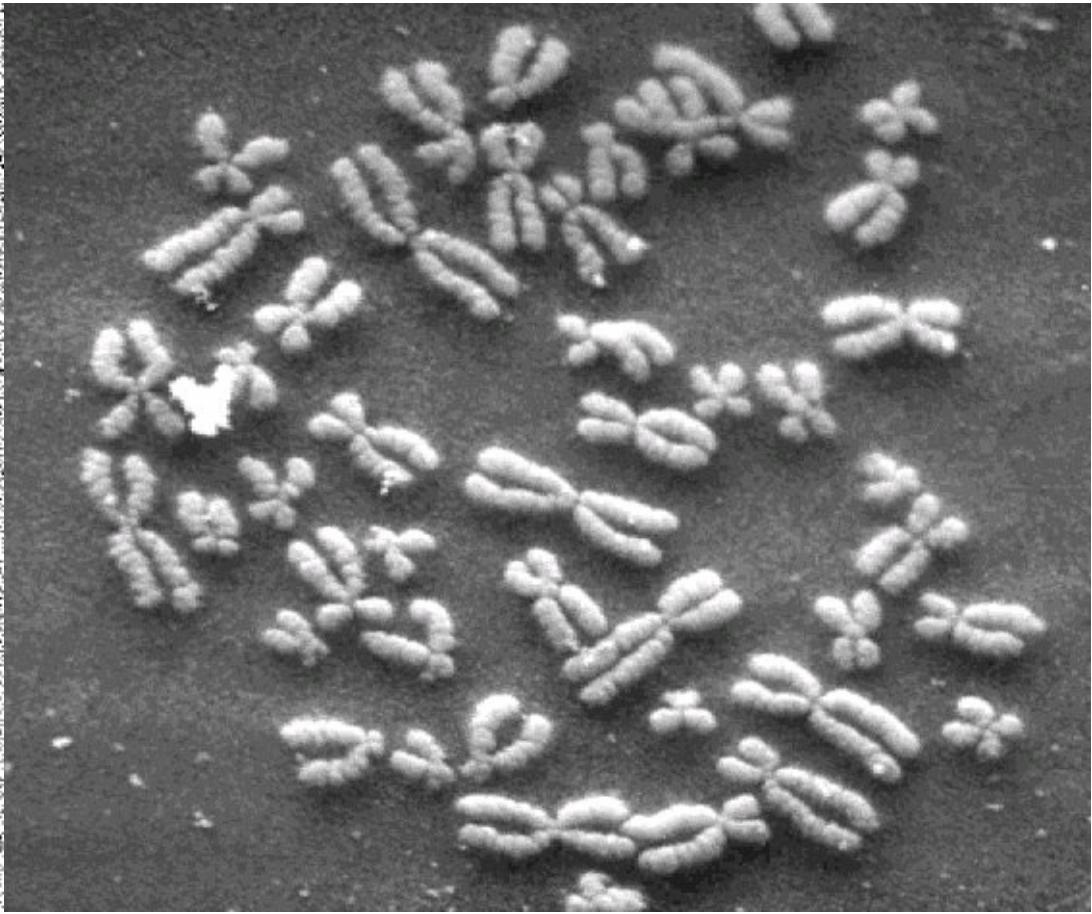
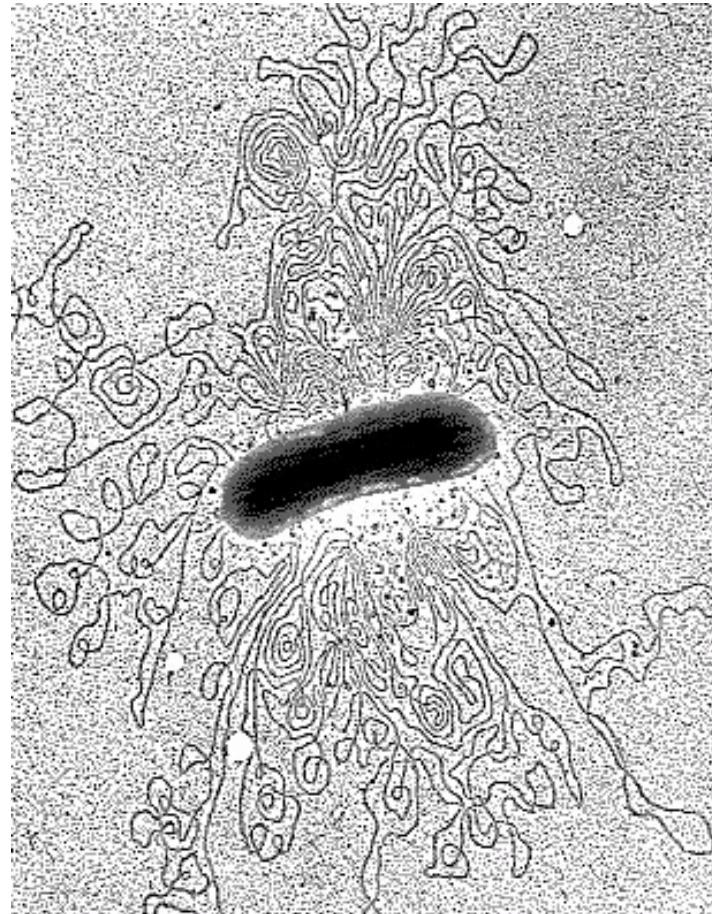
File Edit View Insert Options Window Help

16 32 64 1 2 4 H D N

Address	Value	Hex	ASCII
0x000000	4D5A	7400 B800 0000 2604 0100 FFFF 0000	MZt.,...&...ÿÿ..
0x000100	0000 0001	0000 1E00 0000 0100 0000	.....
0x000200	0000 0000	0000 0000 0000 0000 0000	.....
0x000300	0000 0000	0000 0000 0000 0000 0000	.....
0x000400	2433 2433	454E 5501 00B5 014E 5343 4F00	\$3\$3ENU..µ.NSC0.
0x000500	0306 0016	001A 0001 04FF FF48 0302 0056	.....ÿÿH...v
0x000600	0303 0061	0308 006C 0302 01FF FF7A 0303	...a...l...ÿÿz..
0x000700	CCEA 0383	03EB 03A4 03EC 03BC 03ED 03D2	íê.f.ë.¤.i.¤.i.º
0x000800	03EE 03ED	03EF 0313 04F0 032A 04F1 033C	.i.i.i...ð.*.ñ.<
0x000900	04F2 034D	04F3 0369 04F4 0380 04F5 03B6	.ò.M.ó.i.ó.□.ð.¶
0x000A00	04F6 03CD	04F7 03D8 04F8 03F8 04F9 031E	.ð.Í.÷.§.§.§.ù..
0x000B00	05FA 033F	05FB 034F 05FC 0357 05FD 0363	.ú.?..û.0.û.º.ý.c

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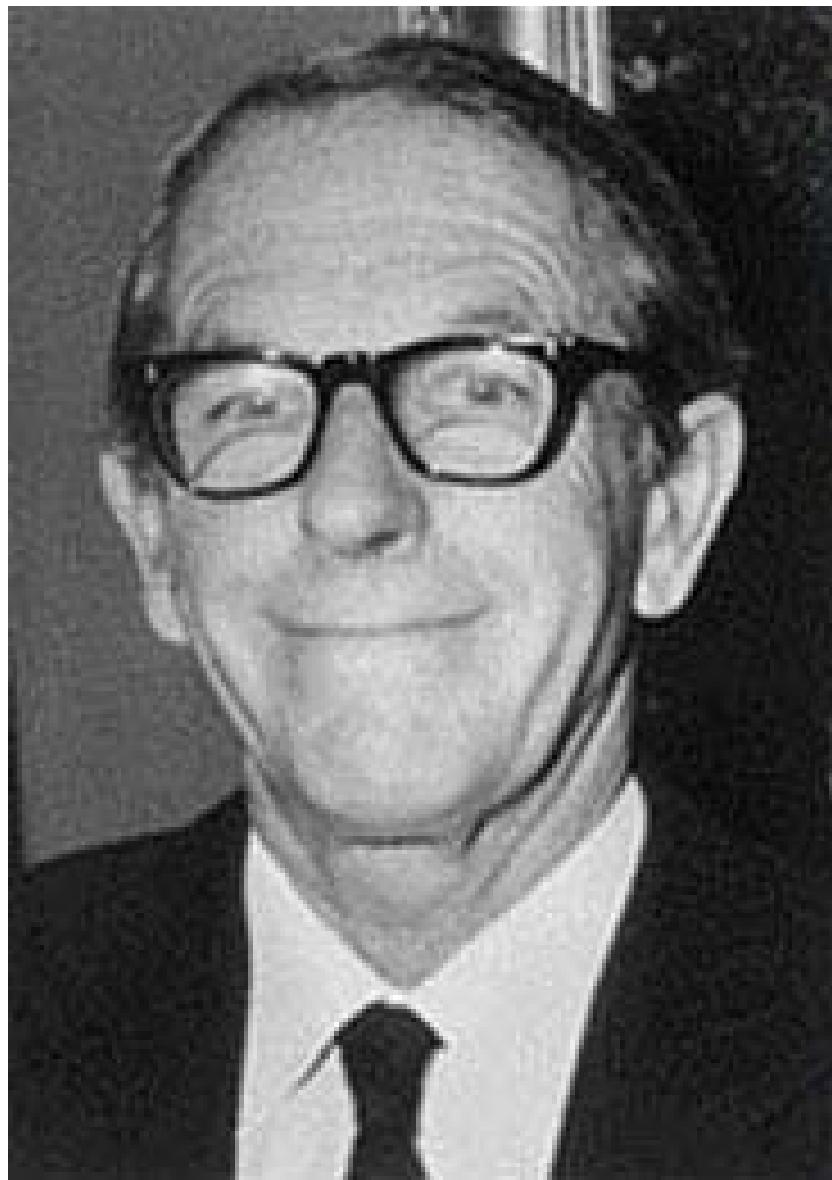
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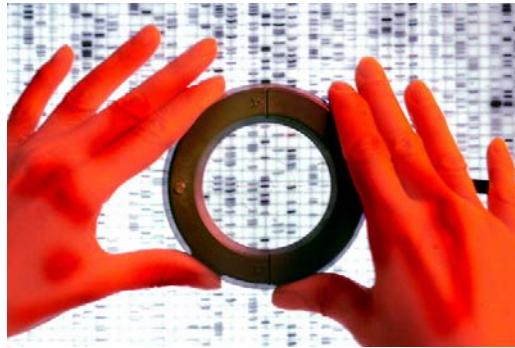
<http://www.colorado.edu/MCDB/MCDB111/intrographics/bacterial%20chromosome.jpg>

<http://academy.asd20.org/kadets/lundberg/ethics/images/chromosomes.jpg>

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GTT-CCT-GCC-ACA-ATA-ATT-TGT-TTA-GAG-ACC-CCT-TGG-GAA-GAA-TGA



Nobel foundation



1980  
500 bp/day (manual)



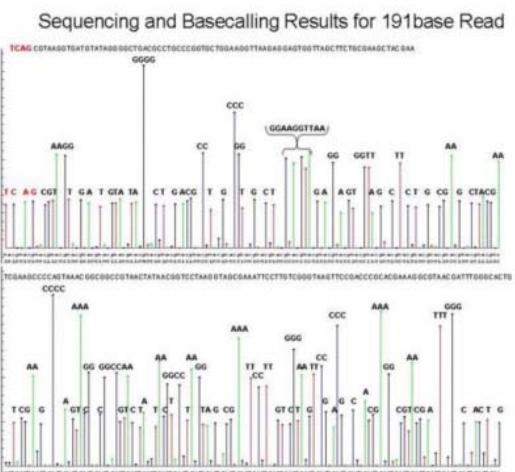
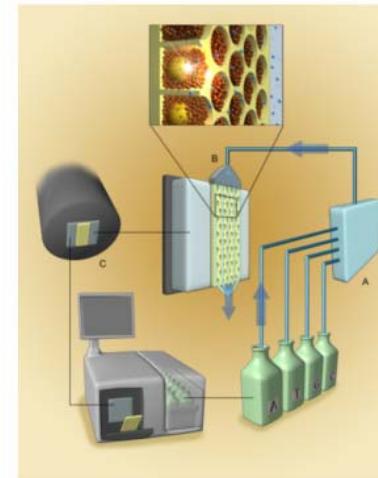
1987  
36,000 bp/day (semi-auto)



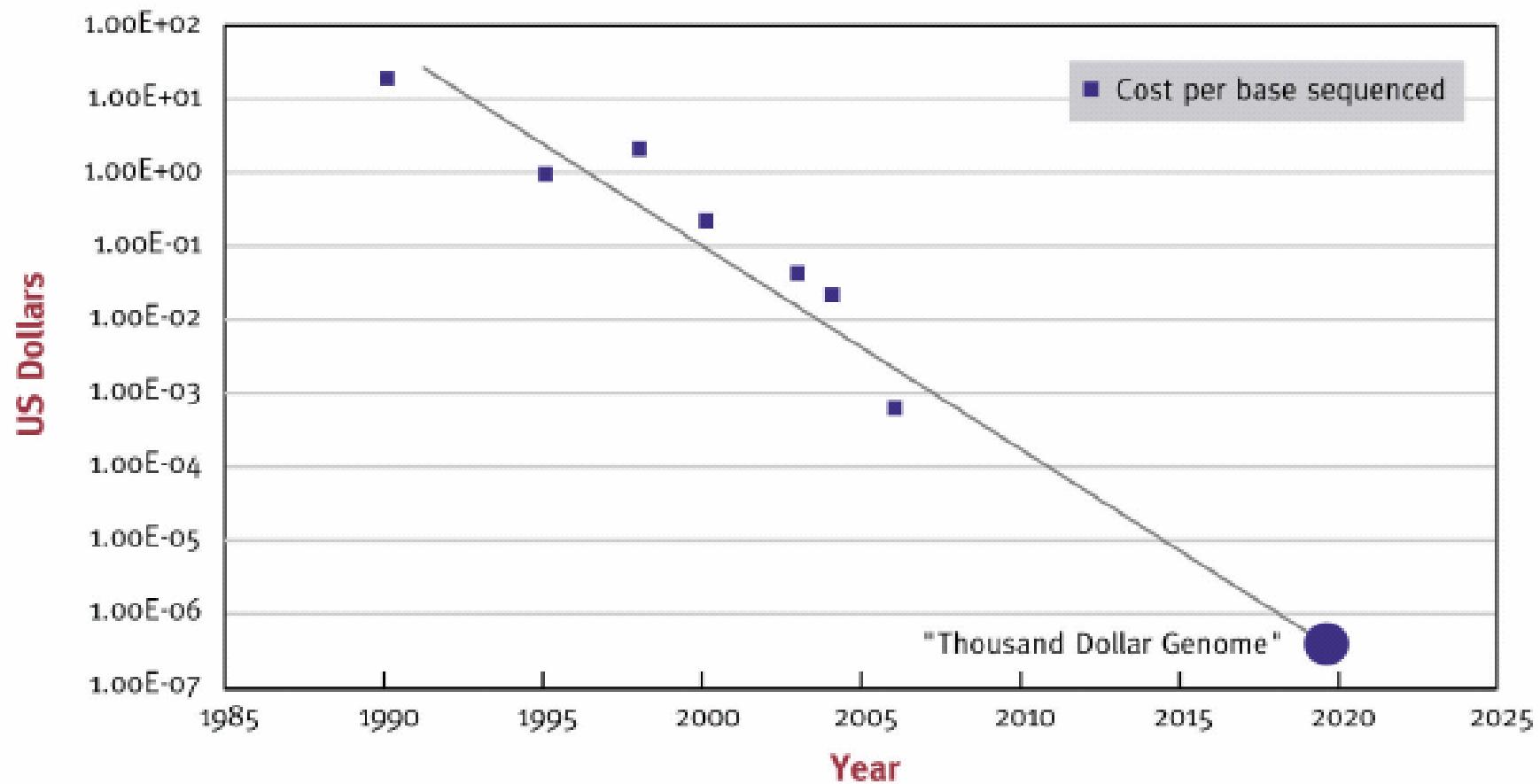
1995  
144,000 bp/day (semi-auto)



1998  
500,000 bp/day (automatic)

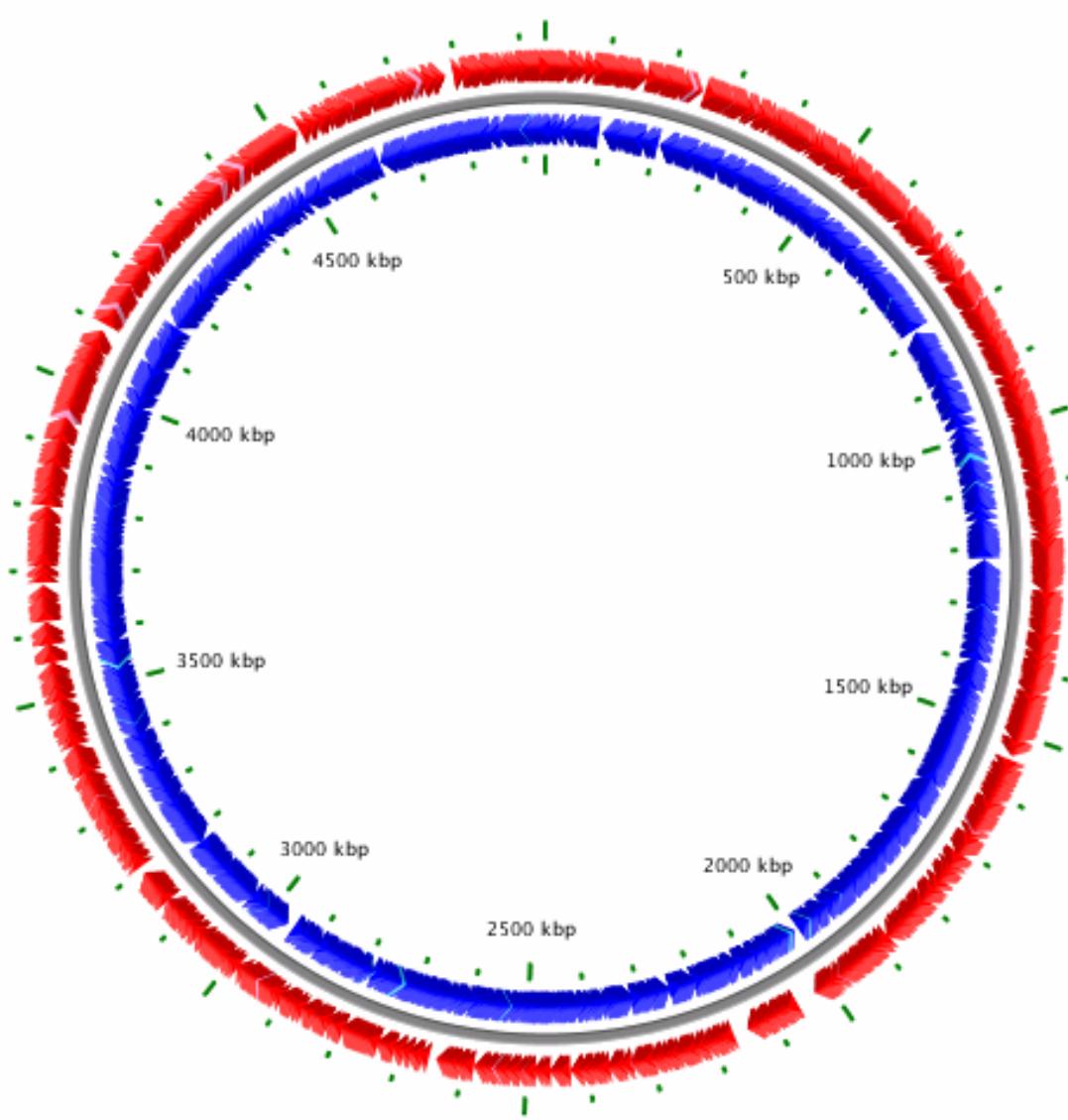


2007 – Sequencing by Synthesis  
1GB bp/day (automatic)



