

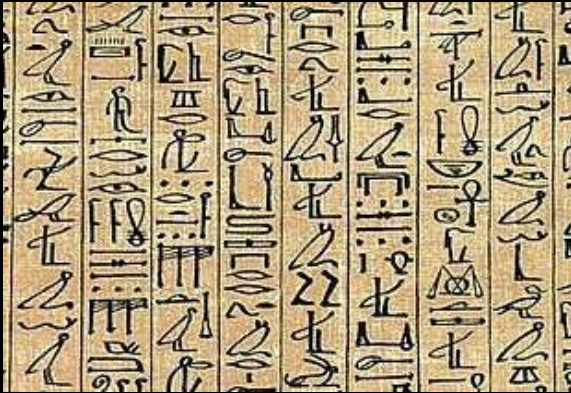
# **From the Quest for Fire to GATTACA: Systems and Synthetic Biology as the Next Wave of Technology Disruption in the Anthropocene**

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**Bethesda, Maryland**  
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# The Journey to the Anthropocene: Mastery of Increasingly Sophisticated Intellectual Challenges and Technological Acceleration

**First Communication  
Revolution 70K YBP**



**Agrarian Revolution  
11K YBP**



**Scientific Revolution  
0.5K YBP**



**Industrial Revolution  
0.25K YBP**



**Digital Revolution  
0.1K YBP**



**Molecular Biology  
Revolution 0.05K YBP**

# **Systems and Synthetic Biology as the Next Major Wave of Technological Disruption in the Anthropocene**

**Understanding the Instructional Rules for Construction and Control of Complex Biological Systems**

**Exploring Biospace:  
Engineering Novel Biological Functions**

**Directed Evolution and Accelerated Evolution:  
From Individual Molecules to Organisms**



# Biological Diversity and Variation: “Endless Forms Most Beautiful”





# **“Endless Forms Most Beautiful”**

## **Systems and Synthetic Biology and Exploring Biospace**



**Form and Function  
(diversity)**

**Instructional Information  
(code)**

**Systems Design  
(rule sets)**

**Design of Novel  
Biological Systems  
(exploring biospace)**



# “Endless Forms Most Beautiful”

## Systems and Synthetic Biology and Exploring Biospace

### Form and Function

- diversity (species)
- variety (individuals)
- specialization (cells)
- hierarchical integration (control systems)

