

#### Synthetic Biology and Sustainability

Presentation at 1<sup>st</sup> Interdisciplinary Symposium:

Global Sustainability – A Nobel Cause

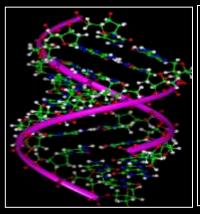
Potsdam, FDR 10 October 2007

Dr. George Poste, Director, Biodesign Institute
Arizona State University

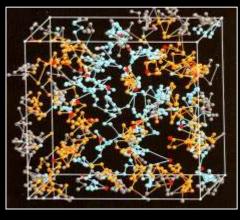
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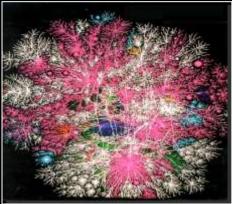
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- elucidation of the design principles and 'rule sets' of biological organization
- design and construction of novel biological functions and life forms

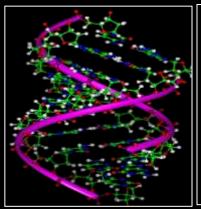


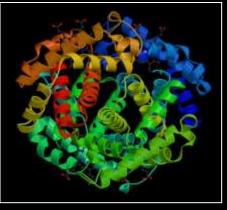


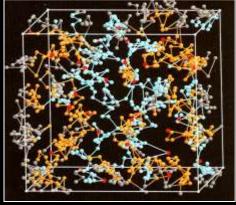


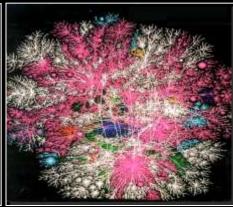


- elucidation of the design principles and 'rule sets' of biological organization
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 understanding the encoded information content and information flow in biological systems

## Biological Design: "Endless Forms Most Beautiful": Limitless Diversity From Combinatorial Assemblies of Limited Building Blocks



## Defining the Rule Sets for Biological Design, Assembly and Function

- common genetic (digital) code in all life forms
- genomes encode a limited series of structural building blocks (protein motifs and programmed assembly)
- combinatorial assembly of protein building blocks generates extravagant structural and functional diversity



### **Exploring Biospace The Design Power of Combinatorial Interactions and Assemblies**

- 30,000 genes
- two genes cooperate to create a function
   =(30,000 x 29,999)/2 = 449,985,000 potential combinations
- 100 genes generate a complex function =10<sup>289</sup> potential combinations
- if any combination of genes can generate a function
   = 2 x 10<sup>72,403</sup> potential combinations
- number of theoretical possibilities for synthetic assembly (biospace) far exceed narrow molecular space sampled in evolutionary time

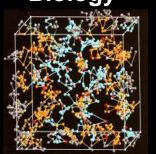
- matching the versatility, elegance and inventiveness of natural systems
- expanding the dimension of explored biospace
- predictive biology: design, simulation and construction of novel functions/organisms