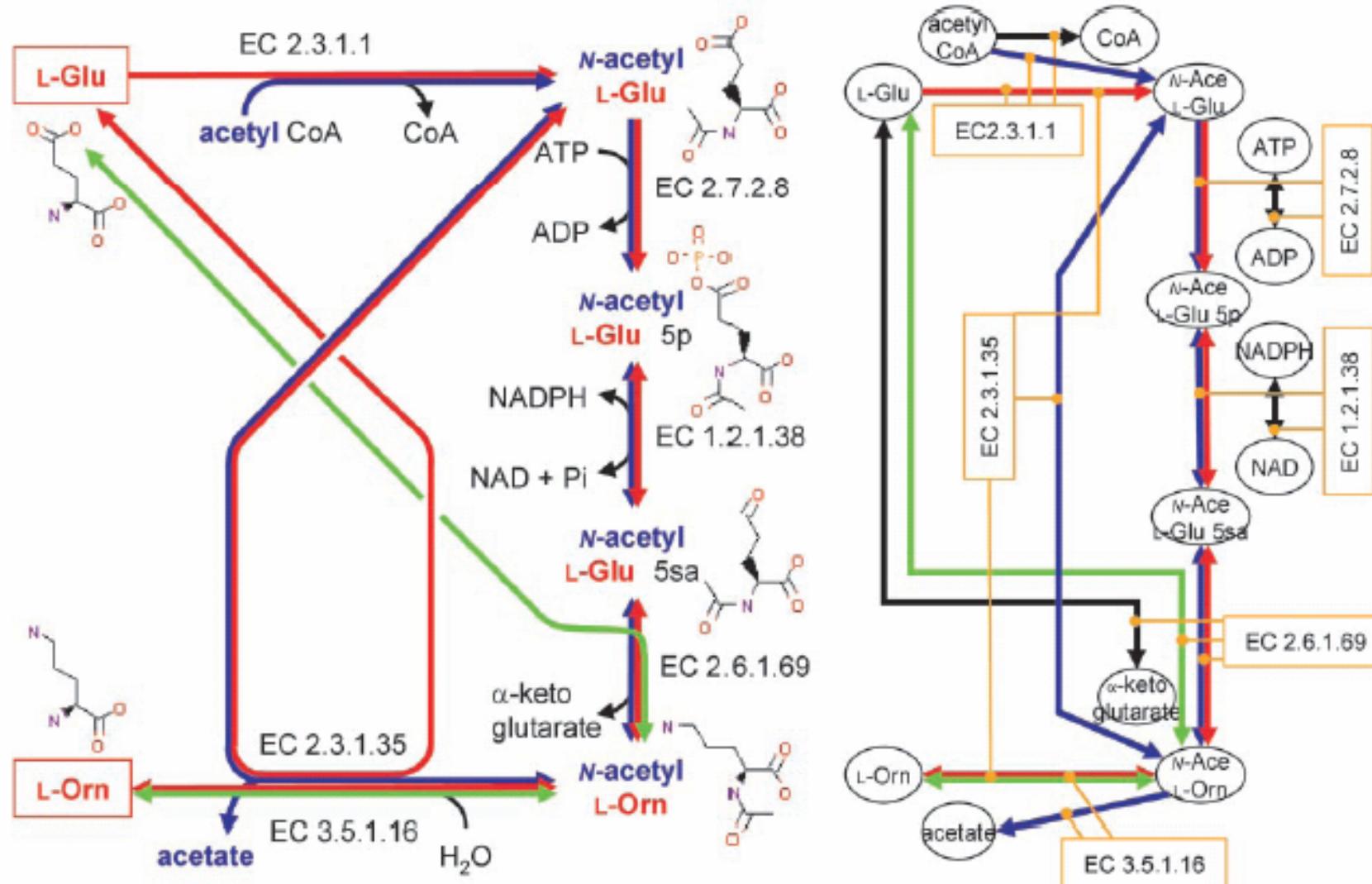
Source: R. Carlson, Bio-era

*Escherichia coli* 536, complete genome

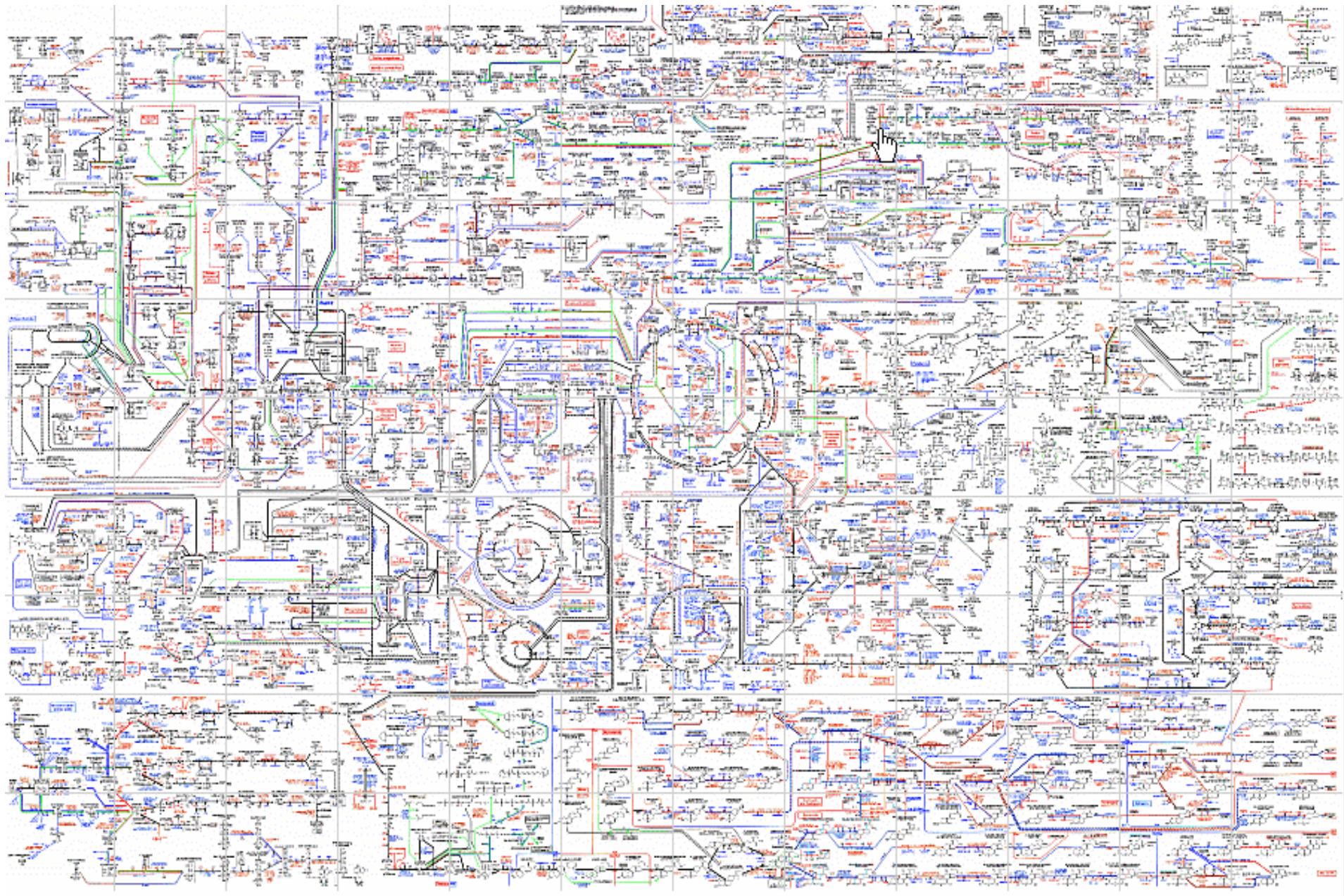


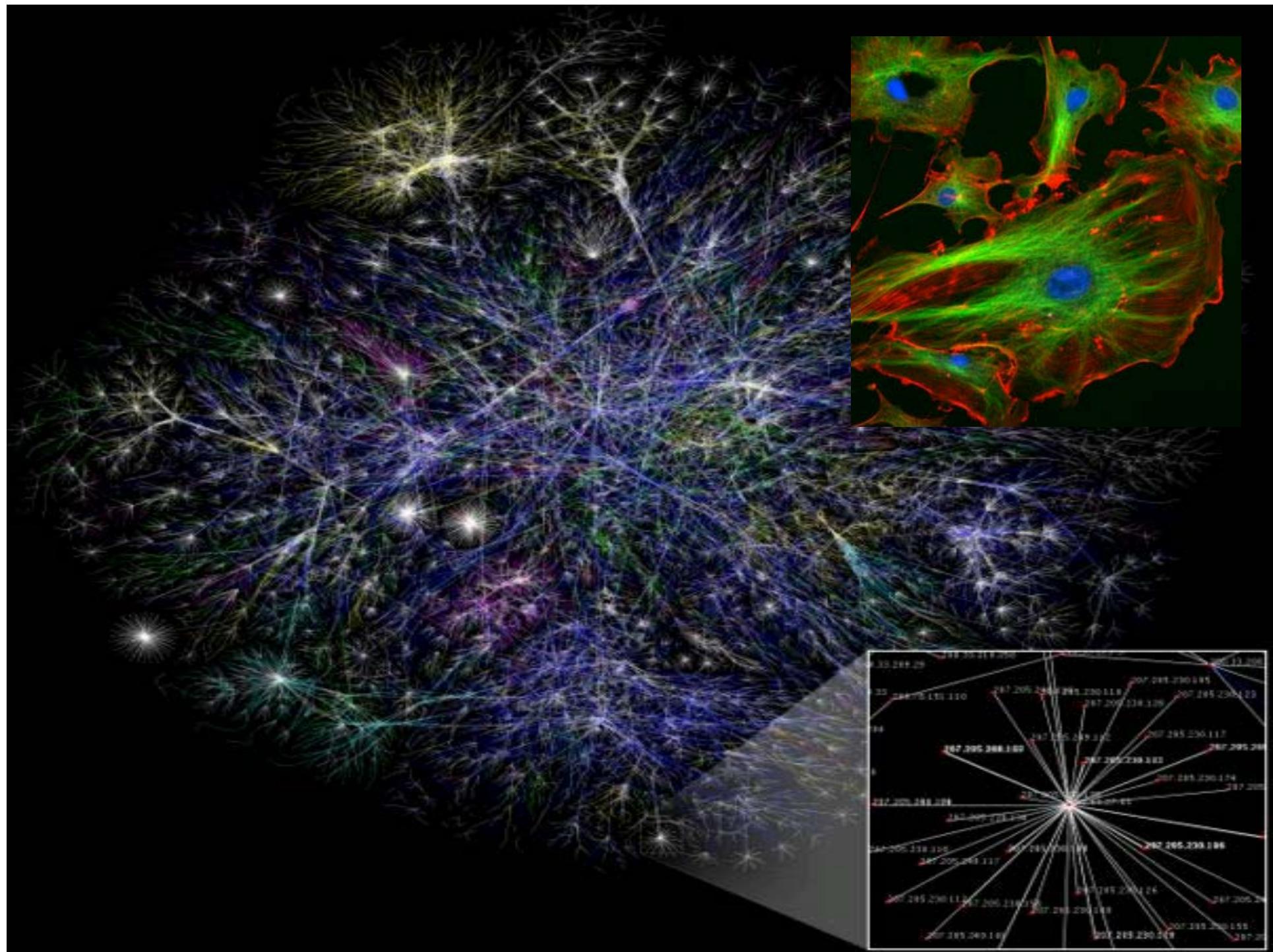
Accession: NC\_008253

Topology: circular; Length: 4,938,920 bp; Genes: 4,732



Ornithine biosynthesis





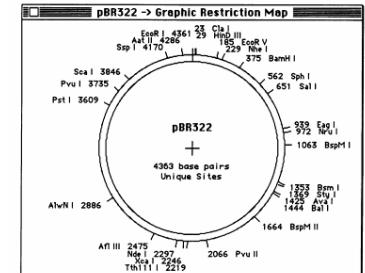
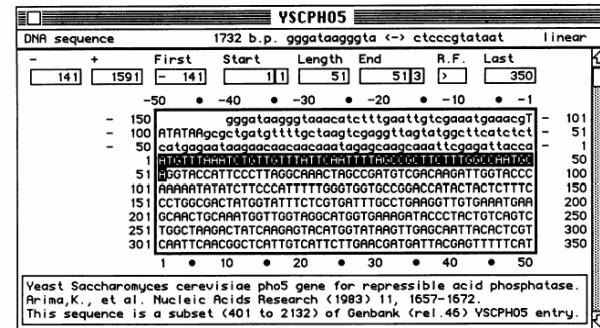


'DNA Strider': a 'C' program for the fast analysis of DNA and protein sequences on the Apple Macintosh family of computers

Christian Marck

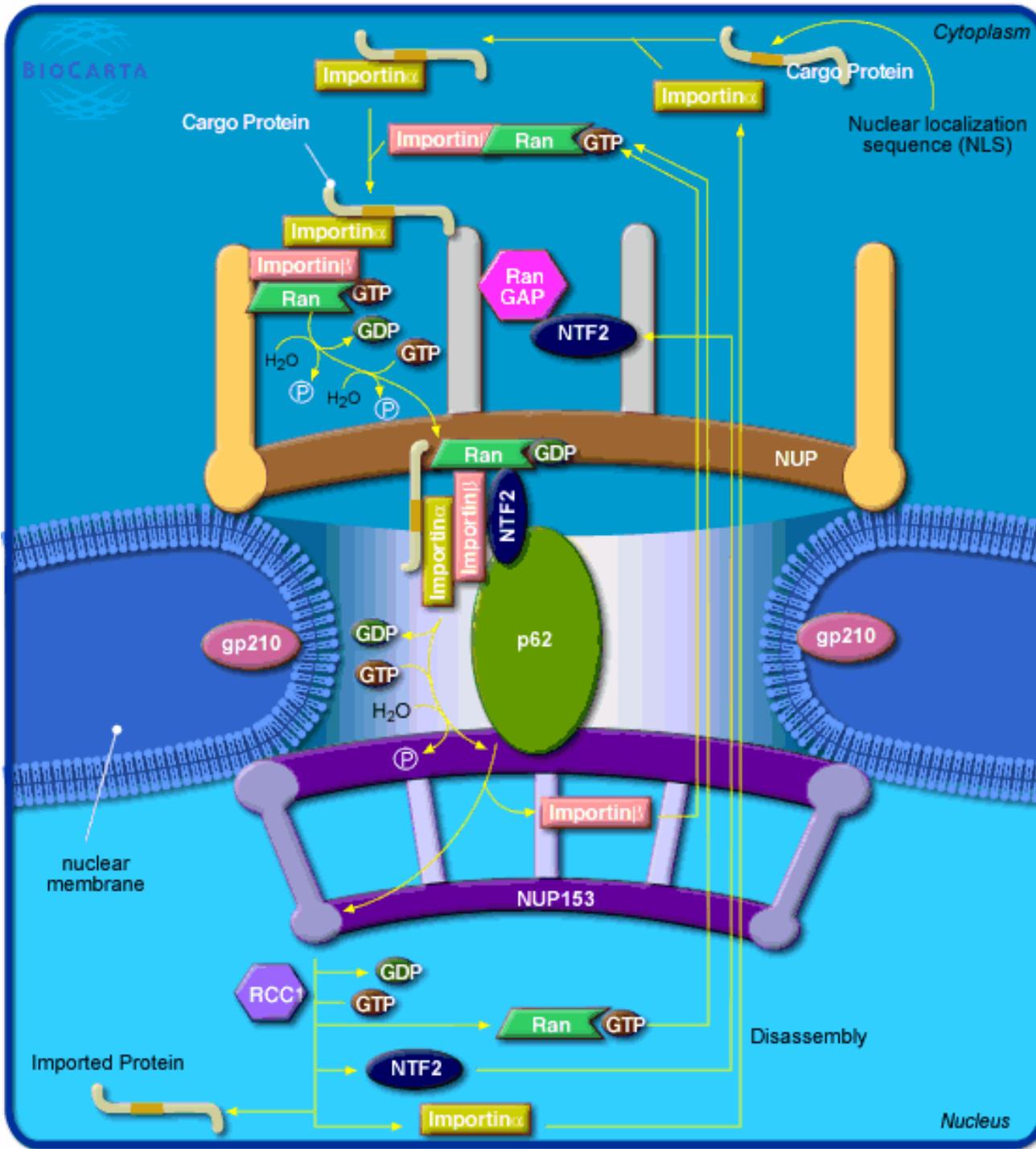
Service de Biochimie, Bâtiment 142, Département de Biologie, Centre d'Etudes Nucléaires de Saclay  
91191 Gif-sur-Yvette Cedex, France

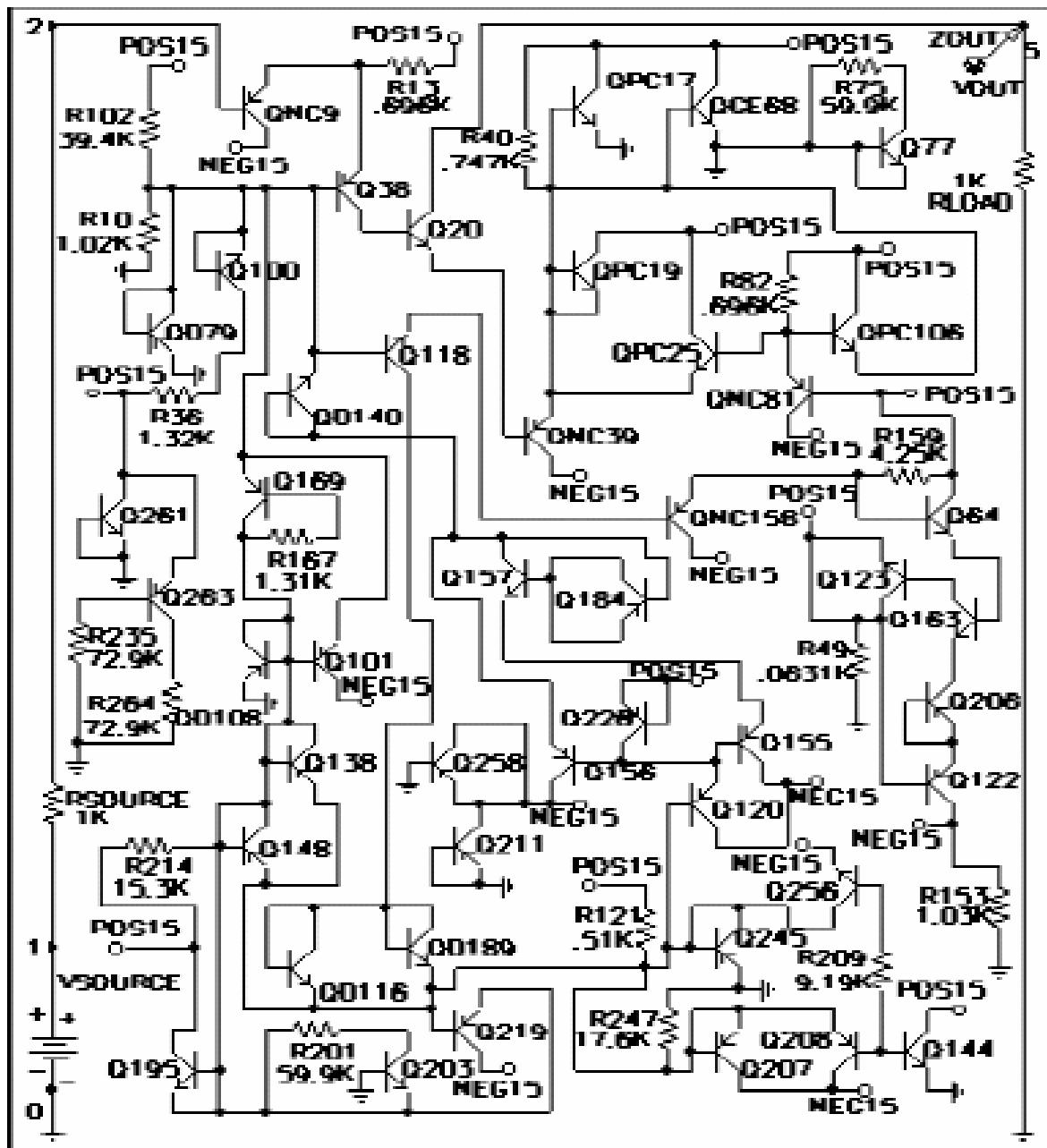
Received August 17, 1987; Revised and Accepted November 15, 1987