### Instructional Information

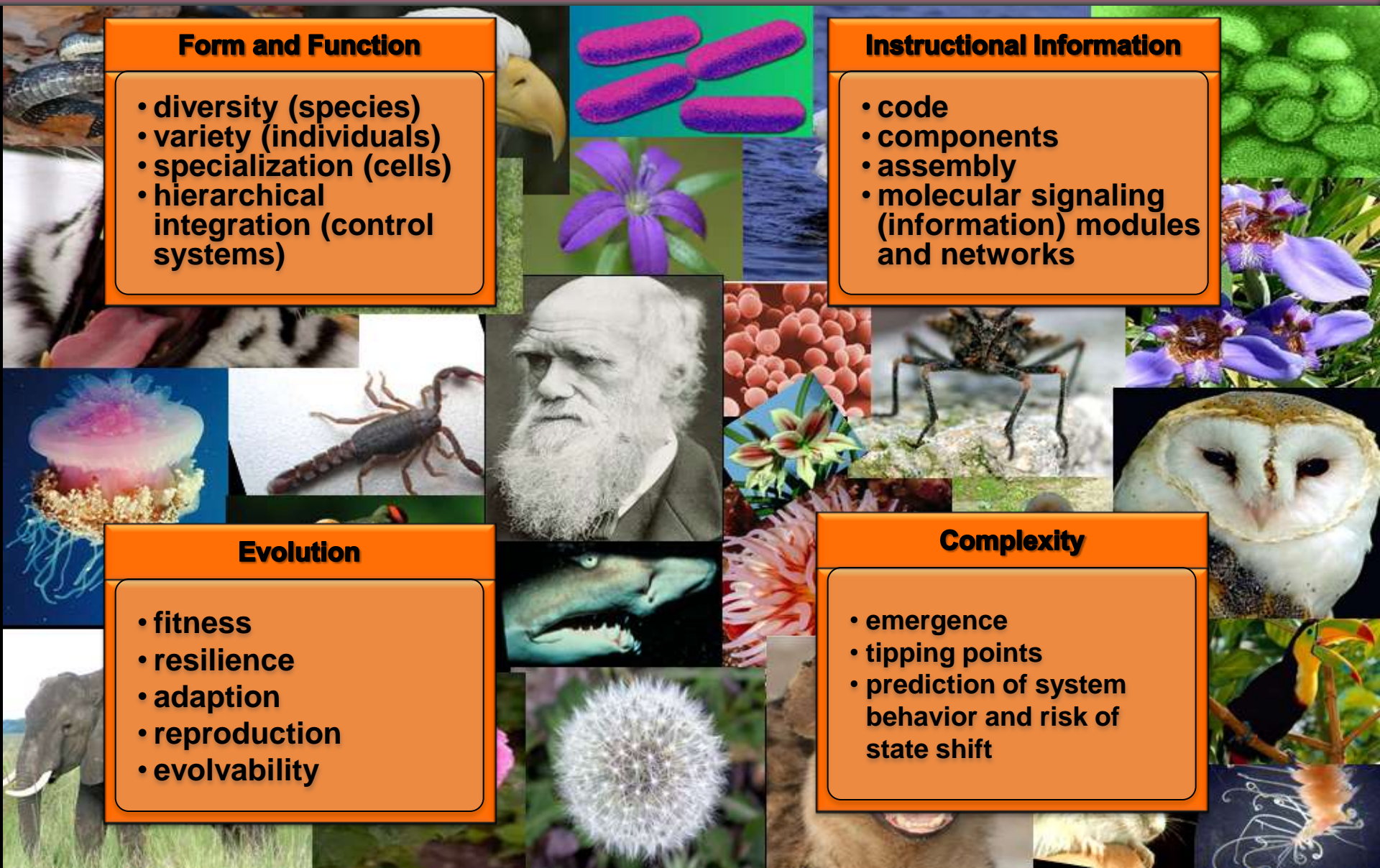
- code
- components
- assembly
- molecular signaling (information) modules and networks

### Evolution

- fitness
- resilience
- adaption
- reproduction
- evolvability

### Complexity

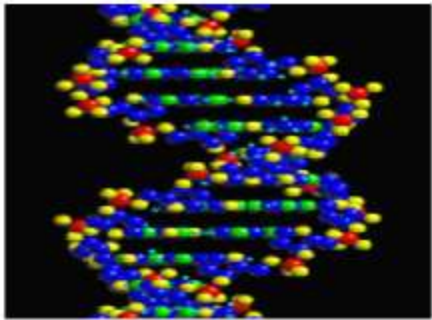
- emergence
- tipping points
- prediction of system behavior and risk of state shift





# **The Economy of Biological Organization: Defining the Common Building Components for Biological Systems and Combinatorial Assembly to Generate Functional Diversity**

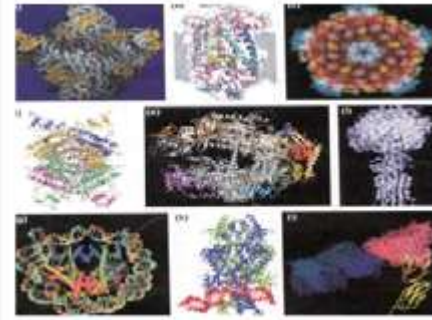
**common genetic  
(digital) code  
in all life forms**



**tool box of  
protein motifs  
for combinatorial  
assembly  
("molecular lego")**



**assembly of  
structurally and  
functionally  
diverse proteins**



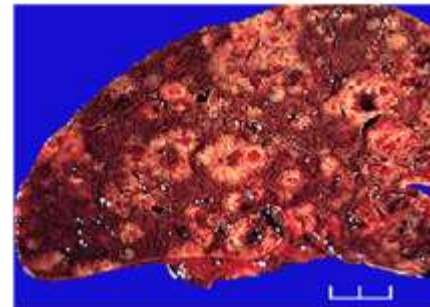
**networks of protein  
interactions and  
complex molecular  
signaling patterns**