## Synthetic Biology: Inter-disciplinary Convergence and Complex Policy Issues

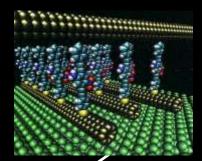
Systems Biology



Computational Biology



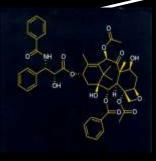
Materials and Nanobiotechnology



Industrial Bioprocess Engineering



#### **Specific Applications**



Occupational Safety



Dual-Use Applications

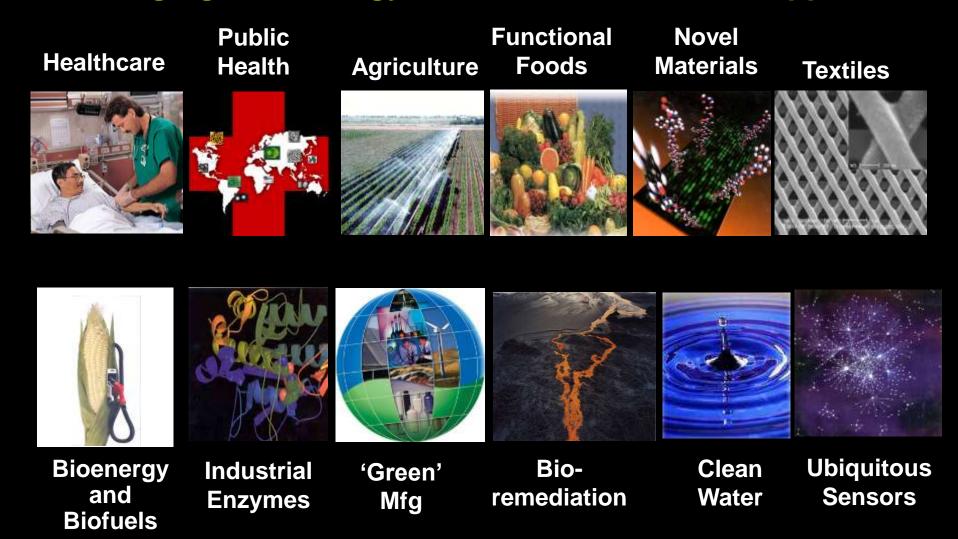


Public and Media Responses



Public Policy and Regulatory Oversight

## Synthetic Biology: An Emerging Technology with Diverse Industrial Applications



Identification of Relevant Genetic Source Code

Minimum
Genomes
(Universal
Producer
Cell)

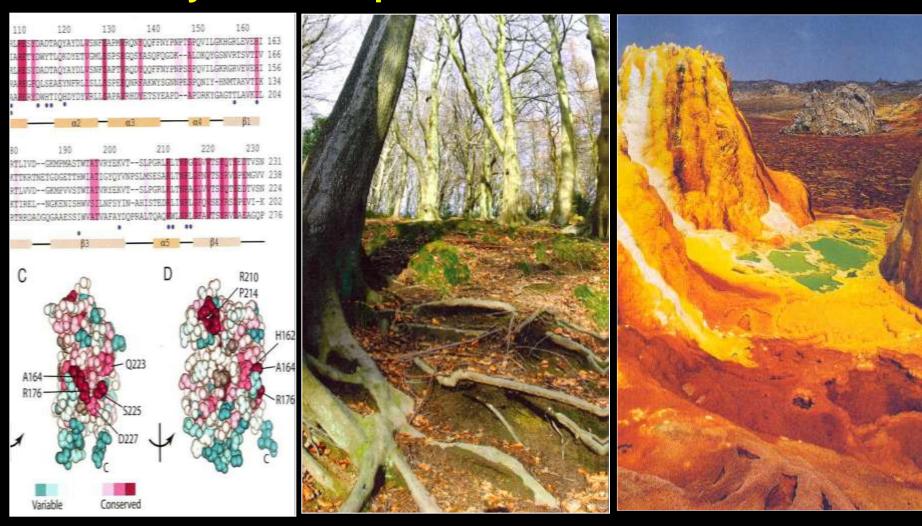
Control
of
Gene
Expression
Networks

 metagenomic screening for desired trait(s)

- "plug and play" programmed expression of gene "cassettes"
- programmed biosynthesis
- induction of novel properties in specific cells/tissues/ organisms

#### **Ecogenomics:**

## Mapping the Extraordinary Genomic Diversity and Biosynthetic Capabilities of Microbial Life



genomics eco-niches extremophiles

## Metagenomics: Sampling the Extravagant Functional Diversity of Microorganisms

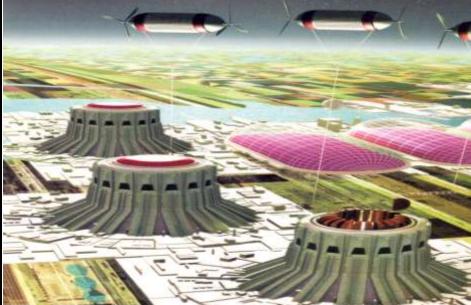
- estimated 100 billion microbial species
- only 6000 species cultivated and characterized
- massive repertoire of uncharacterized genes/proteins/metabolomes
- metagenomic sampling
  - mass screening of complete genomes of unknown/unculturable organisms
  - high throughput profiling to identify transfer of gene(s) with desired function into 'universal acceptor' organisms