## IBM's BlueGene/L: world's fastest supercomputer, 3 years running

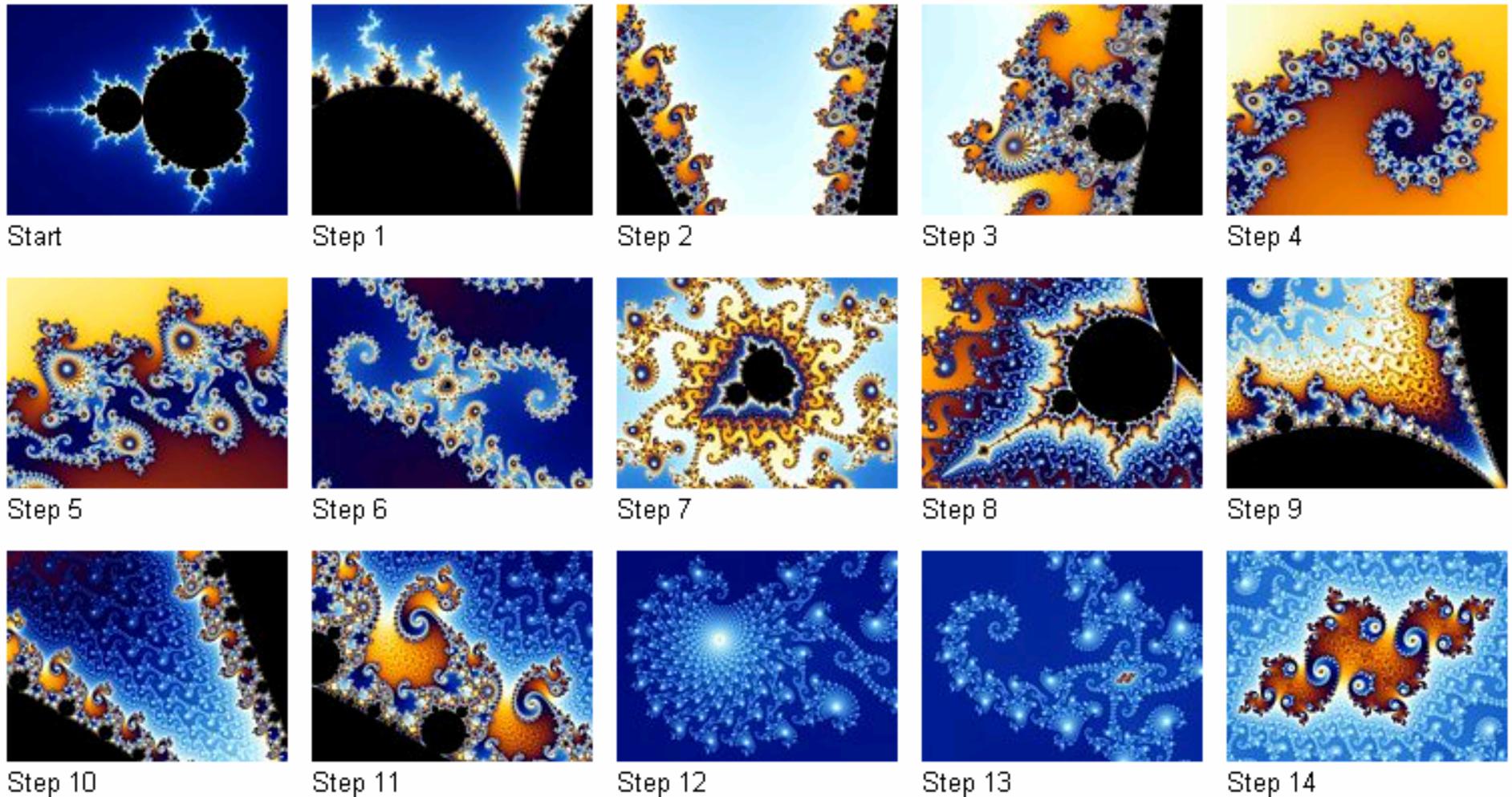
478.2 TFLOPS with 131072 nodes



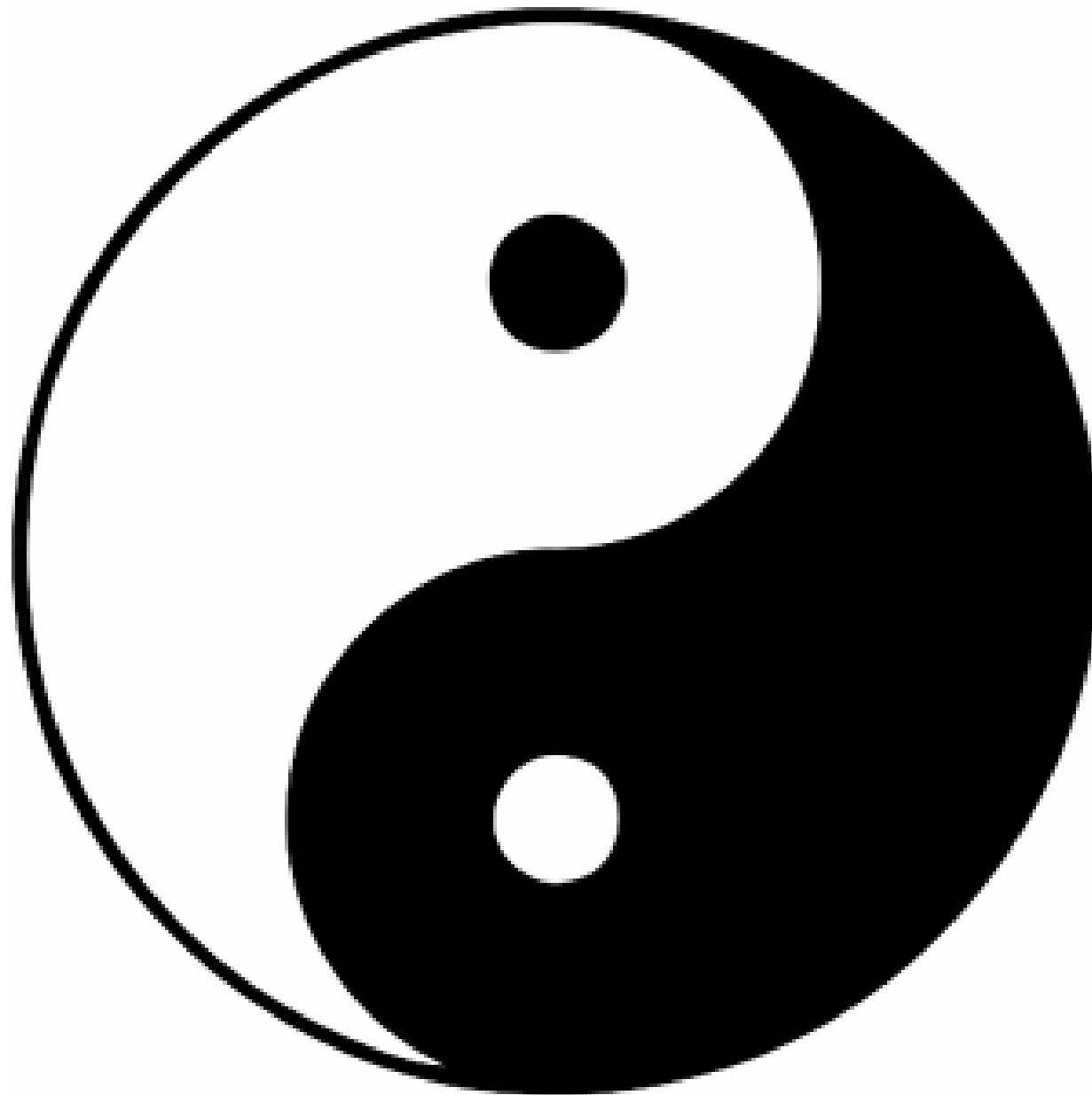


J.R. Koza et al.  
Automatic creation of computer  
programs for designing  
electrical circuits using genetic  
programming.

$$M = \left\{ c \in \mathbb{C} : \sup_{n \in \mathbb{N}} |f_c^n(0)| < \infty \right\}.$$



Reduction | Complexity



If we can't build it, we don't understand it.

Genetic “Engineering”

Writing code

**Biochemical Method for Inserting New Genetic Information into DNA of Simian Virus 40: Circular SV40 DNA Molecules Containing Lambda Phage Genes and the Galactose Operon of *Escherichia coli***

(molecular hybrids/DNA joining/viral transformation/genetic transfer)

DAVID A. JACKSON\*, ROBERT H. SYMONS†, AND PAUL BERG

Department of Biochemistry, Stanford University Medical Center, Stanford, California 94305

*Contributed by Paul Berg, July 31, 1972*

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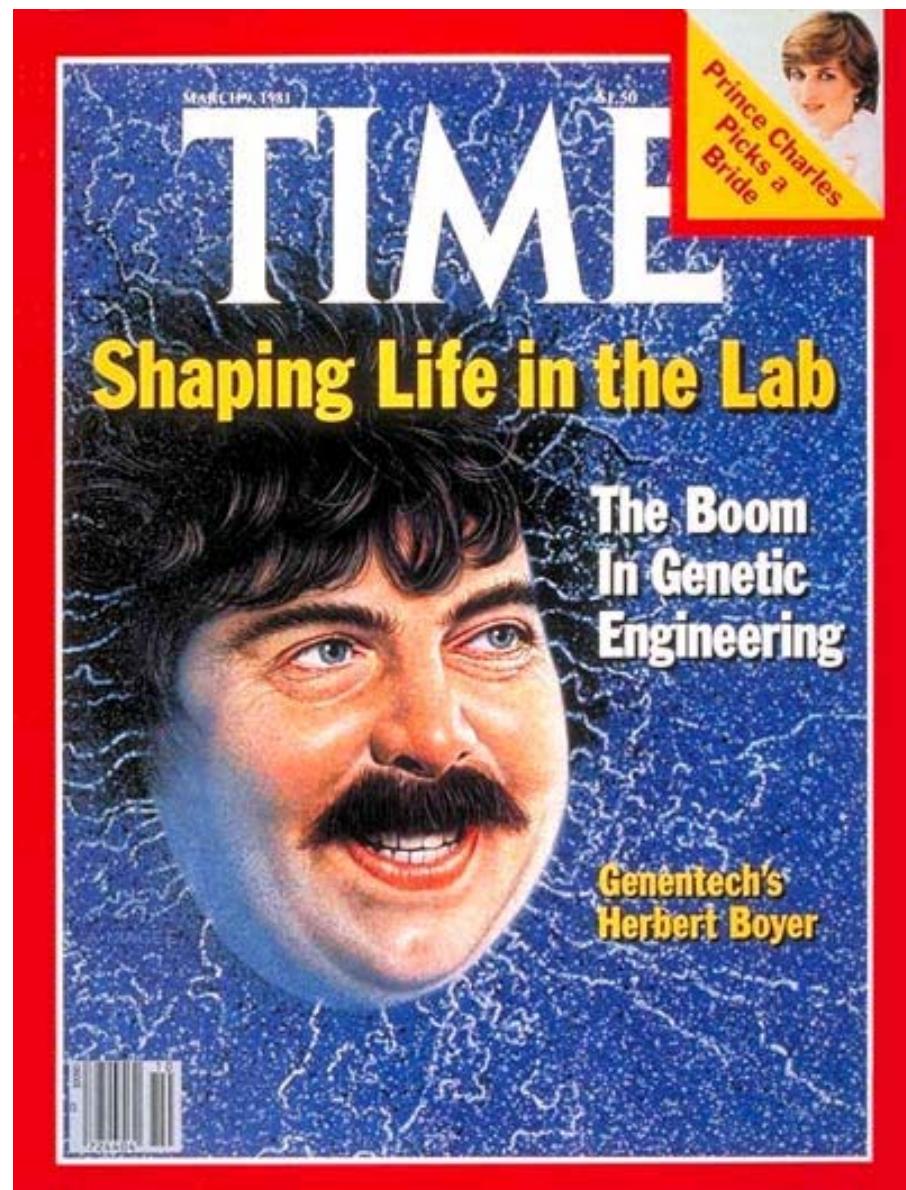
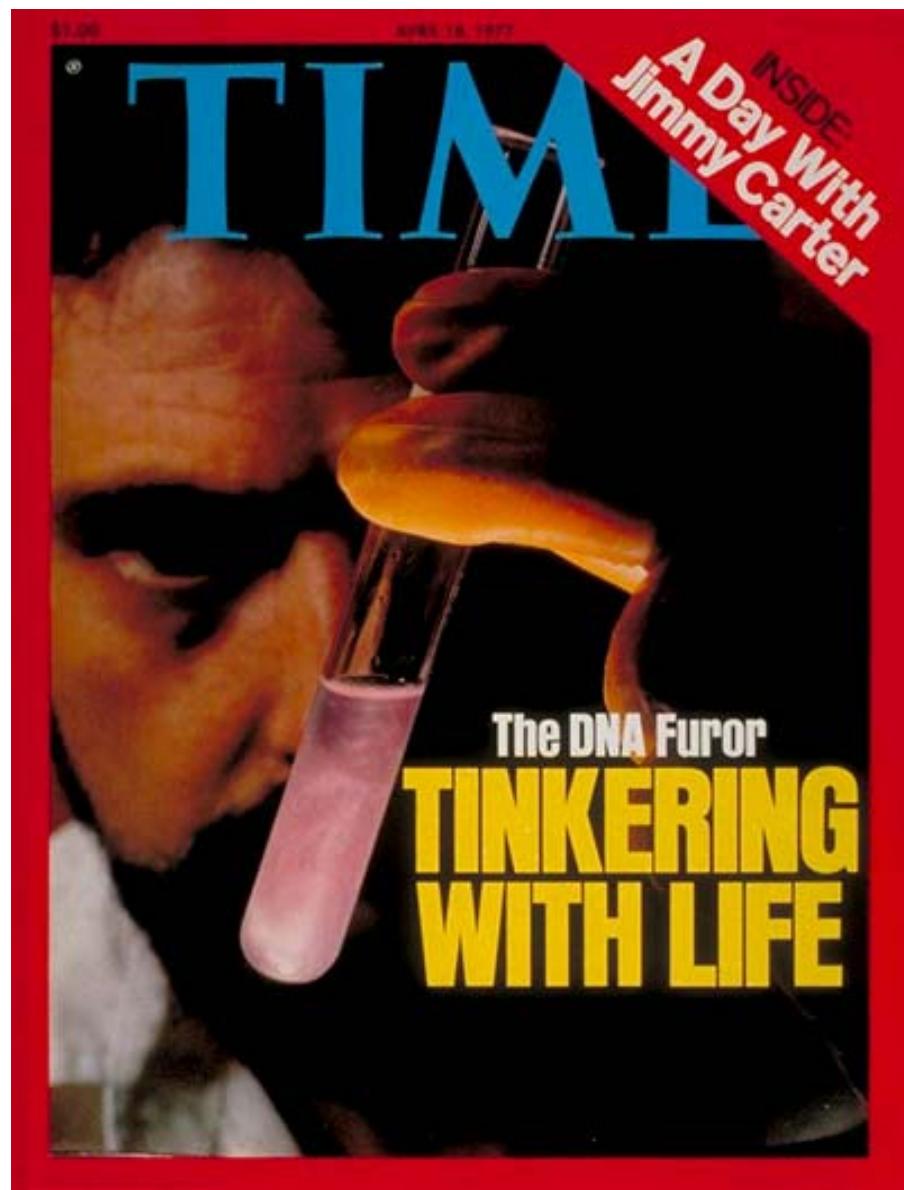
**Construction of Biologically Functional Bacterial Plasmids *In Vitro***

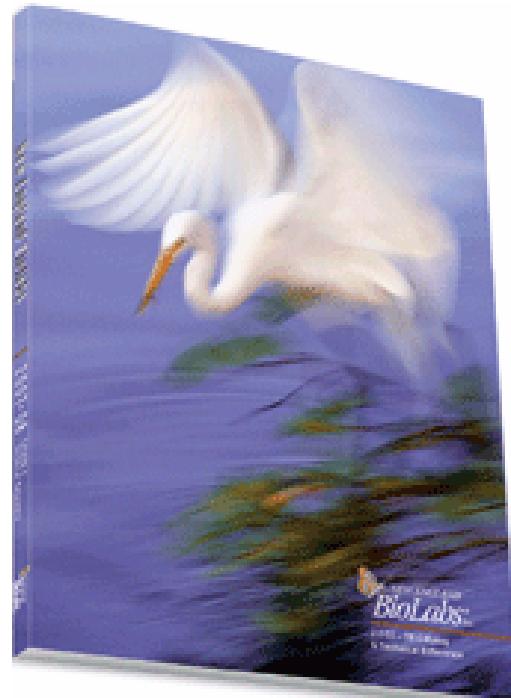
(R factor/restriction enzyme/ transformation/ endonuclease/ antibiotic resistance)

STANLEY N. COHEN\*, ANNIE C. Y. CHANG\*, HERBERT W. BOYER†, AND ROBERT B. HELLING†

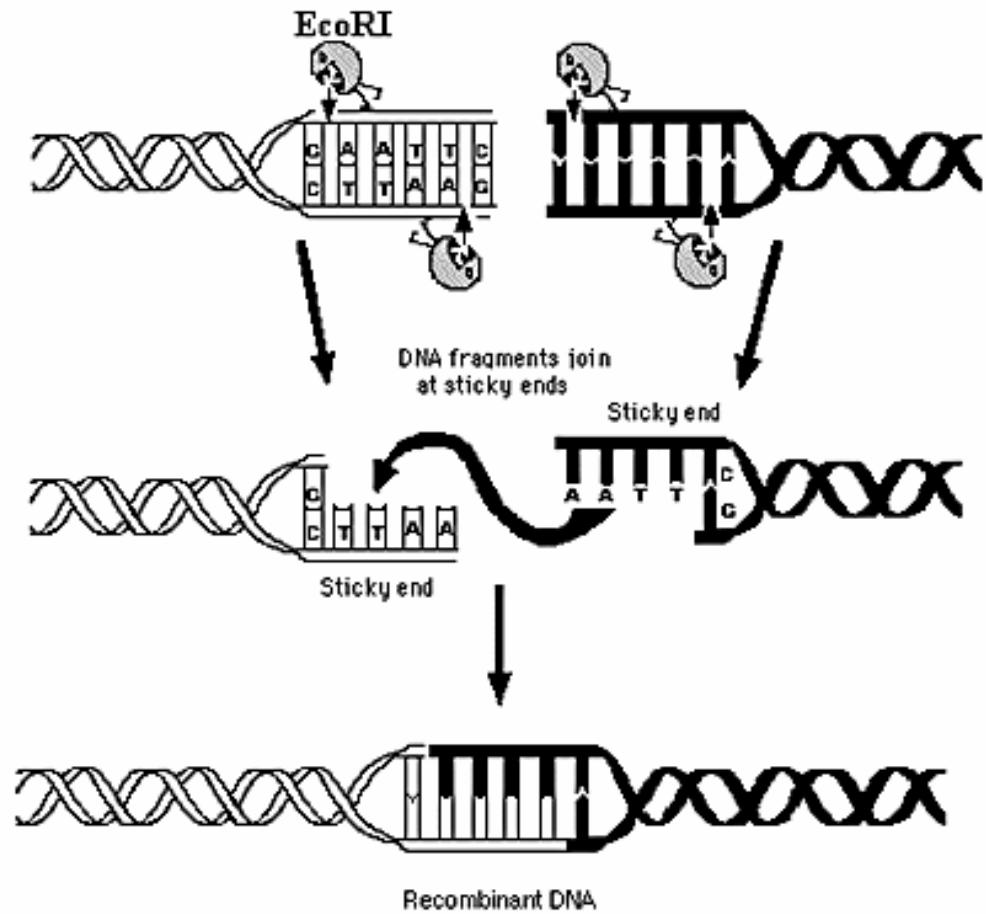
\* Department of Medicine, Stanford University School of Medicine, Stanford, California 94305; and † Department of Microbiology, University of California at San Francisco, San Francisco, Calif. 94122

*Communicated by Norman Davidson, July 18, 1973*





Over 3500 RE's available

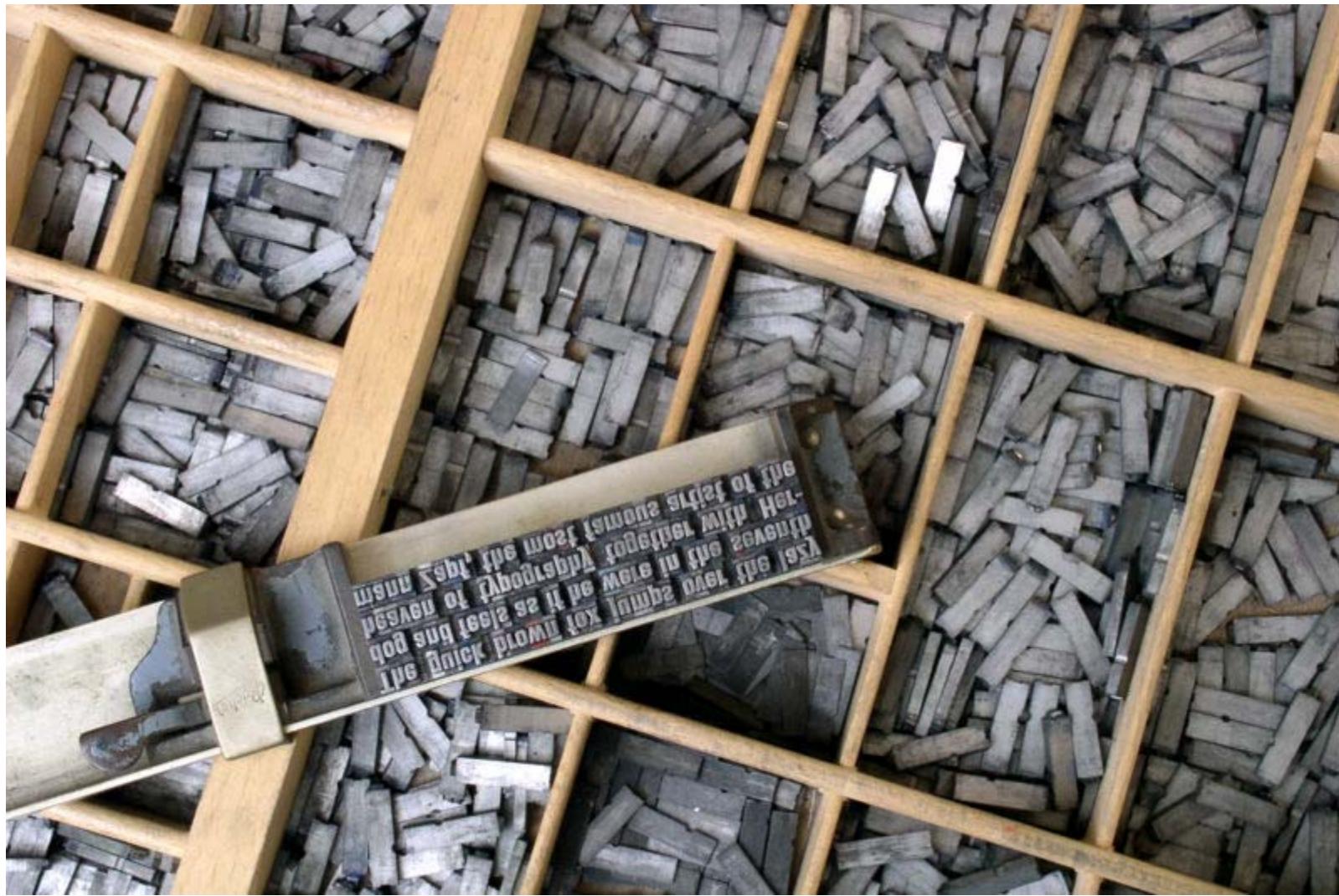


**Restriction Enzyme  
Action of EcoRI**

if you can WRITE DNA,

You'RE no LONGER liMITED

to "What IS IT" but TO what you could MAKE •



[http://en.wikipedia.org/wiki/Image:Metal\\_movable\\_type.jpg](http://en.wikipedia.org/wiki/Image:Metal_movable_type.jpg)