**complex  
organism design**



**stable  
networks and  
information fidelity  
(health)**



**dysregulated  
networks and  
altered information  
patterns (disease)**

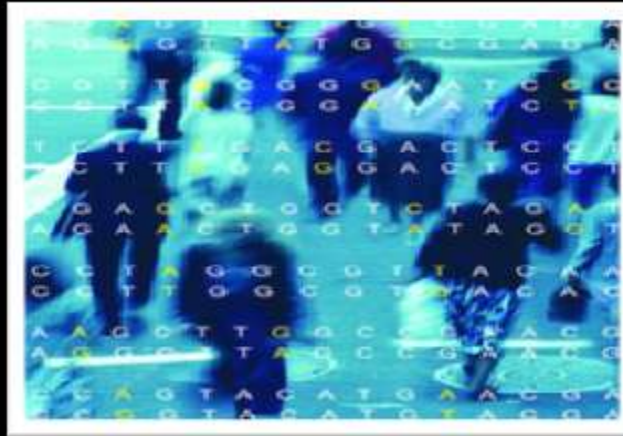


**therapeutic  
modulation  
of  
perturbed networks**

# Understanding the Rule Sets for Regulated Information Flow and Processing in Dynamic Biological Systems



**"It"**  
**(hardware/wetware)**



**"Bits"**  
**(software and  
encoded design)**



**"Exploring Biospace"**  
**(directed design of novel  
biological systems)**

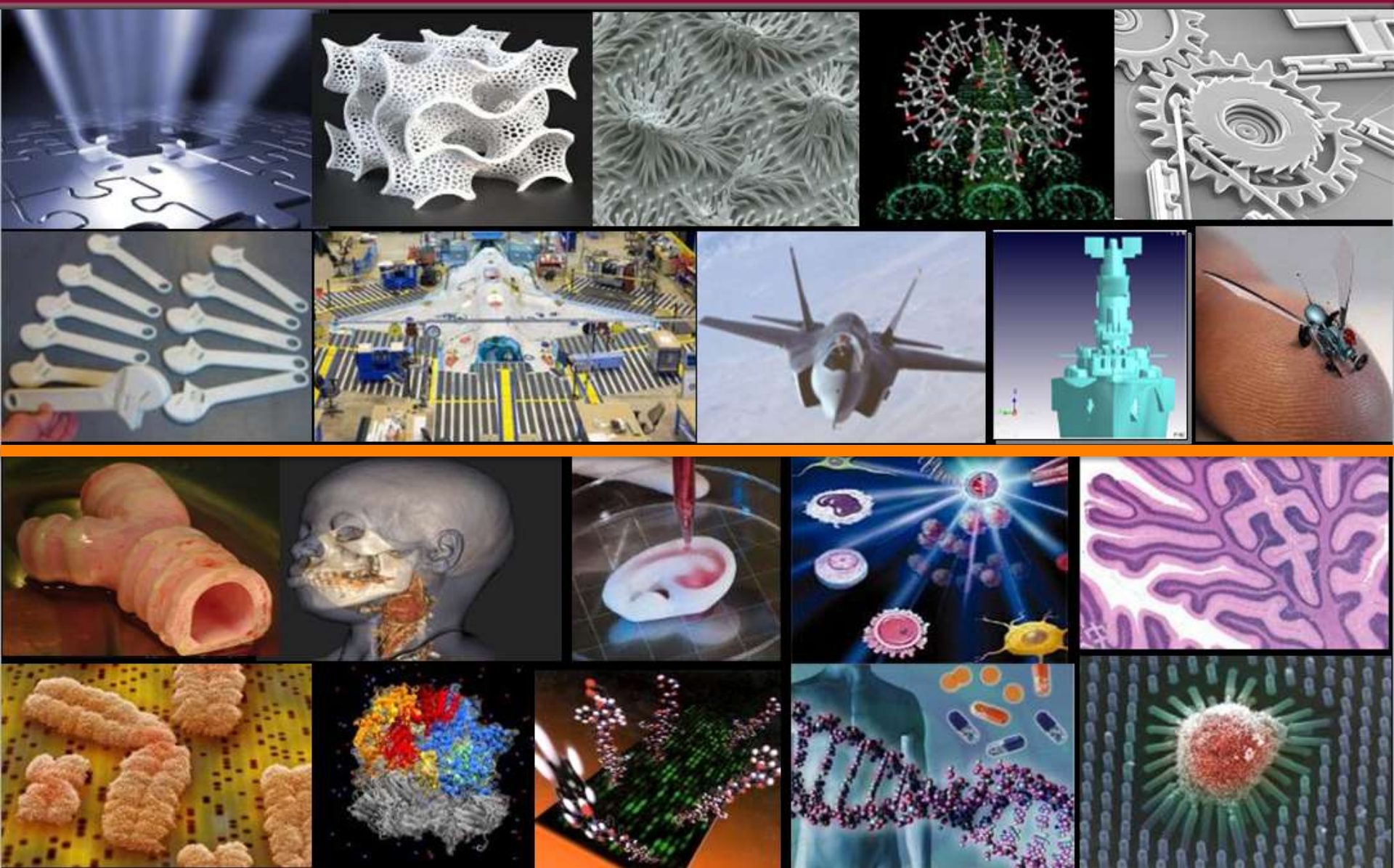


# **Digital Biology (Code) and Synthetic Biology (Construction): “It from Bits”**

- **program and assemble new biological functions and organisms based on knowledge of the ‘rule sets’ underlying hierarchical biological systems**
- **reprogramming existing biological signaling pathways and networks**
- **expanding the dimension of explored biospace**