#### The Quest for Sustainable Energy Sources



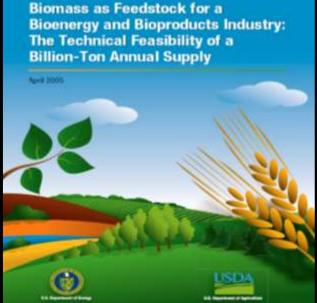


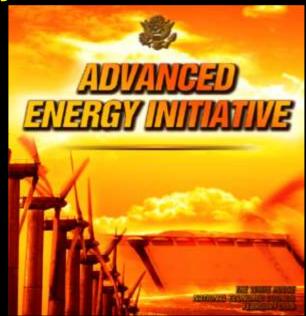




#### **Bio-Inspired Systems for Energy Production**













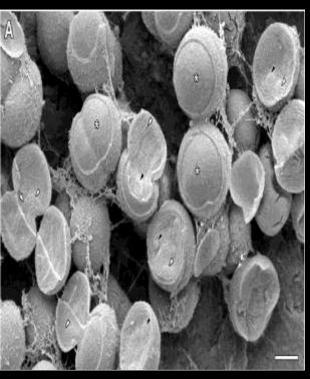
#### Production of Biodiesel from Biomass: A Widely Differing Yield Spectrum\*

Source	Yield	
	US gal/acre	L/Km <sup>2</sup>
<ul><li>Soybean</li></ul>	40-50	35-45,000
<ul><li>Rapeseed</li></ul>	110-145	100-130,000
<ul><li>Mustard</li></ul>	140	130,000
Jatropha	175	160,000
• Palm oil	650	580,000
<ul><li>Photosynthetic Microbes</li></ul>	10-20,000	9-18,000,000

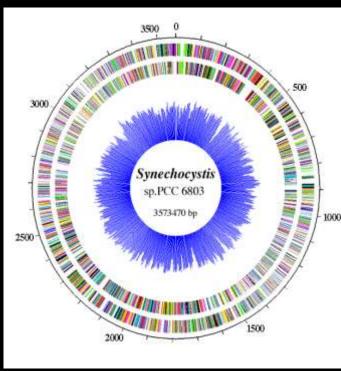
<sup>\*</sup>http://en.wikipedia.org/wiki/Biodiesel



## Biomass to Energy: The Merits of Cyanobacteria as a Feedstock







- well characterized, tractable genome for trait enhancement
- no celluose
- compositional homogeneity
- rapid assay of genetic modifications

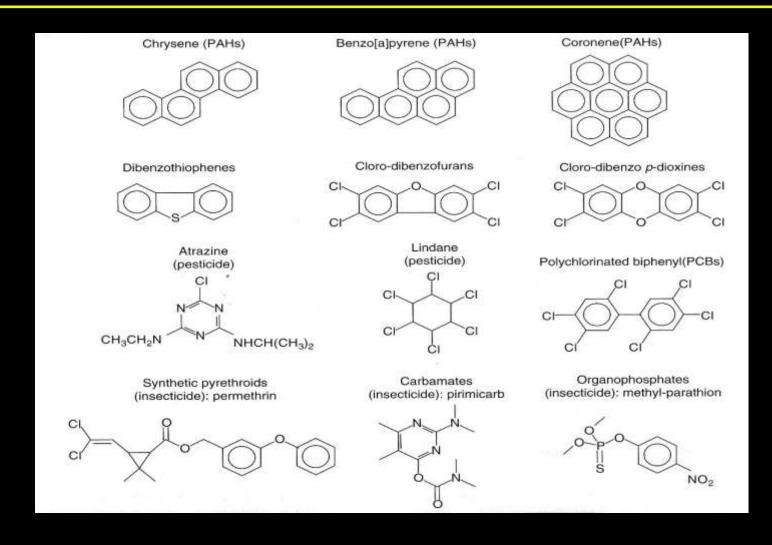
- siting flexibility
- no or limited transport costs
- CO<sub>2</sub> fixation