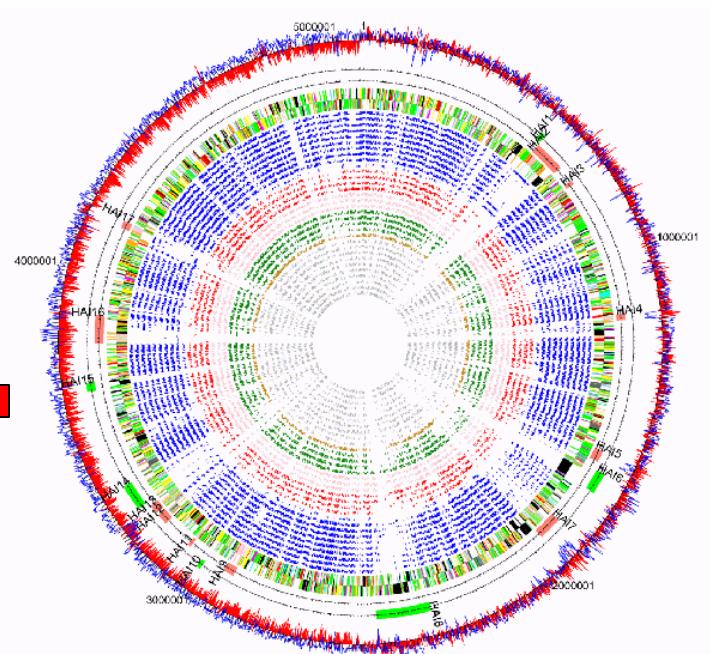


## Physical DNA

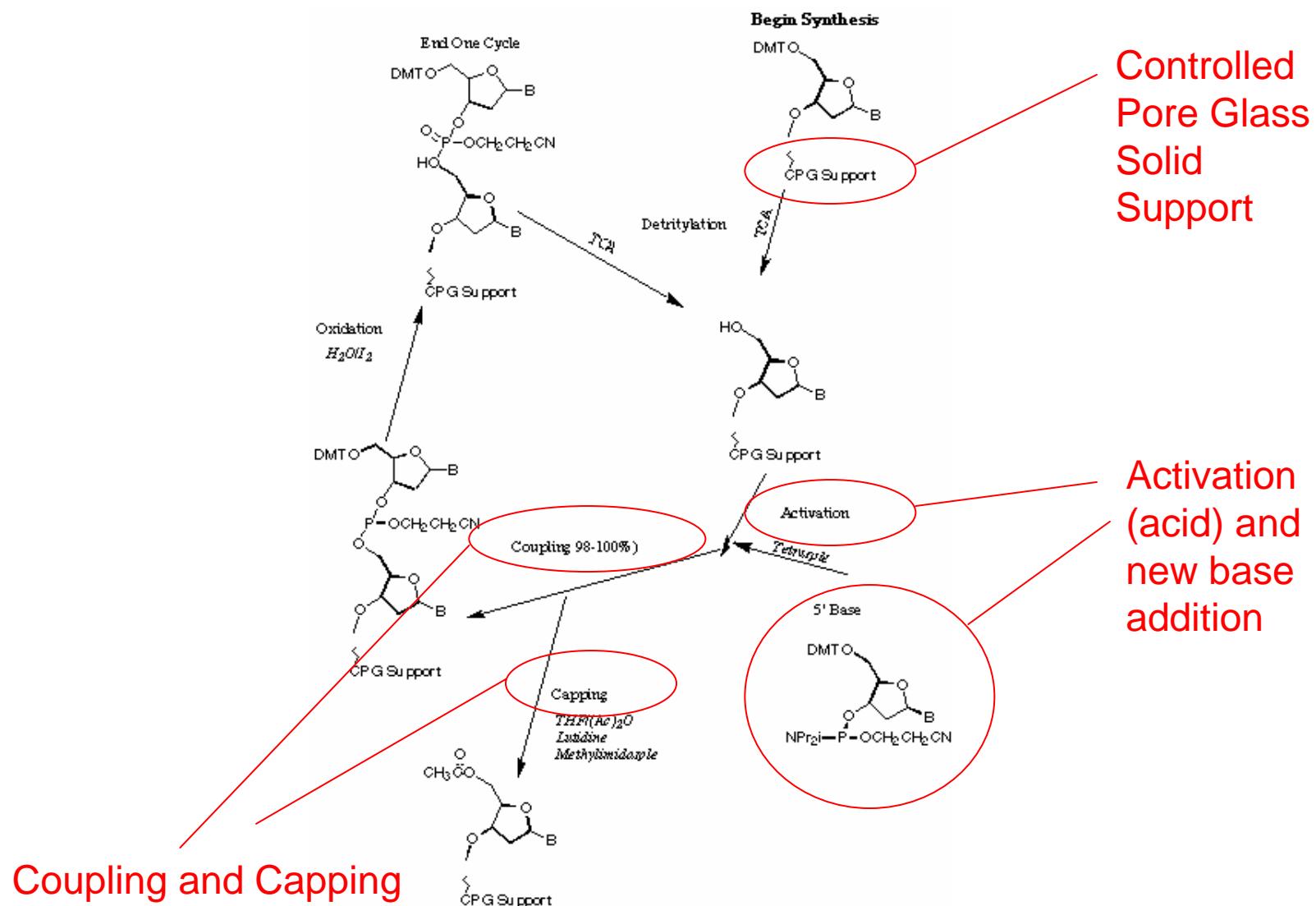
## sequencing

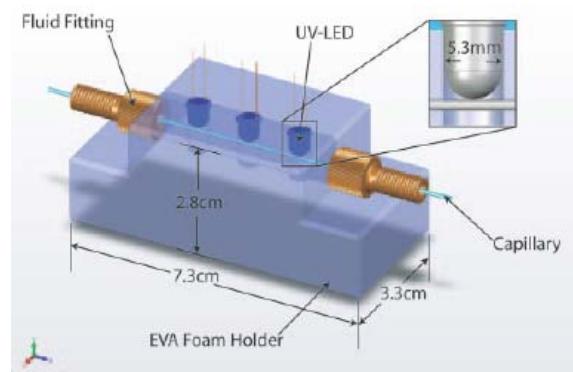
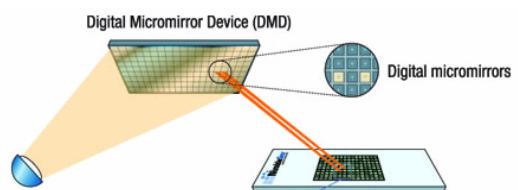
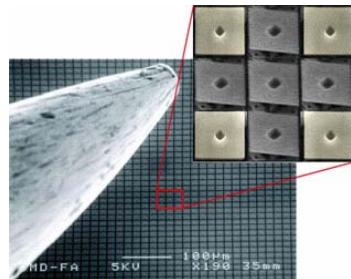
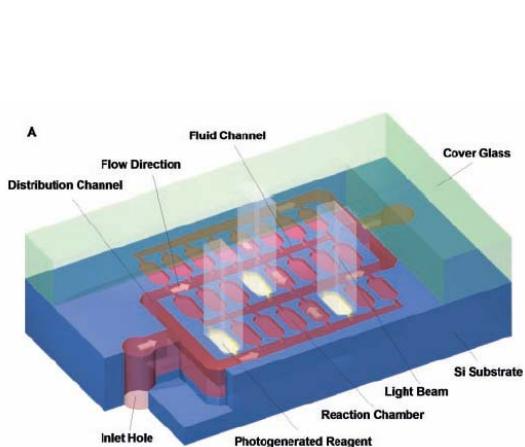


## synthesis



# Digital DNA





# MILLIGEN / MILLIPORE DNA SYNTHESIZER CYCLONE PLUS 8400

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1 of 9



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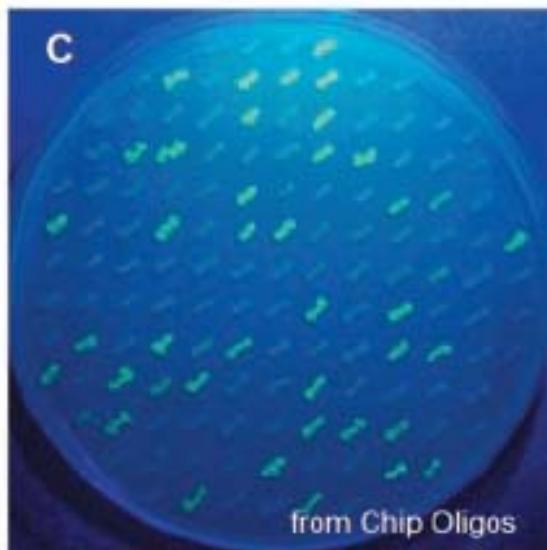
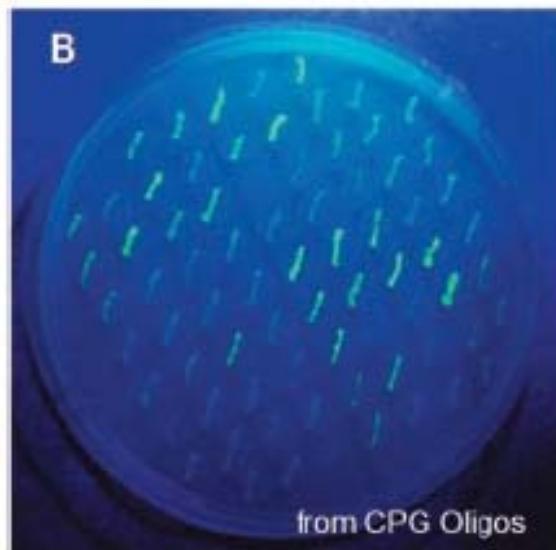
Standard Flat Rate Shipping Service  
Service to [United States](#)

Ships to: Worldwide

Item location: N. E. OKLA., United States

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EGFP gene 714 bp

**D**

Oligonucleotide set 1

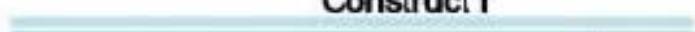


Oligonucleotide set 2



↓ ligation

Construct I



Construct II



↓ PCR

DNA Construct

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buy now at our best rate ever!  
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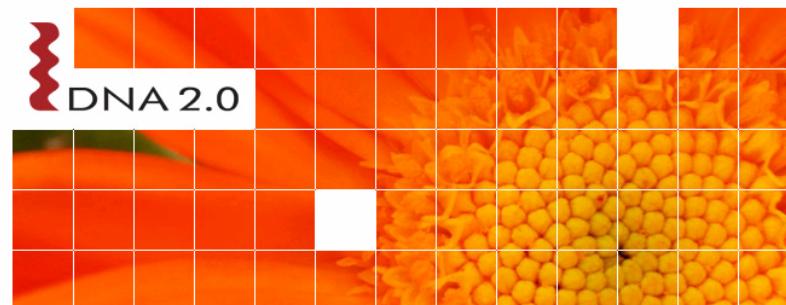
### DNA 2.0

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Your Building Blocks of Life  
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### Blue Heron Bio GeneMaker

Call now for promotional pricing!  
**Synthetic** Gene Specialists.  
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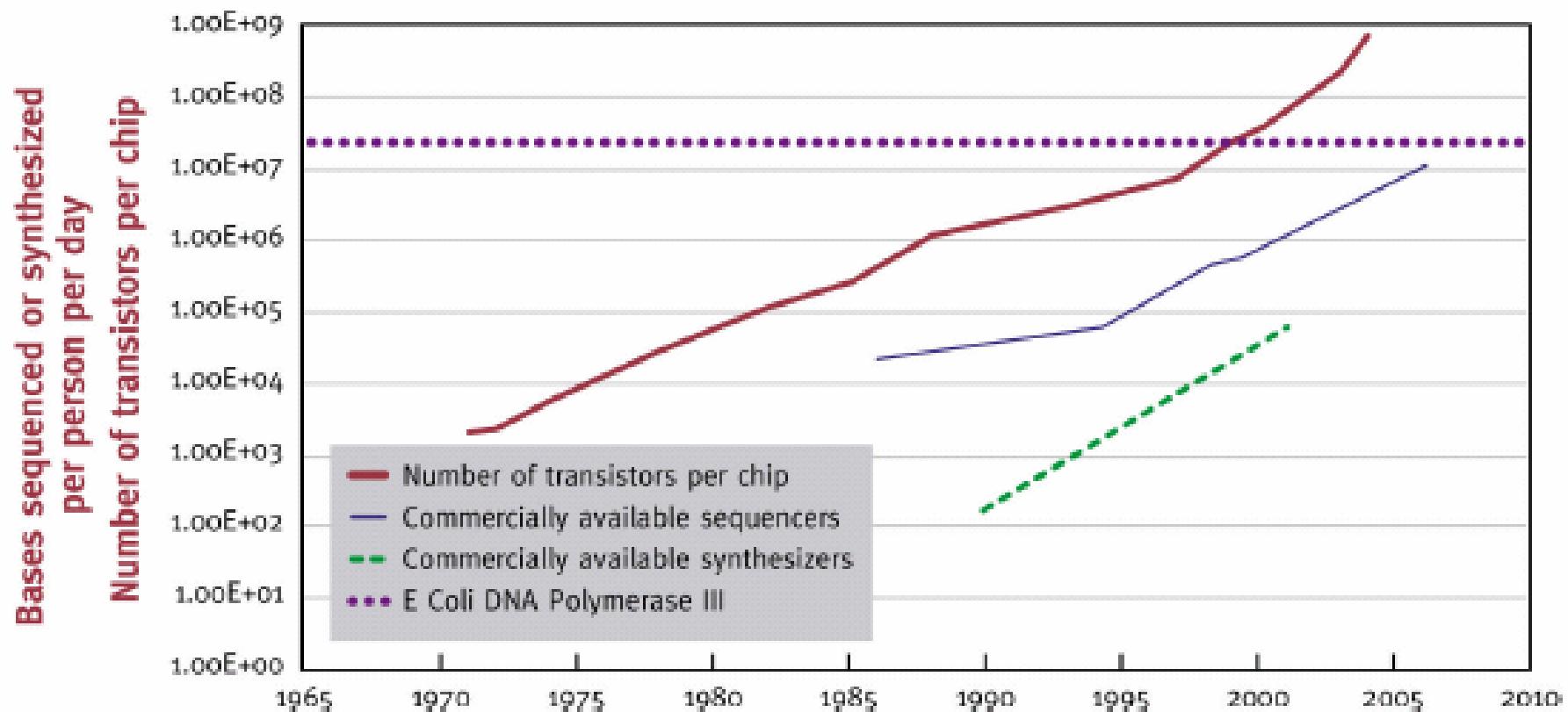
11/23/2007



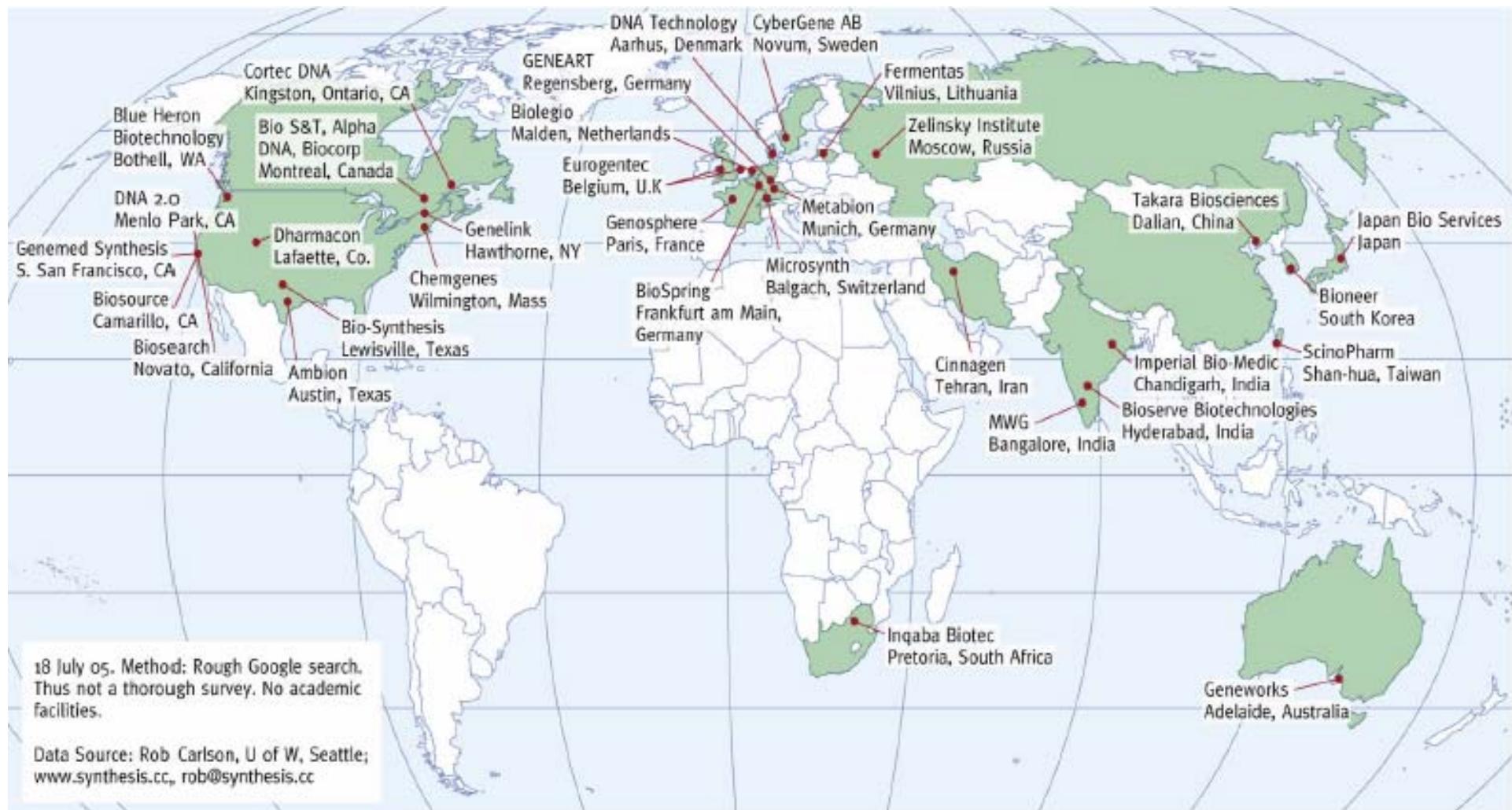
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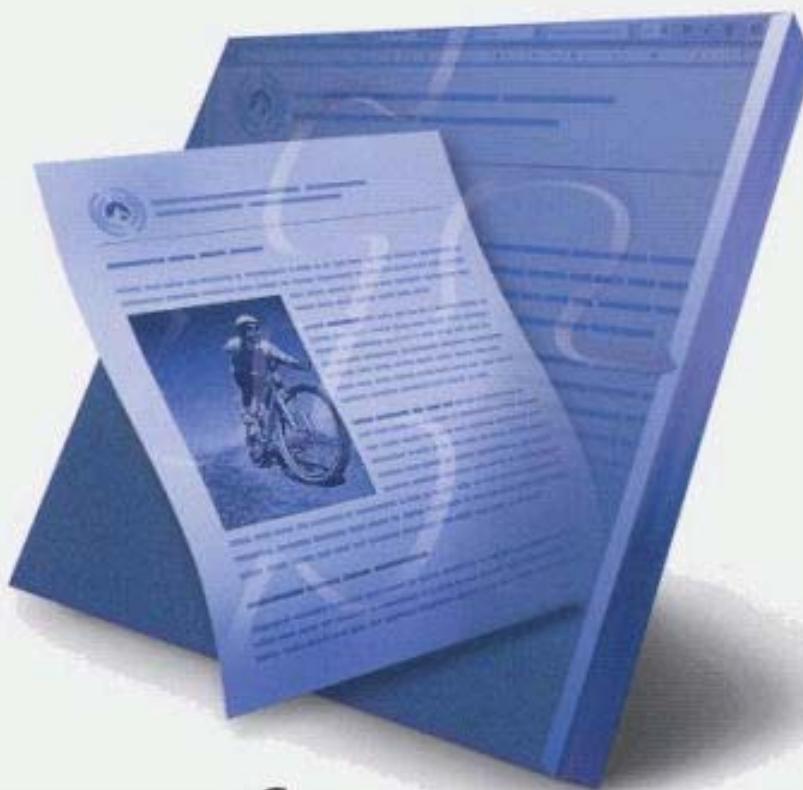
Source: R. Carlson, Bio-era.