  - **design, simulation and construction of novel functions/organisms with no known natural evolutionary counter part**
  - **novel biotic: abiotic combinations**
- **“directed evolution” and “accelerated evolution”**

# Advanced Manufacturing

## Digital Programming of New 3-D Fabrication and Assembly Technologies

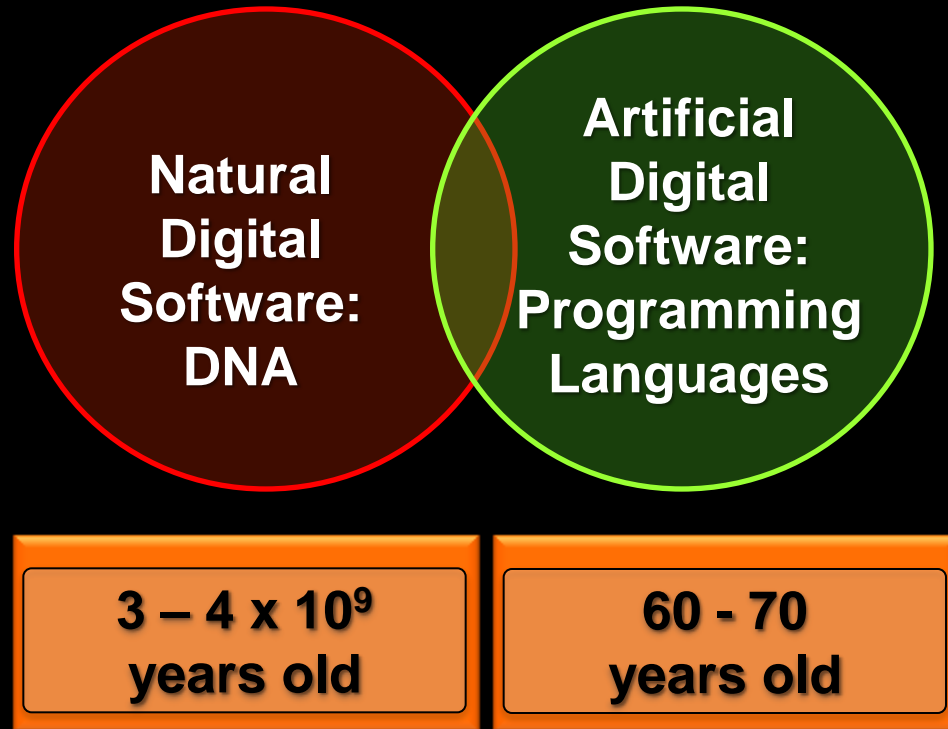




# **Programmable Matter: Computer-Controlled 3D Assembly of Structures of Increasing Complexity**

- **digital code for automated assembly of complex multi-scale structures**
- **uncoupling of design from fabrication and rise of point-of-need (PON) production capabilities**
- **3D printing**
  - **spatio-temporal assembly at nano-/ Ångstrom-level scale**
  - **abiotic materials**
  - **biotic materials**
  - **abiotic:biotic hybrids**
- **4D systems**
  - **self-assembly, repair and reconfiguration**
  - **dynamic adaptive behavior: repair, reconfiguration**