## Water Supply and Safety A Source of Future Geopolitical and Economic Instabilities



#### Microbial Genomics and Synthetic Biology: New Technology Platforms for Bioremediation and Improved Efficiency of Wastestream Management



#### Bio-Inspired Engineering: Removal of Perchlorate Contamination by Chemical Reduction by Biofilms of Hydrogen Producing Bacteria

- ✓ pilot scale project at La Puente, CA (400 L/min)
- influent: 25 mg/L of NO<sub>3</sub><sup>-</sup> and
   60 μg/L ClO<sub>4</sub><sup>-</sup>
- √ ≈ 95% removal of ClO<sub>4</sub>-, from 60 μg/L to below the CA action level of 4 μg/L
- $\checkmark$  ≈ 98% NO<sub>3</sub><sup>-</sup> removal of 25 mgN/L to ≈ 0.5mgNO<sub>3</sub><sup>-</sup>/L
- √ essentially 100% H₂ usage



#### e.Waste



- US produces 2 million tons per year
- escalating toxicity in developing countries with outsourced salvage activities
- lead, cadmium, mercury and flame retardants
- EC Restriction of Hazardous Substances Directive (ROHS)

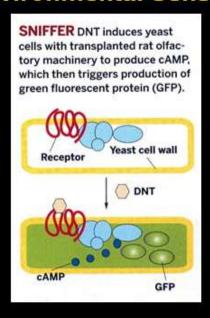
#### Biological Sensor Systems for Environmental Monitoring and Ecosystem Status

Genetically-Engineered Arabidopsis thaliana
Change-Color when Exposed to Landmine
Degradation Products