Sources: Source: R. Carlson, G. Epstein, A. Yu (2005)

Designed for  
  
Microsoft®  
Windows NT®  
Windows®98

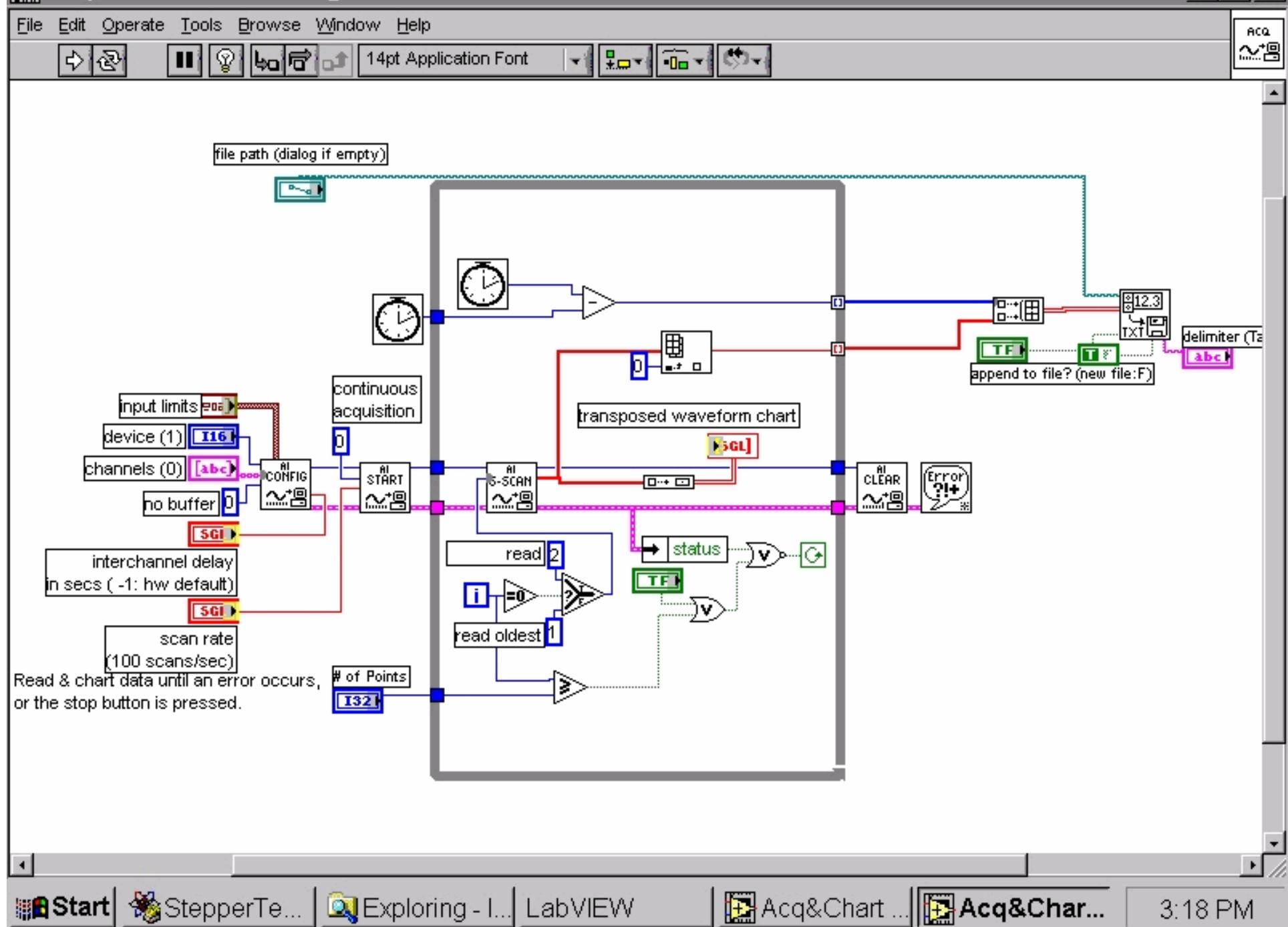
**Microsoft®**

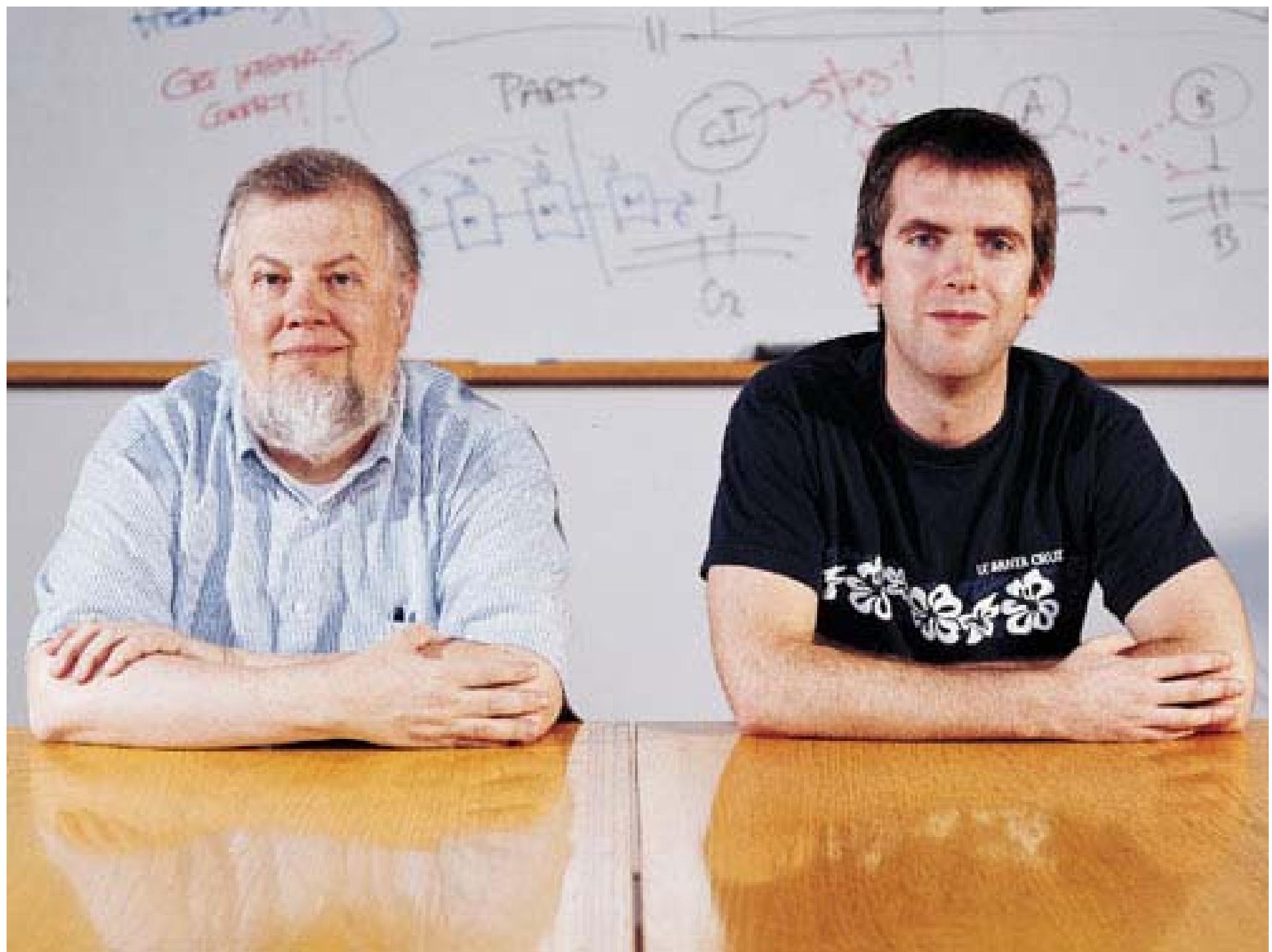


Microsoft®  
**Word 2000**   
Microsoft Office Application

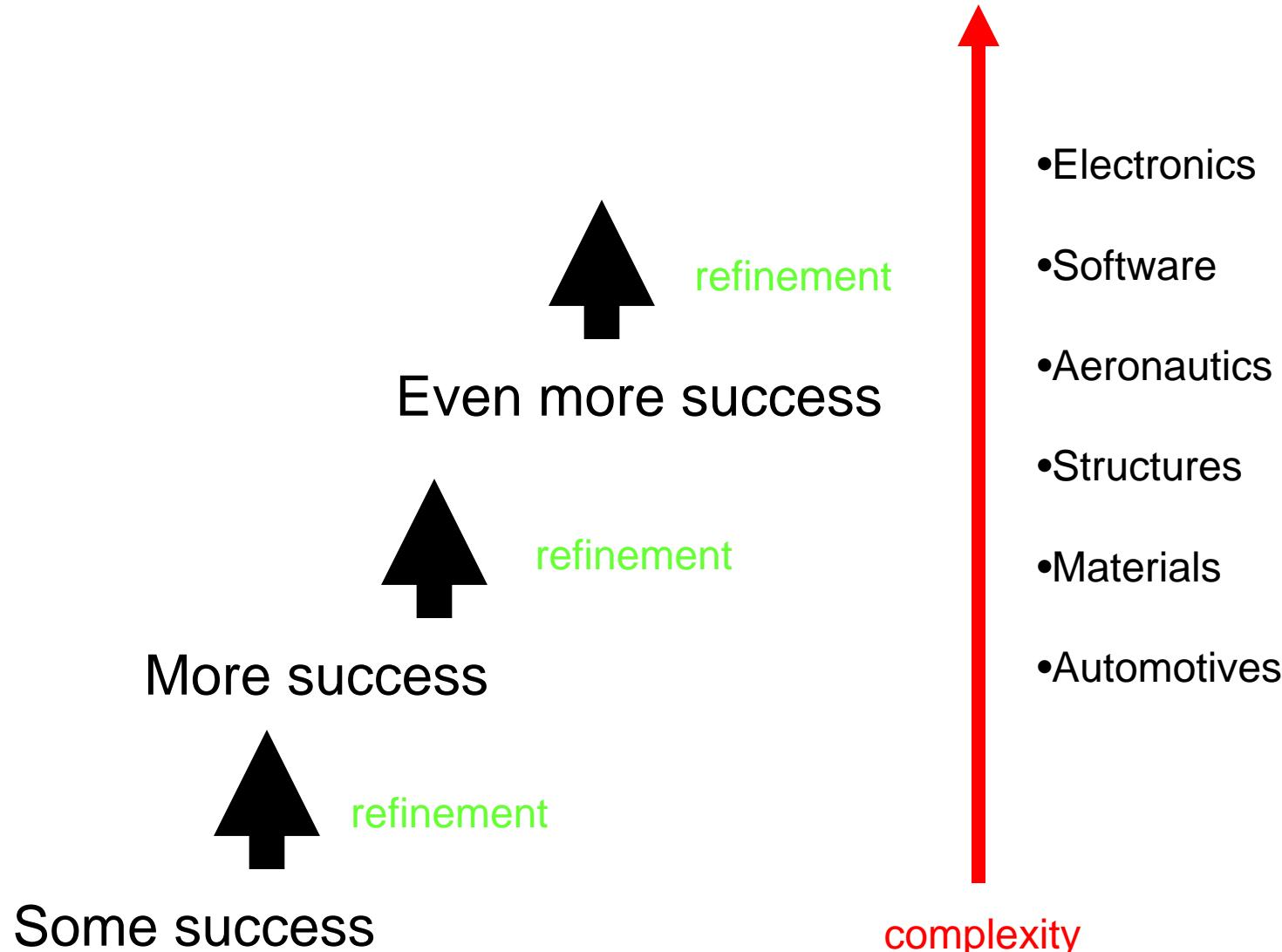
The Microsoft Office Word Processor

Acq&Chart to xls.vi Diagram \*





# Engineering process...



# F1760

Sender Device

# B0015

terminator

Name: B0015

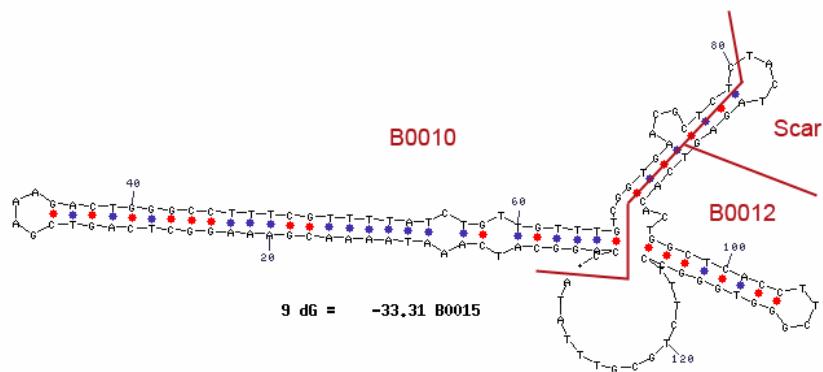
Type: Double terminator

Length 129 bp

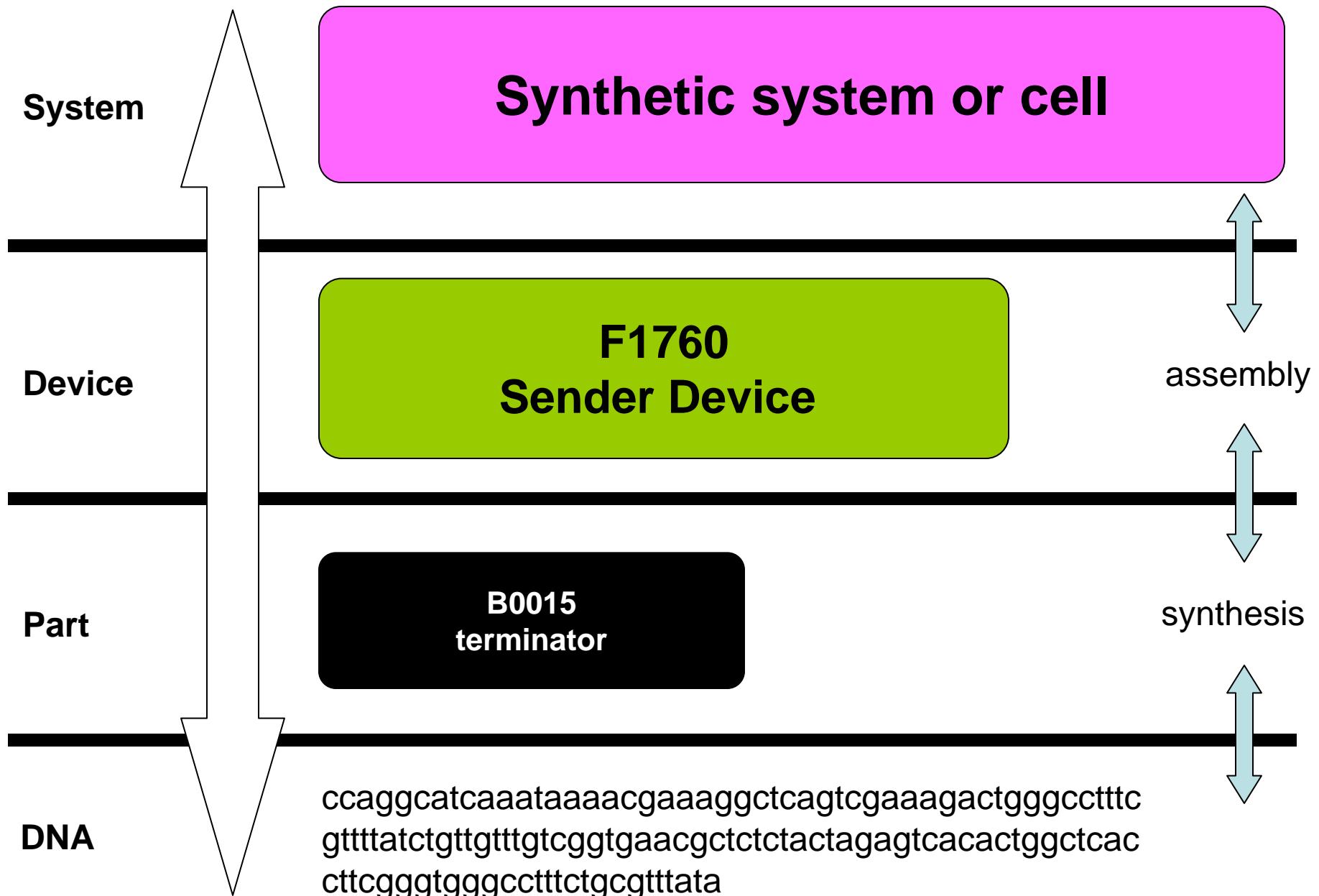
Designed by: Reshma Shetty

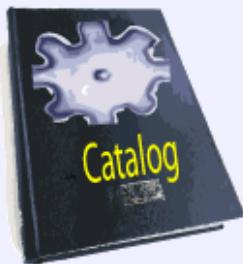
Forward efficiency: 0.984

Reverse efficiency: .295



## STANDARDIZED DATA




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# Transcriptional Regulators

**Available repressible regulators (normally ON) [-?](#)**
[Show 0 more parts](#)
[Edit](#)

<a href="#">-?</a>	<b>Name</b>	<b>Description</b>	<b>Direction</b>	<b>Control <a href="#">-?</a></b>	<b>Output Low High</b>	<b>Length</b>
A	<a href="#">W BBa_I14032</a>	promoter P(Lac) IQ	Forward			37
A	<a href="#">W BBa_R0040</a>	promoter (tetR, negative)	Forward	aTc, tetracycline		54
A	<a href="#">W BBa_R0051</a>	promoter (lambda cl regulated)	Forward	lambda cl		49

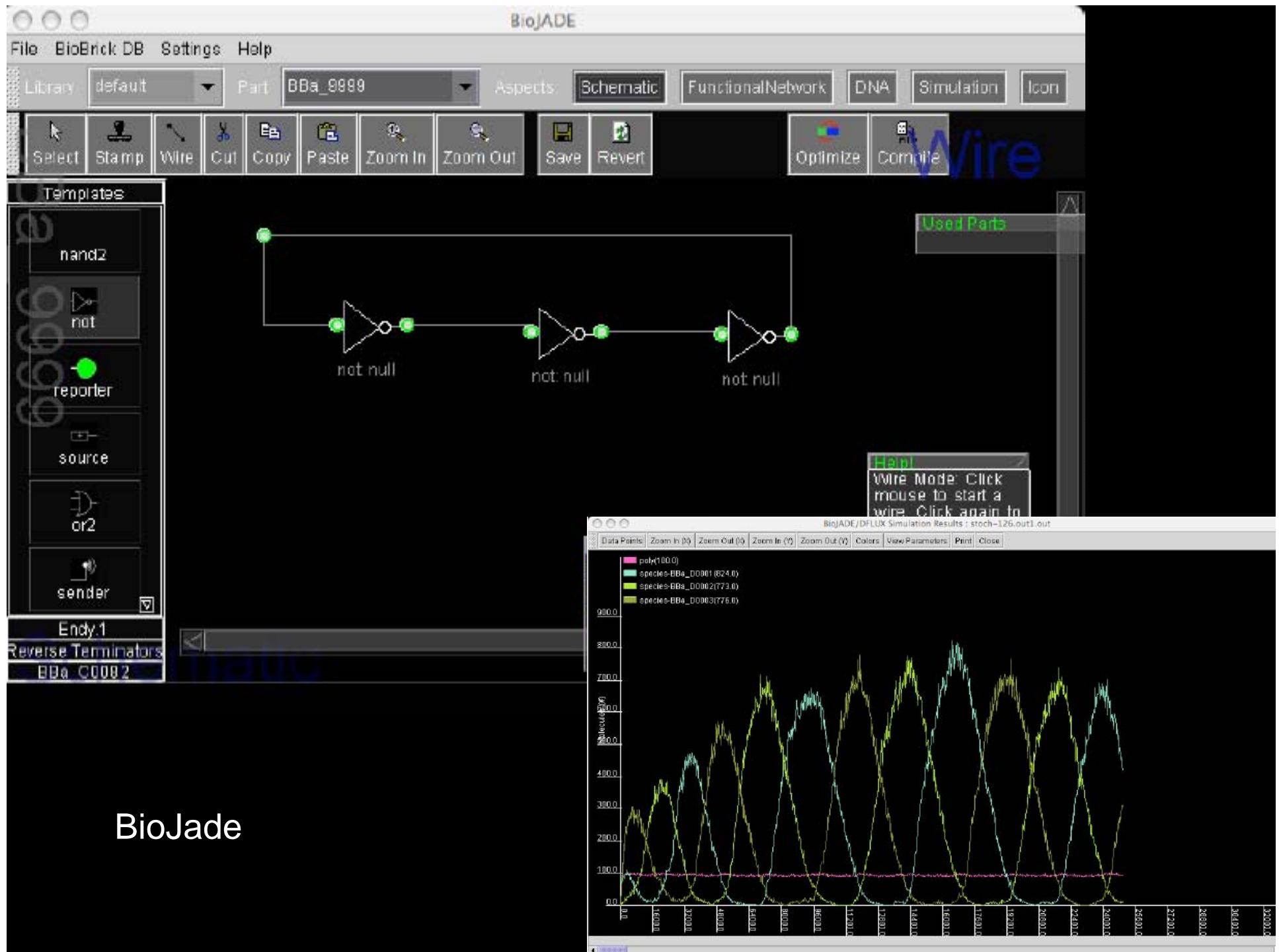
**Available inducible regulators (normally OFF) [-?](#)**
[Show 0 more parts](#)
[Edit](#)

<a href="#">-?</a>	<b>Name</b>	<b>Description</b>	<b>Direction</b>	<b>Control <a href="#">-?</a></b>	<b>Output Low High</b>	<b>Length</b>
A	<a href="#">BBa_I12007</a>	Modified lambda Prm promoter (OR-3 obliterated)	Forward	cl		82
A	<a href="#">BBa_R0062</a>	Promoter (luxR & HSL regulated -- lux pR)	Forward	luxR, HSL		55
A	<a href="#">BBa_R0079</a>	Promoter (LasR & PAI regulated)	Forward	PAI		157
A	<a href="#">BBa_R0080</a>	Promoter (AraC regulated)	Forward	araC		149

**Available other regulators**
[Show 172 more parts](#)
[Edit](#)

<a href="#">-?</a>	<b>Name</b>	<b>Description</b>	<b>Direction</b>	<b>Control <a href="#">-?</a></b>	<b>Output Low High</b>	<b>Length</b>
A	<a href="#">W BBa_I0500</a>	Inducible pBAD/araC	Forward	araC, arabinose		1210
A	<a href="#">W BBa_I13453</a>	Pbad promoter				130
A	<a href="#">W BBa_J13002</a>	TetR repressed POPS/RIPS generator	Forward	ATc		74
A	<a href="#">W BBa_J13023</a>	3OC6HSL+LuxR dependent POPS/RIPS generator				117
A	<a href="#">W BBa_J23100</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23101</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23102</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23103</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23104</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23105</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23106</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23107</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23108</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23109</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23110</a>	constitutive promoter family member				35
A	<a href="#">W BBa_J23111</a>	constitutive promoter family member				35





BioJADE

## How to use this site:

GenoCAD™ is an experimental tool allowing you to build and verify complex genetic constructs derived from a library of standard genetic parts.

© 2007 Virginia Bioinformatics Institute  
contact information

## Design

- 1 Think of a construct
- 2 Build its structure
- 3 Select its parts
- 4 Download your sequence

[Start a Design ▶](#)

## Validate

- 1 Upload your sequence
- 2 Click validate
- 3 View structure
- 4 Download lexical analysis!

[Validate ▶](#)



# Training DNA programmers

*Synthetic biology requires an unprecedented level of interdisciplinary cooperation, fewer laboratory skills, and less overhead than conventional genetic engineering.*

## Cooperative yet competitive

Open education format...

Shares:

- DNA parts
- DNA code
- Protocols
- Experience
- Publications
- One big rule: share back!





# iGEM 2007 Wiki

International Genetically Engineered Machine Competition

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## Results of the Jamboree



sat & sun, nov 3-4

iGEM 2007 is now officially concluded! Congratulations to all!

- [Results](#)
- [See the medal winners](#)
- [Media](#) (including links to videos and [flickr™ photo gallery](#))
- [Learn about iGEM 2008](#)

## iGEM?

Hundreds of undergraduates all over the world spend their summer making Synthetic Biology a reality by participating in the annual International Genetically Engineered Machine competition.

iGEM through the years

- [2008](#)
- [2007](#)
- [2006](#)

[Learn More](#)

## calendar

Jamboree roster + fees due *fri 12 Oct 12*

iGEM wiki frozen + parts *fri postmarked 26 Oct 07*

Jamboree! *sat-sun 3-4 nov 07*

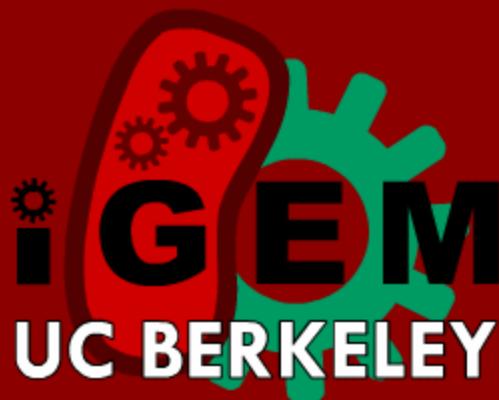
Registry + BioBrick foundation workshops *sun-tue 4-6 nov 07*







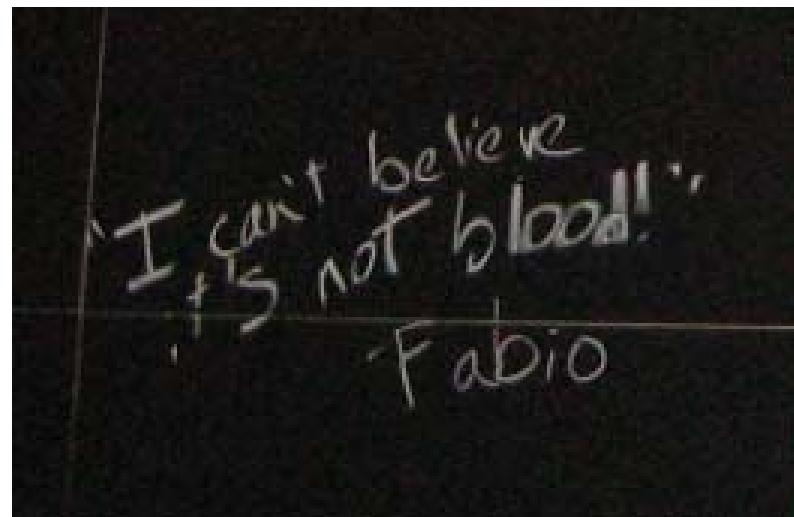
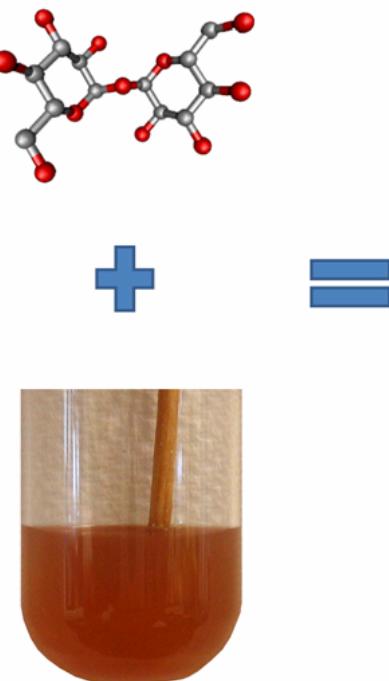


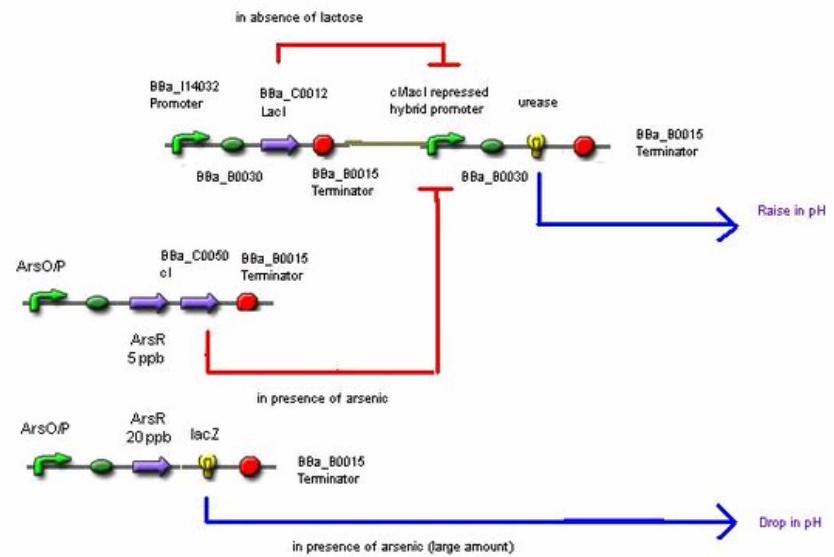


# BACTOBLOOD

The necessity of inexpensive, disease-free, and universally compatible blood substitutes is undisputed. There are currently no blood substitutes approved for use in the US or the UK, and whole blood is almost always in short supply. Developing countries have the greatest need for blood transfusions, yet many lack the necessary donation and storage infrastructure and the required pool of healthy donors. To address this problem, we are developing a cost-effective red blood cell substitute constructed from engineered *E. coli* bacteria. Our system is designed to safely transport oxygen in the bloodstream without inducing sepsis, and to be stored for prolonged periods in a freeze-dried state.

*Support for Berkeley iGEM 2007 was generously provided by SynBERC and The Camille and Henry Dreyfus Foundation, Inc.*







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**Sci-Tech**



Under a magnification of 6836x, this scanning electron micrograph (SEM) depicts a number of Gram-negative Escherichia coli bacteria of the strain O157:H7. (Courtesy of Centers of Disease Control and Prevention)

**Alberta team trying to turn E. coli into fuel**

Updated Mon. Nov. 5 2007 8:12 AM ET [The Canadian Press](#)

CAMBRIDGE, Mass. -- A University of Alberta team trying to turn E. coli into fuel has earned a first place finish in the energy category at an international genetic engineering competition at MIT.

Andrew Hessel, a consultant with the Alberta Ingenuity Fund who has helped teams from three Alberta universities compete, says while none of the teams from the University of Calgary, the University of Lethbridge or the U of A made it to the final competition, he calls it a fantastic educational experience.



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A JOINT PROGRAM OF THE CALIFORNIA INSTITUTE FOR QUANTITATIVE BIOMEDICAL RESEARCH (QB3)  
AND LAWRENCE BERKELEY NATIONAL LABORATORY (LBNL)

The California Institute for Quantitative Biomedical Research (QB3) and Lawrence Berkeley National Laboratory (LBNL) have joined forces to accelerate the growth of synthetic biology, a new field that promises major new advances in preventing and treating disease, generating new energy sources, and preventing and mitigating environmental threats.

Opening in spring 2005 in a spacious, modern building in west Berkeley, the Berkeley Center for Synthetic Biology gives renowned scientists and engineers the chance to pool their talents and collaborate in new ways, with enormous potential benefits for California's citizens in the form of advances in biomedicine and energy renewables and economic growth.

Synthetic biologists study the control and design of biological components and new organisms to solve a host of important health, energy, and environmental problems that cannot be solved using naturally occurring biological entities. The inherently



QB3 and LBNL scientists occupy lab space in a building renovated in 1997 for biotech research, previously leased by Bayer, featuring large labs, viral suites, and tissue culture rooms. UCSF Mission Bay and numerous biotech firms are nearby.

## MIT establishes groundbreaking biological engineering major

February 17, 2005

The Massachusetts Institute of Technology faculty yesterday approved a new course of study for undergraduates, in biological engineering, the first entirely new curriculum established at the Institute in 29 years.

MAY 3, 1982

\$1.50

# TIME

## COMPUTER GENERATION A New Breed of Whiz Kids

1.3%  
INFLATION VANISHES!  
At Least for  
A Month



# TRS-80 COMPUTER

CAT. NO. 68-2030

# whiz kids™

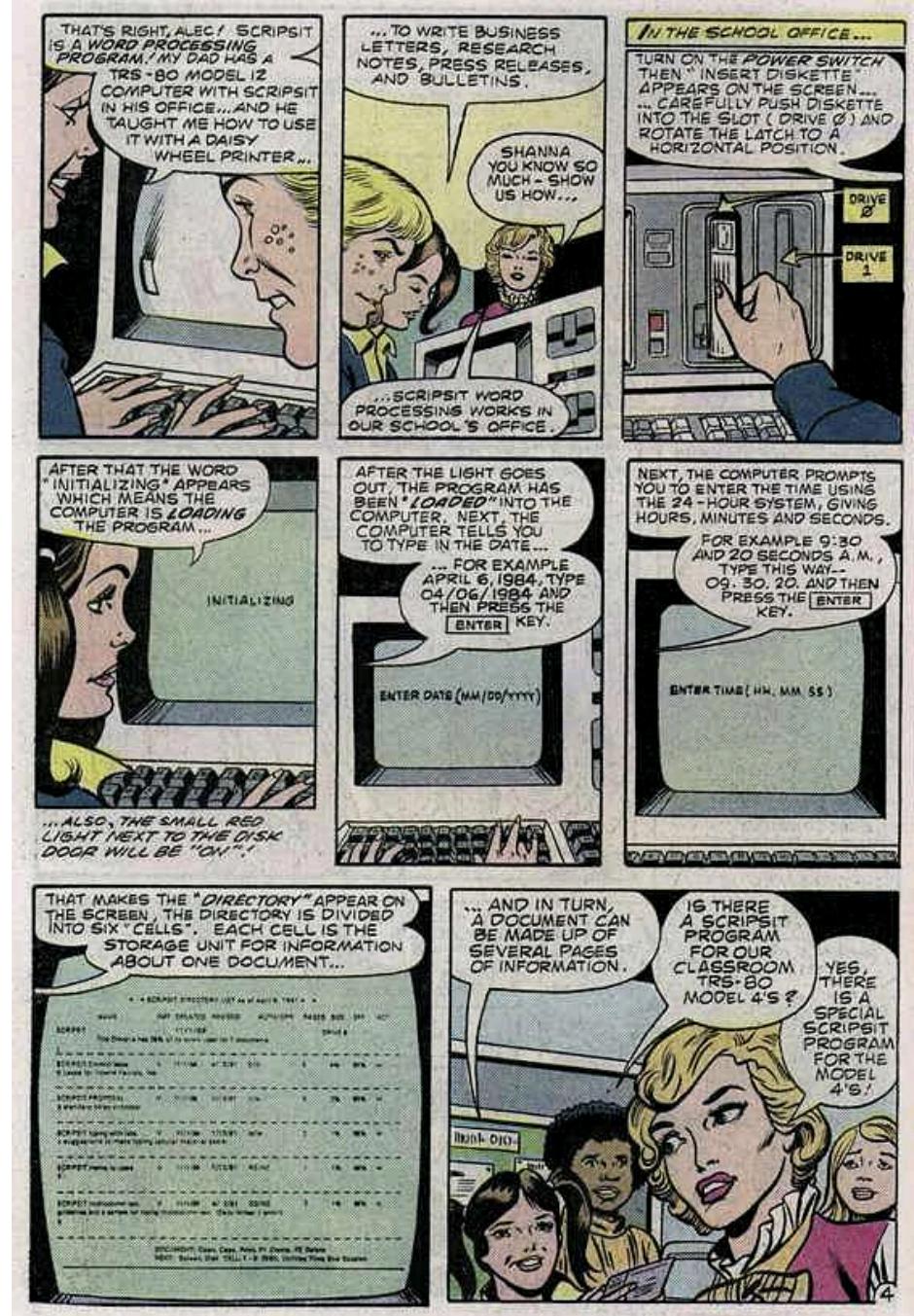
ALEC AND  
SHANNA  
STARRING IN

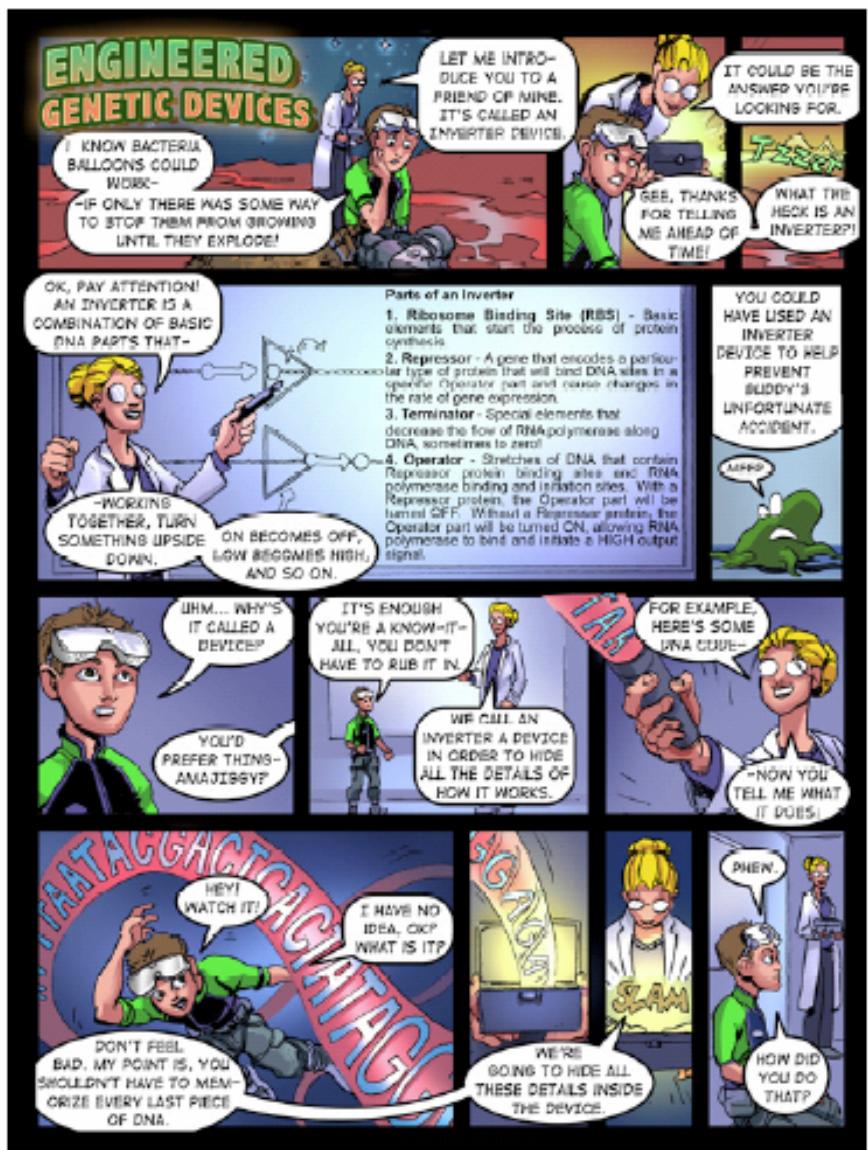
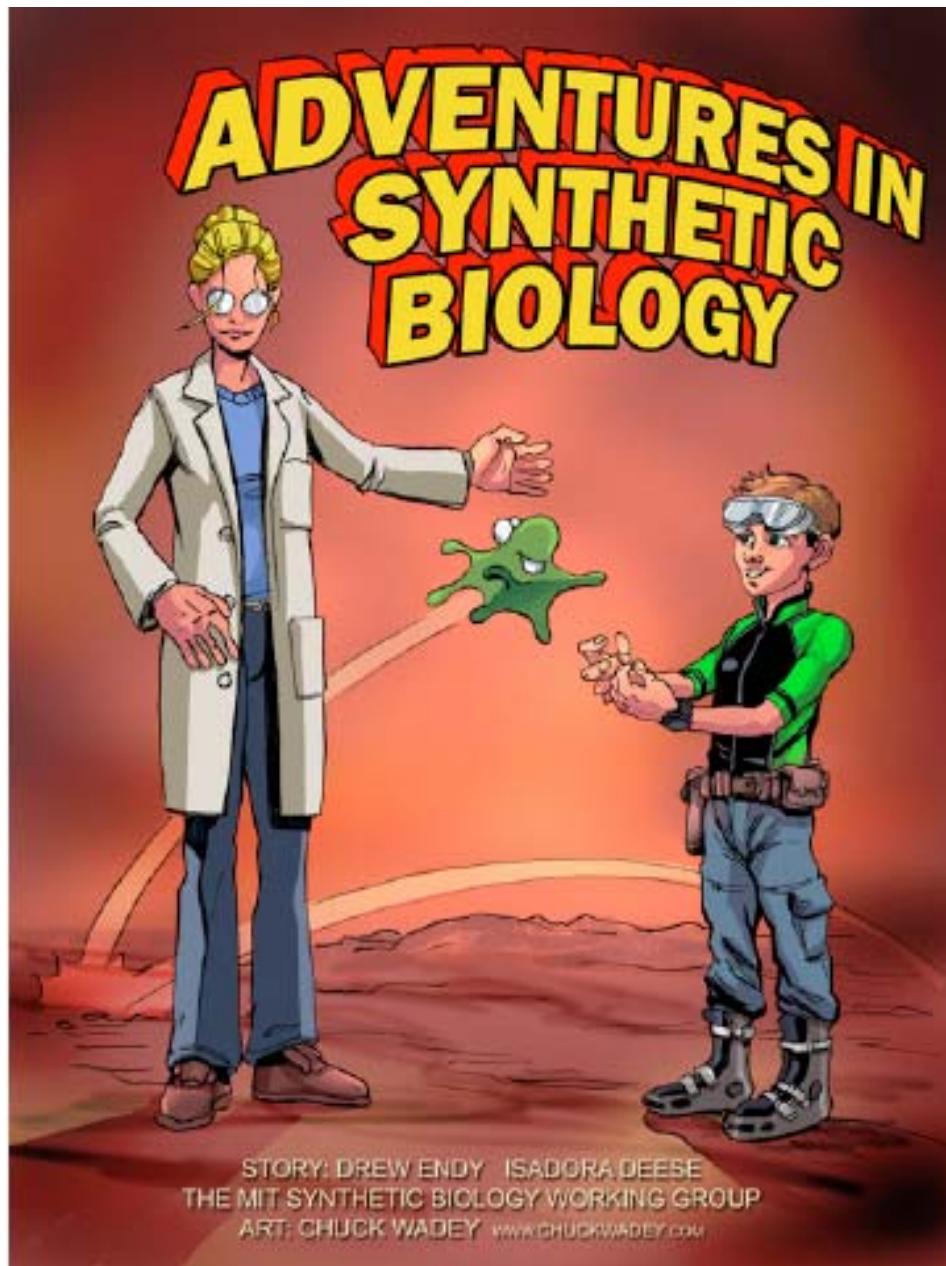
# THE COMPUTER TRAP

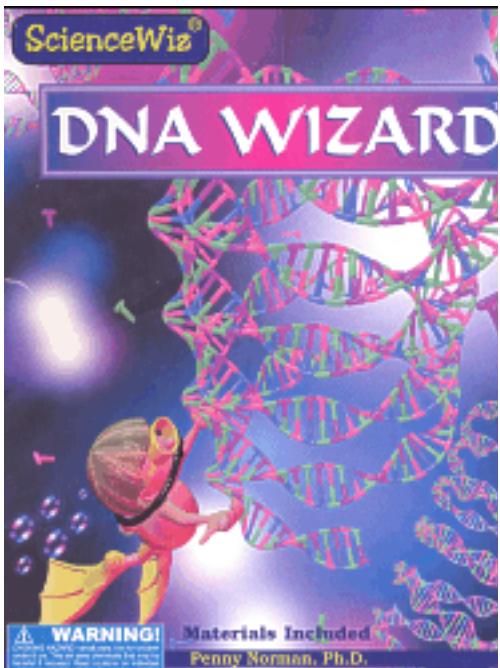
A comic book illustration of four people on a boat deck at night. A man in a green sweater is on the right, another man in a brown jacket is in the center, a woman in a purple sweater is on the left, and a man in a blue shirt is partially visible behind the woman. They are working with ropes and equipment. The background shows a city skyline across the water. A yellow starburst in the bottom left contains the text "COMPLIMENTS OF Radio Shack® The Name in Classroom Computing™" and "DICK AYERS AND GENE STONE". A small speech bubble from the woman says "I'm scared!".