Aresa Biotection, Demark

#### Engineering Sentinel Organisms as Environmental Sensors



D. N. Dhanasekaran et al (2007) Nature Chem. Biol.

DOI: 10.1038/nchembio882

#### **EIDs: Global Reach and Global Consequences**



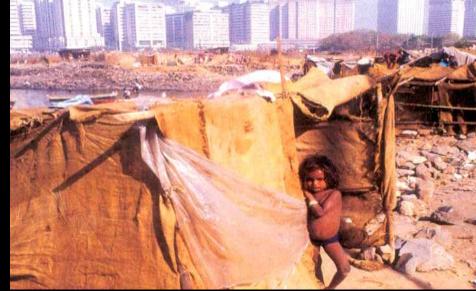






## The Lack of Public Health Infrastructure in Developing Countries: Urbanization and New Zoonoses

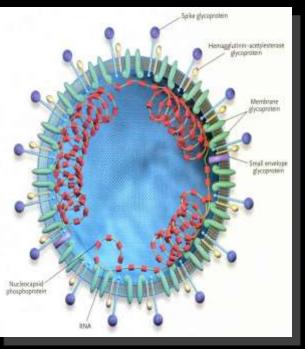


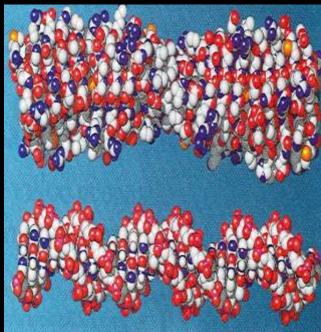


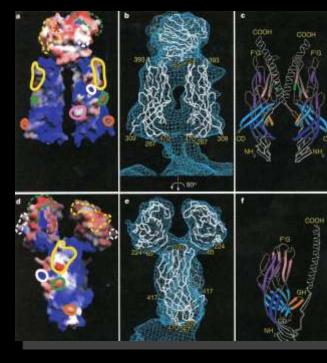




# Engineered Microorganisms and Cells as Novel Delivery Systems for Drugs and Vaccines







#### No One Walks Alone

- every adult human carries two pounds of bacteria
  - 100 trillion non-human cells
- microorganisms as key substrates for therapeutic/diagnostic interventions
  - in-body sensors
  - regulated production of biomediators
  - gene-centered implants, copies, upgrades
- genetics will be scaleable and upgradeable
- plug and play genetics : the "undo" button



## **Synthetic Biology and Engineering Enhanced Traits** in Food, Feed and Fiber Products



## Directed Evolution and Design of Novel Enzymes for Catalysis of Lignocellulosic Biomass



- cellulosic energy crops
- agricultural, forestry and mill residues
- food processing residues
- municipal solid waste
- non-recycled paper
- construction and demolition wood

#### Synthetic Biology and Novel Industrial Process Chemistry

The Natural World (Evolutionary Biospace)

The Synthetic World (Exploring Unexplored Biospace)

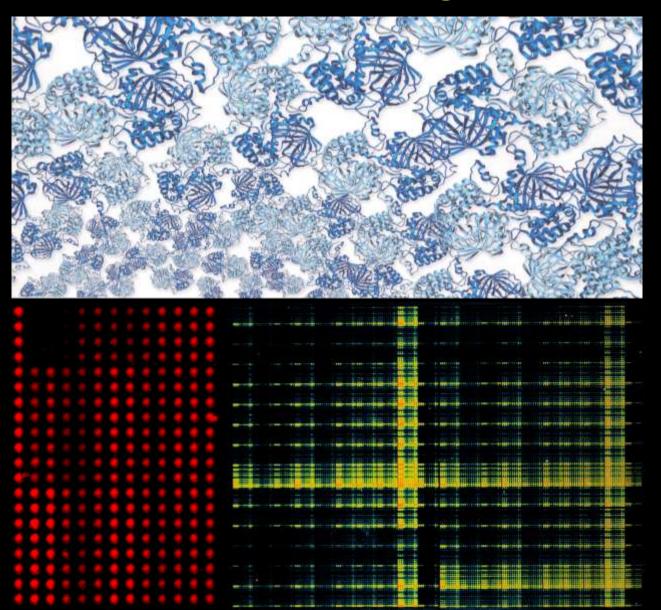
mapping protein families and functional design

metagenomics
and
expanded
knowledge
of
evolved
'biospace'

directed
(accelerated)
evolution
and
expansion of
novel
'biospace'



#### **Accelerated Evolution and Design of Novel Catalysts**



# Exploring Biospace: Iterative Selection of Novel Variants for Substantial Functional Performance Improvements Versus Naturally Occurring "Parent System"





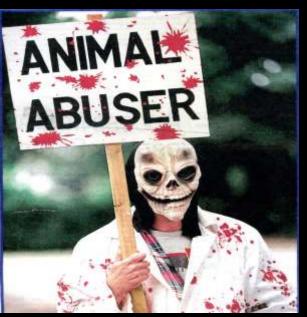


#### **Activity Versus Starting Material**

	Reduced	Same	Increased
cycle 1			
cycle 2			
cycle 3			
cycle 4		•	



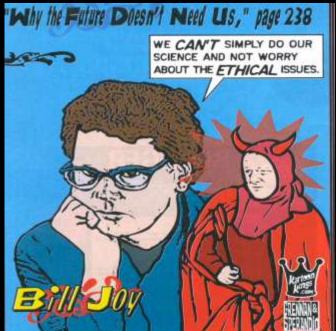
#### **Controversy and Divisiveness of New Technologies**