COMPLIMENTS OF  
**RadioShack®**  
The Name in Classroom Computing™

DICK  
AYERS  
AND  
CHIC  
STONE







# Projects with DNA

For ages 8 and up  
Adult Supervision Required

Materials included except for the items listed.  
Through play, hands-on projects, patterns and puzzles  
this book and kit explores the amazing DNA story.

Build a DNA ladder.

Extract DNA

Heat SHOCK!

Decode the code of life

Grow glowing cells

Is it a boy or girl?

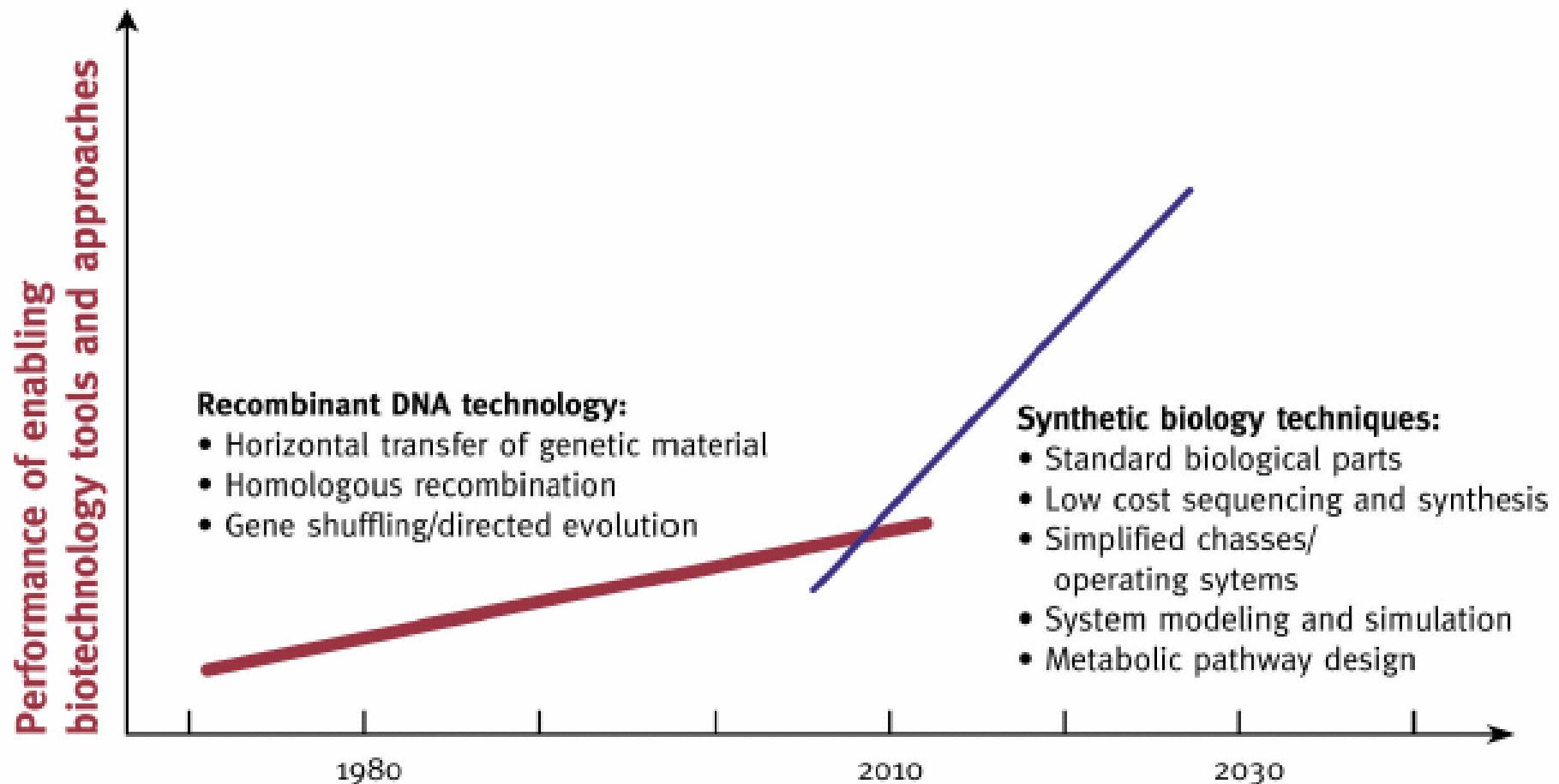
Ooey, Gooey, DNA!

Dress up for sterile techniques.

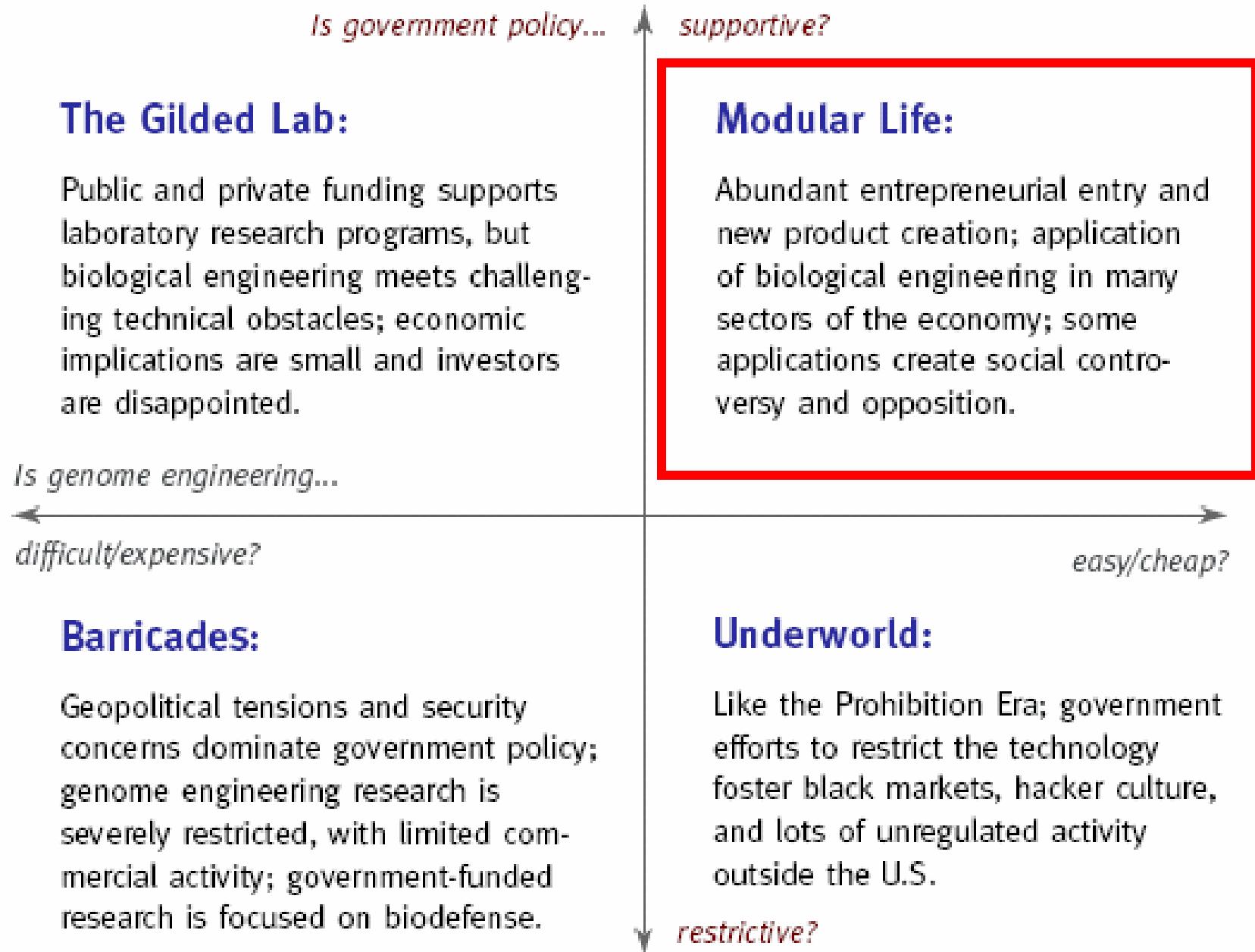
Quality time, quality learning, quality play.

Solve the chromosome puzzle.

Looking forward



Source: Bio era



Source: Bio era



Photo Credit: By Joby Warrick -- The Washington Post Photo

### Genome Transplantation in Bacteria: Changing One Species to Another

Carole Lartigue, John I. Glass, \* Nina Alperovich, Rembert Pieper, Prashanth P. Parmar, Clyde A. Hutchison III, Hamilton O. Smith, J. Craig Venter

The J. Craig Venter Institute, Rockville, MD 20850, USA.

## science theguardian

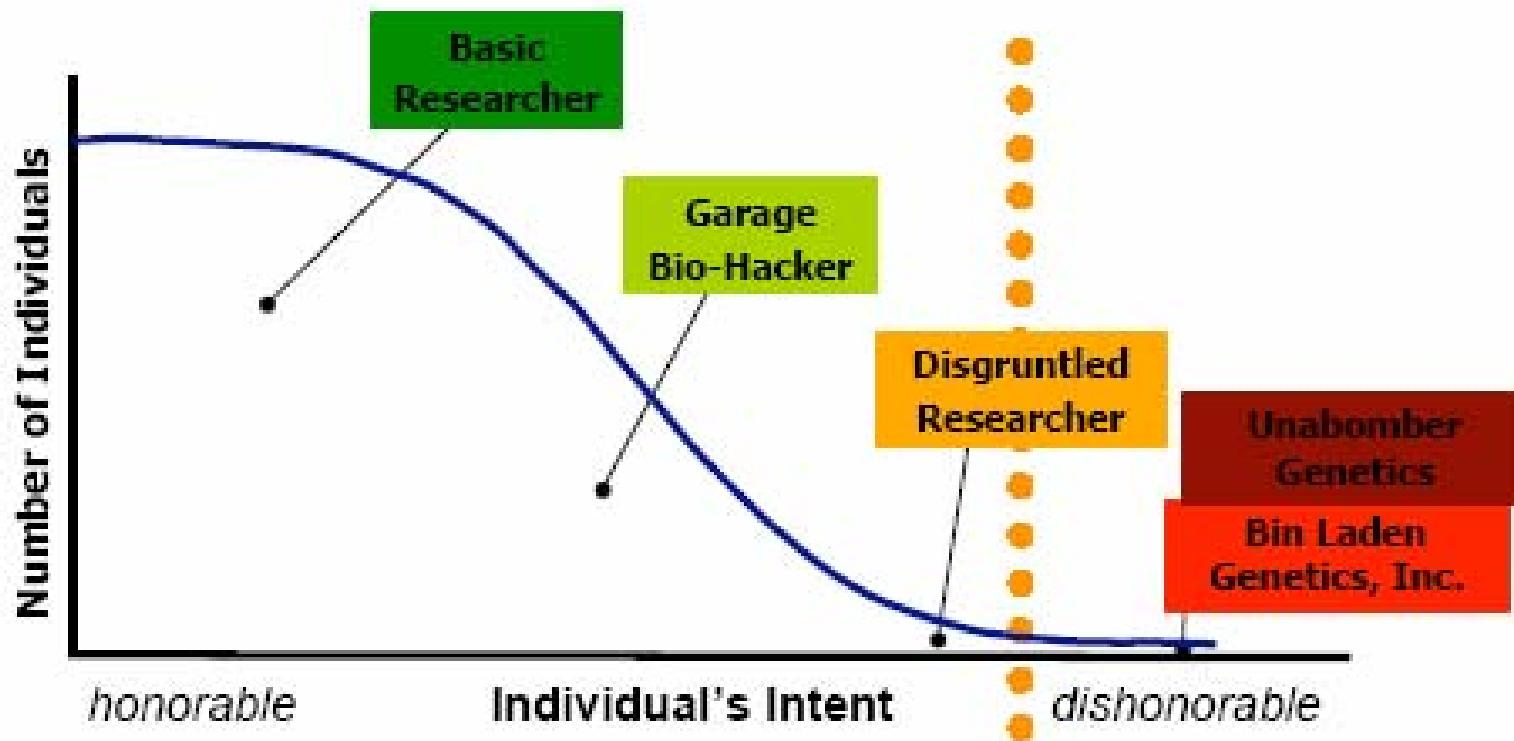
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## I am creating artificial life, declares US gene pioneer

- Scientist has made synthetic chromosome
- Breakthrough could combat global warming

## *Suite of solutions*

---



## SYNTHETIC GENOMICS | *Options for Governance*

---

Michael S. Gartland, The J. Craig Venter Institute, Rockville, Maryland; Drew Endy, Massachusetts Institute of Technology, Cambridge, Massachusetts; Gerald L. Epstein, Center for Strategic and International Studies, Washington, District of Columbia and Robert M. Friedman, The J. Craig Venter Institute, Rockville, Maryland

October 2009

J. Craig Venter®  

dicaprio

hanks



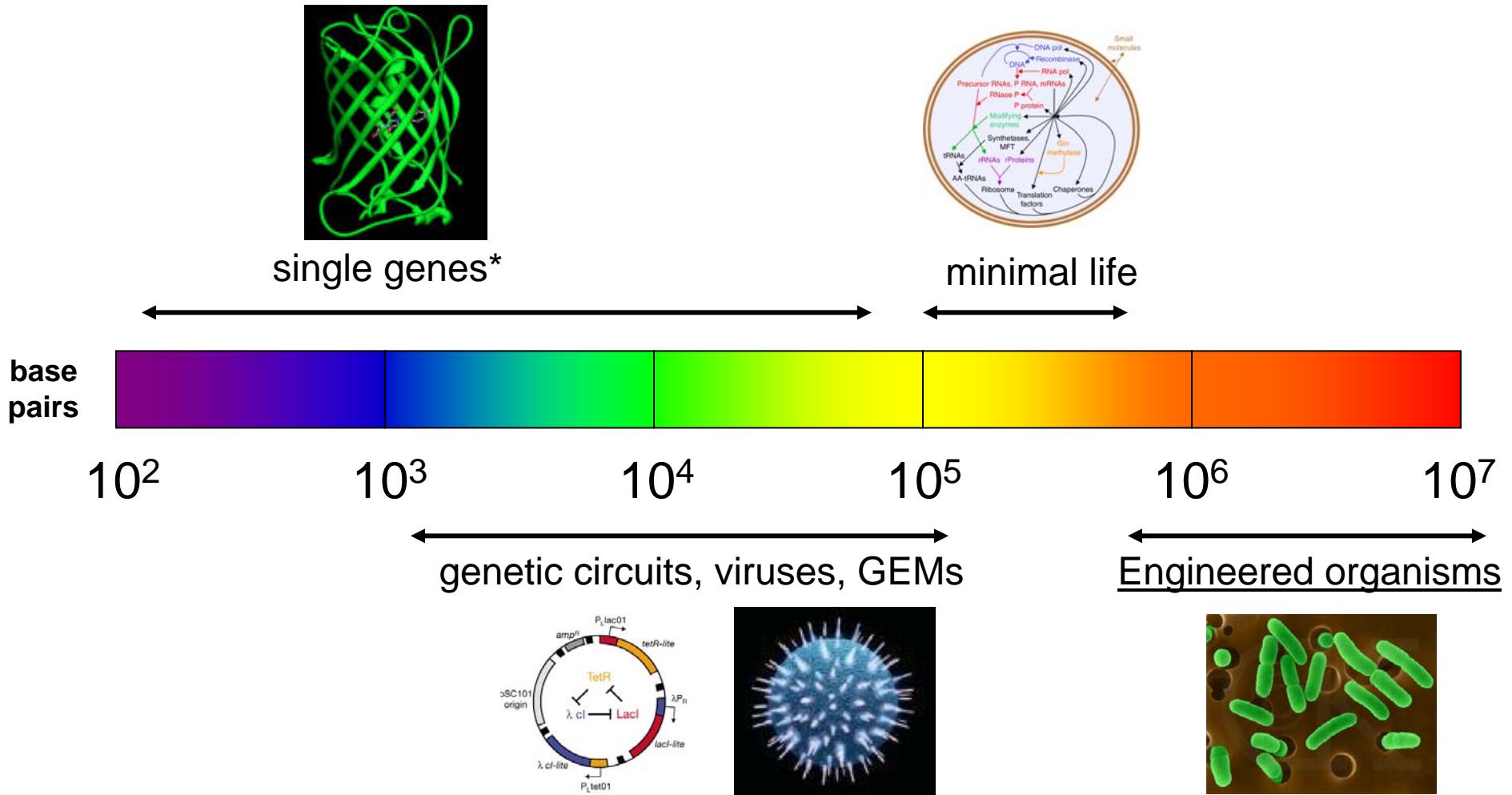
DELMARVUE PICTURES PRESENTS A KEMP COMPANY AND SPLENDID PICTURES PRODUCTION A PARKS/MACDONALD PRODUCTION A STEVEN SPIELBERG FILM  
LEONARDO DICAPRIO TOM HANKS "CATCH ME IF YOU CAN" CHRISTOPHER WALKEN MARTIN SHEEN NATHALIE BAYE DEBRA ZANE DEBORAH MIMS HANKIN  
MARK RYAN RYAN W. ABAGNALE WITH STAN REISBERG JOHN WILLIAMS BERNARD ZOPPES MICHAEL KAHN A.C.E. JEANNE OPPENWAL DANUSZ KAMINSKI A.S.C.  
PROD. DESIGNER DANIEL LIPINSKI CIN. DARRYL KEMP LAURIE MACDONALD MICHAEL SHANE AND TONY ROMANO EDIT. STEVEN SPIELBERG WALTER E. PARKES  
DIRECTOR OF PHOTOGRAPHY STEVEN SPIELBERG C.P. STEVEN SPIELBERG PICTURES

Catch them in Cinemas

**Q&A: Former fraudster Frank Abagnale offers IT security advice**

Next generation biotechnology

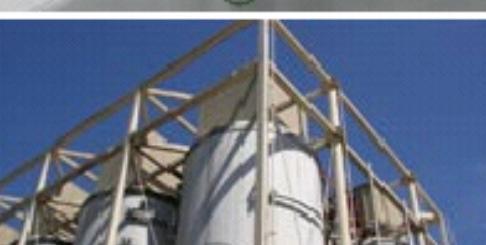
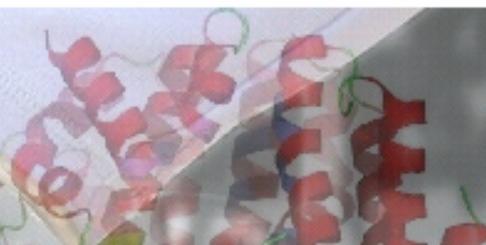
# Applications dependent on synthetic capabilities





# LS9, INC.

the renewable petroleum company™





Realizing the Promise of Synthetic Biology

COMPANY PROFILE

TECHNOLOGY

CURRENT PROJECTS

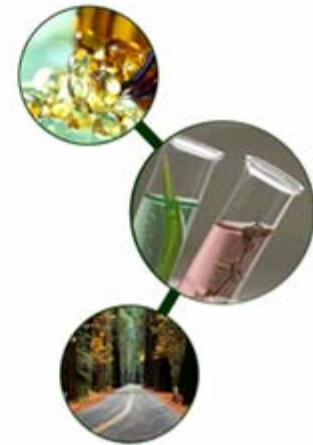
NEWS

CAREERS

## Welcome



Amyris Biotechnologies is translating the promise of synthetic biology into solutions for real-world problems. Building on advances in molecular, cell and systems biology, we are engineering microbes capable of producing high-value compounds to address major global health and energy challenges. We are employing these living chemical factories to produce novel pharmaceuticals, renewable fuels, and specialty chemicals.



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## Stop Cloning Genes SYNTHESIZE THEM!