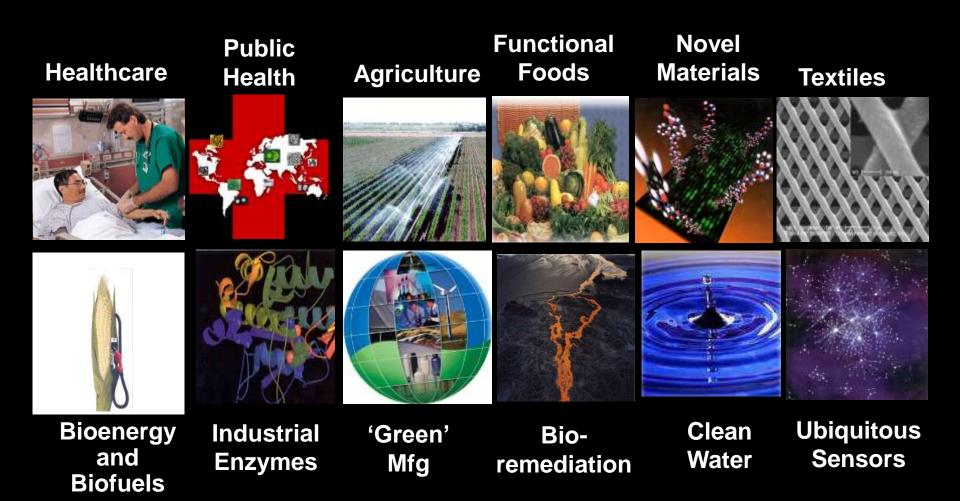


- complex policy issues
- design of novel life forms and societal response
- dual-use applications
- public and media attitudes to perceived risks and benefits
- proactive, predictable, evidence-based regulatory frameworks

#### The End of the Darwinian Interlude

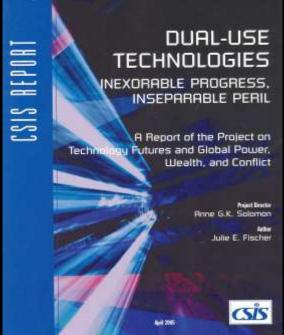
- early 'biotic' world
  - massive lateral gene transfer and 'loose' definition of species
  - rapid evolution as a communal affair
- the Darwinian interlude
  - majority period in evolution (3 billion plus) years
  - slow pace of change and species 'isolationism'
- synthetic biology
  - revival of pre-Darwinian era of horizontal gene transfer

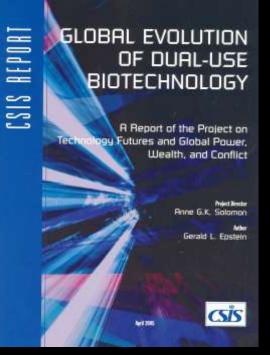
emerging technology with myriad applications across diverse industrial sectors



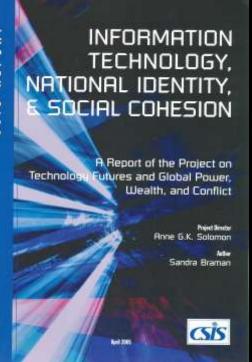
- new bioprocess technologies based on the 'design principles' of biological systems will progressively transform industrial manufacturing methods
  - cost-effective use of renewable resources
  - reduced depletion of scarce/non-renewable resources
  - mitigation of adverse environmental impacts via bioremediation of waste streams and contaminated environments
  - economic and security benefits of bioenergy and biofuels
  - novel goods and services

- anticipated powerful driver of industrial innovation and market disruption
- new aspirants, new cross-sector relationships and new markets
- the next era in the evolution of human mastery of the environment
  - agronomic, industrial, informational, genetic, biomimetic, designed life forms
  - enhancement and eugenics?





CSIS REPORT



CSIS REPORT

AND GLOBAL POWER,
WEALTH, AND
CONFLICT

A Report of the Project on
Technology Futures and Global Power,
Wealth, and Conflict

Philit Vector and Lieber
Annie G.K. Soloman

TECHNOLOGY FUTURES