Rapid, high-quality,  
low-cost gene synthesis



Gene  
Synthesis



Constructed  
Variant  
Libraries



Constructed  
Operons &  
Operon Variant  
Libraries



Genomic  
Engineering





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## Investors

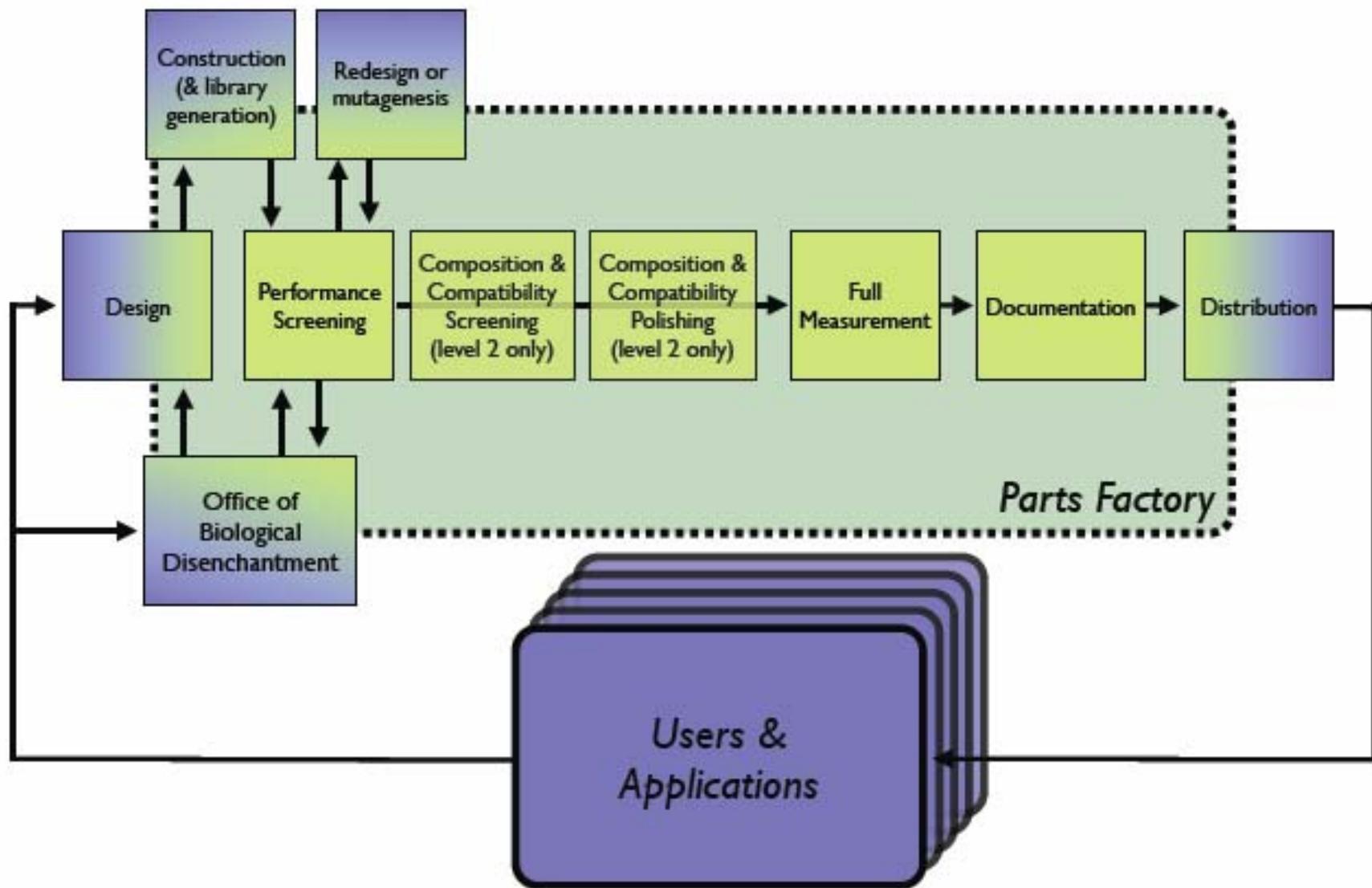
**Funded by a syndicate of well-respected, top-tier venture capital firms, Archemix has raised over \$100 million to date.**

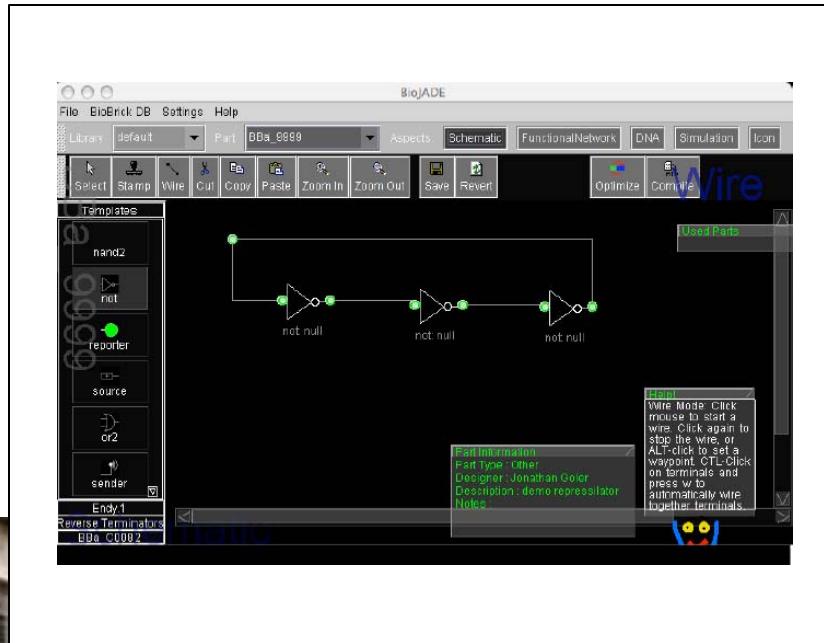
The Series B round (\$53M, completed 1Q 2004) was led by Highland Capital Partners, and included all of the Series A investors and Athenian Venture Partners.

The Series A round (\$52M, completed 3Q 2002) included lead investors Atlas Venture, Prospect Venture Partners and SV Life Sciences. Other investors included Rho Ventures, Care Capital, MDS Capital, POSCO BioVentures, and US Trust Private Equity.

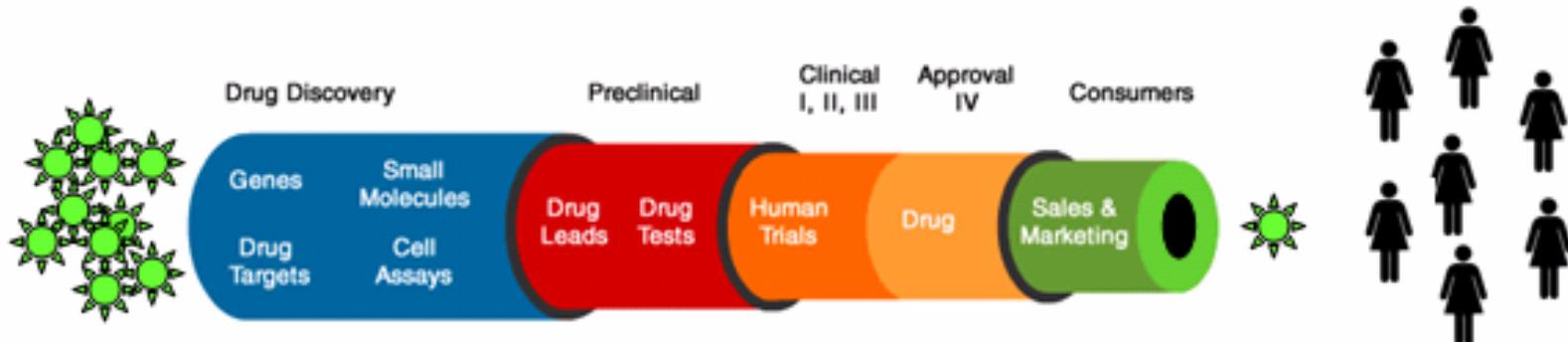
## Archemix and Merck KGaA Sign Strategic Alliance

*Collaboration to Focus Primarily on Cancer Therapeutics Using Novel Aptamer Technology*



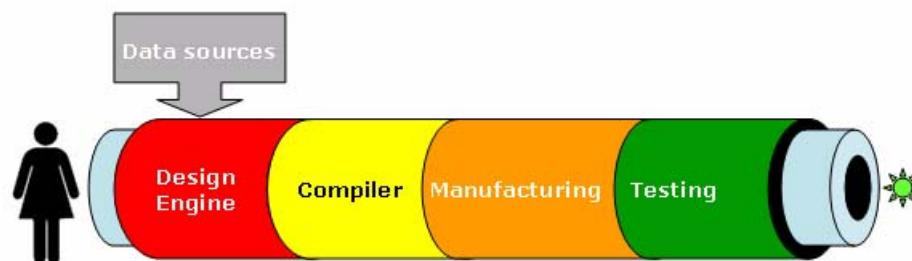


## Toward a New Drug Development Pipeline Suitable for Personalized Cancer Therapeutics

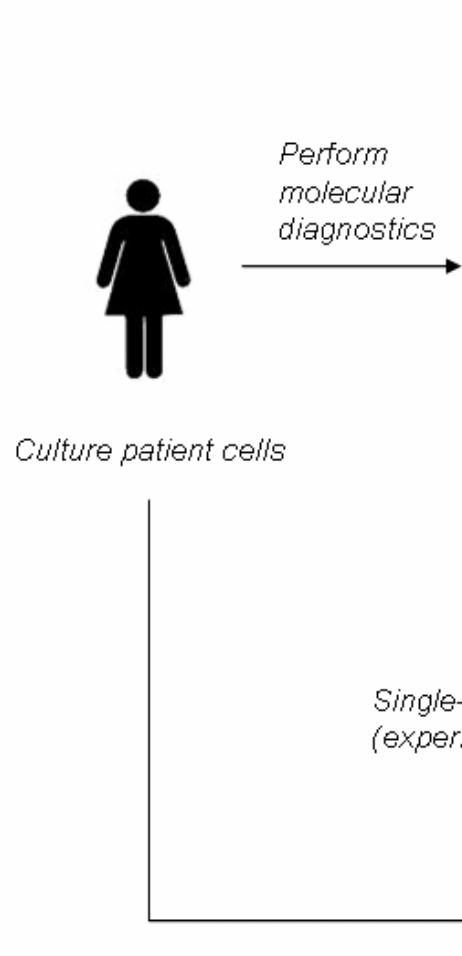


Ideally, a next-generation pipeline for effective cancer drugs would:

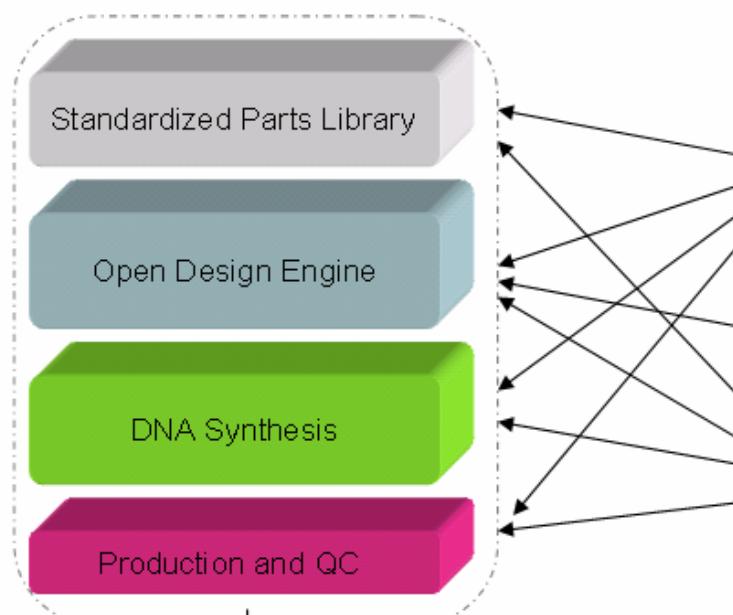
- As an output, produce personalized medicines
- Be able to produce a targeted medicine for every cancer, newly diagnosed or recurring
- Allow for rapid and sustainable development of these outputs
- Have favorable drug development economics, ie. the cost of producing drugs decreases over time
- Make a failure of any drug in the pipeline a trivial event
- Be able to grow in output capacity to satisfy the needs of a global market
- Continuously incorporate the latest understanding and data from molecular biology, oncology, and related sciences, plus growing clinical experience with personalized medicines
- And, finally, ensure the highest possible standards of safety and oversight



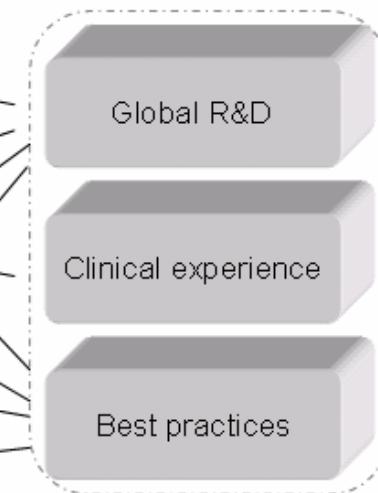
## Clinical presentation



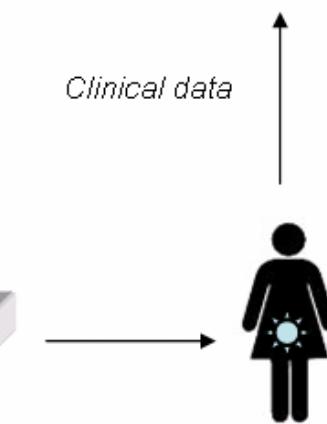
## Open Bio-Fabrication

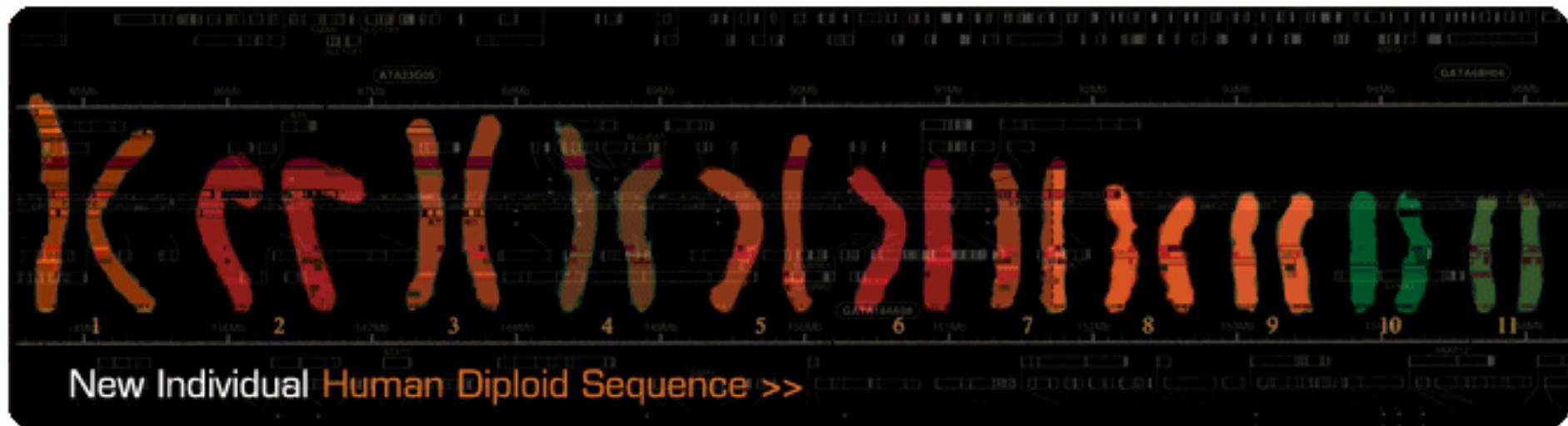


## Shared knowledge base



*in vitro testing should show drug affects only cancer cells, otherwise failures analyzed and design process repeated*





ARCHON GENOMICS  PRIZE

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the X PRIZE about each of us.

Revolution Through Competition. 

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1866: Gregor Mendel discovers the laws of inheritance.

200,000 years ago: *Homo sapiens* walks the Earth.

2003: The Human Genome Project maps a single person's genome.

**2007: 23andMe introduces the first Personal Genome Service.**

Unlock the secrets of your own DNA. Today.

175,000 years ago: The mother of all present-day humans is born in Africa.

1953: Watson and Crick uncover the double-helix structure of DNA.

Welcome to 23andMe, a web-based service that helps you read and understand your DNA. After providing a saliva sample using an at-home kit, you can use our interactive tools to shed new light on your distant ancestors, your close family and most of all, yourself.

## Part 1: What are genes?



Find out about the basics of cells, chromosomes, and the genes contained in your DNA.

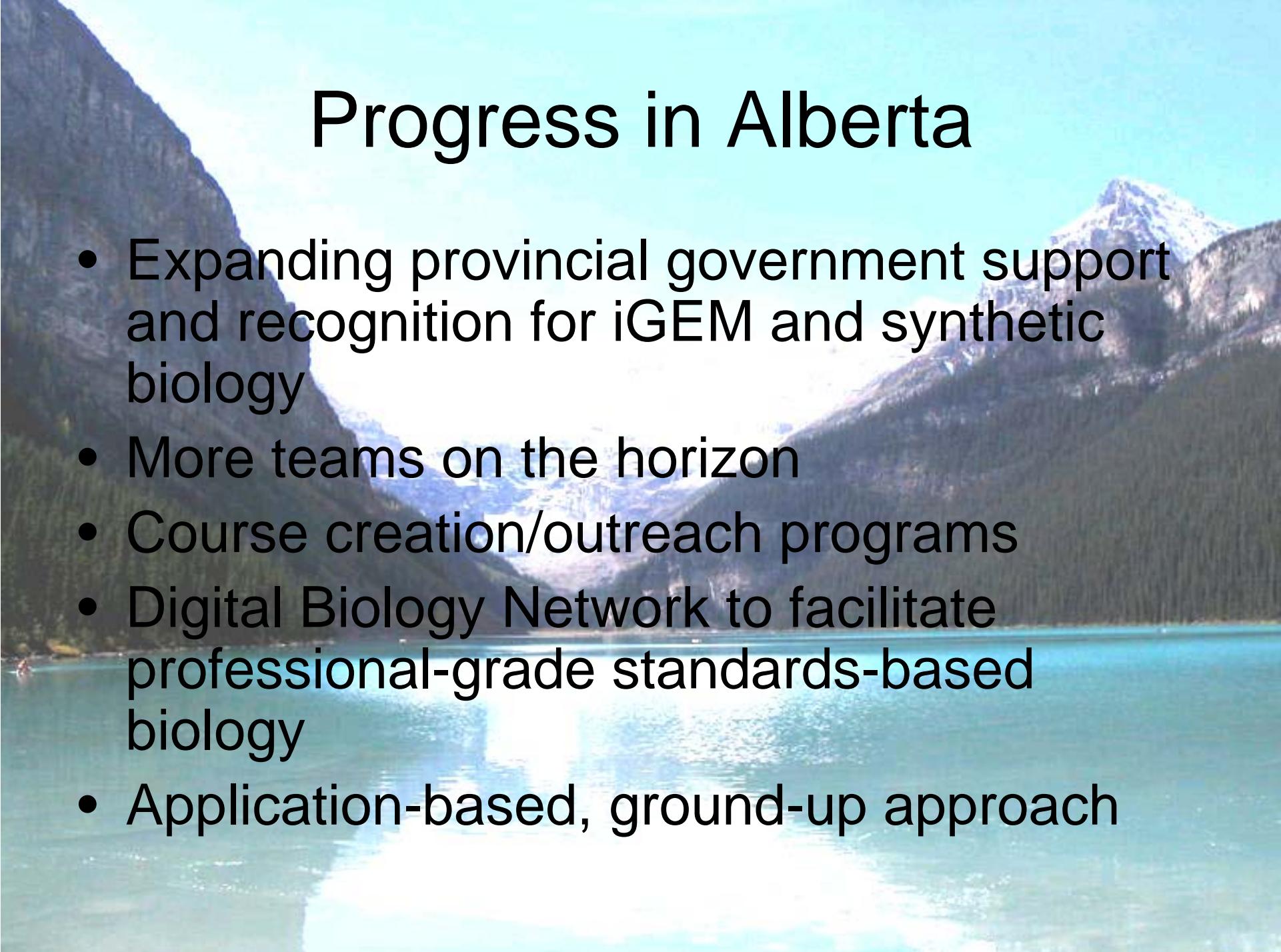
## Part 2: What are SNPs?



Learn about the variations in human DNA called SNPs, and how they can be used to understand relationships between people.

# Opportunities for Canada if..

- Can think **BIG** and move **Fast**
- *Tear down barriers* between professions and institutions and think global in scale
- Streamline and reform biological engineering education
- Invest in core resources (informational and physical)
- Select engaging and valuable applications where it can become recognized leader
- Support next-generation industry development



# Progress in Alberta

- Expanding provincial government support and recognition for iGEM and synthetic biology
- More teams on the horizon
- Course creation/outreach programs
- Digital Biology Network to facilitate professional-grade standards-based biology
- Application-based, ground-up approach



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Alberta Ingenuity operates  
the Alberta Heritage Foundation for Science and Engineering Research,  
a \$1 billion endowment to support research.

# Ingenuity

NEWS

**Ingenuity launches bold new \$100 million research program**  
October 10, 2007 - Ingenuity Accelerators focus on ensuring future prosperity of the province.