# Digital Convergence



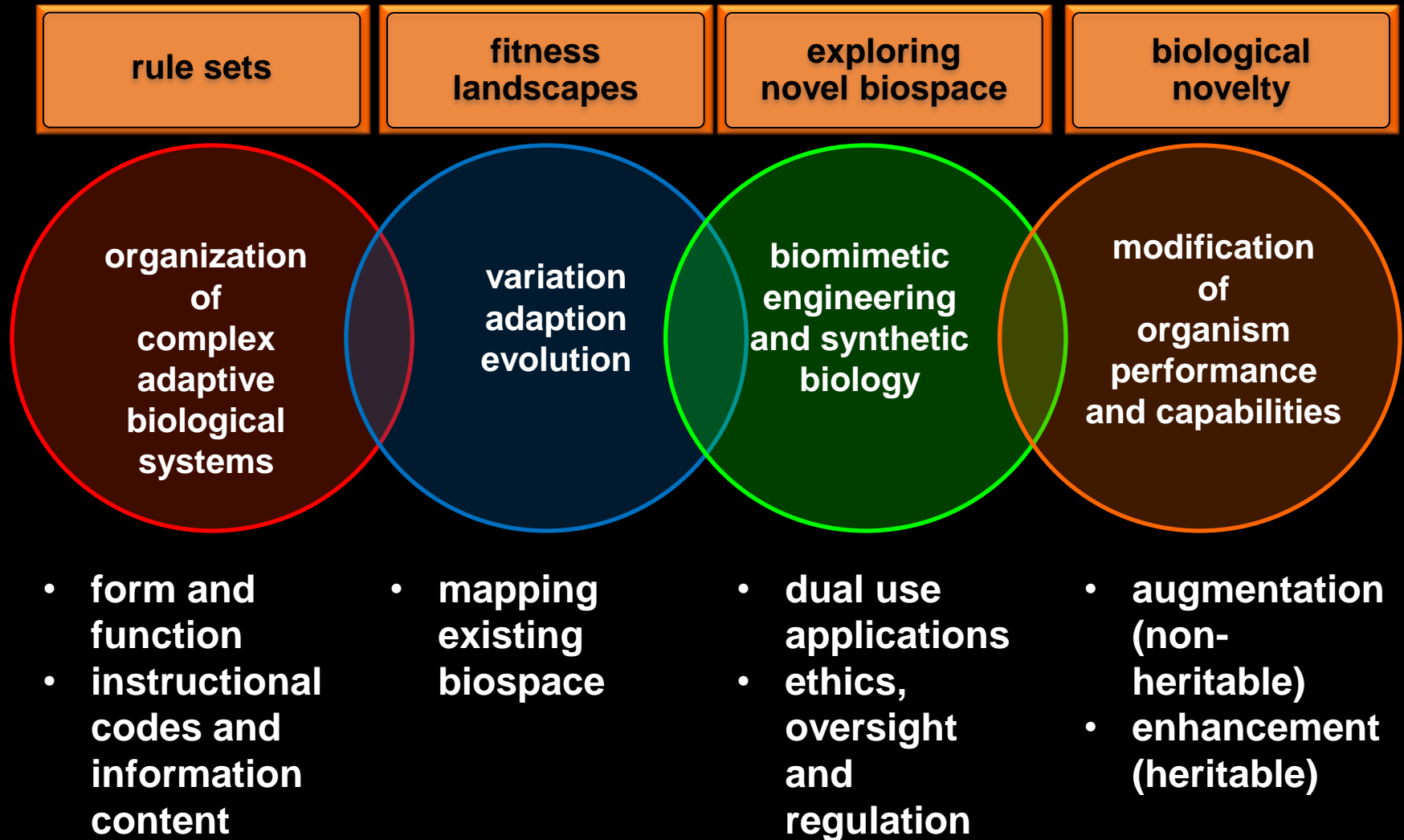


# Exploring Biospace

## The Power of Combinatorial Interactions and Molecular Assembly

- number of theoretical possibilities for synthetic assembly (biospace) far exceed narrow molecular space sampled in evolutionary time to date
- estimated 22,000 human genes
- two genes cooperate to create a function  
$$= (22,000 \times 21,999) / 2 = 241,989,000$$
 potential combinations
- 100 genes generate a complex function  
$$= 10^{65794}$$
 potential combinations

# Mapping the Coding Information for Organizational Complexity, Diversity and Variation in Biological Systems





# Technological Progress

**From Reactive Precarious Survival to  
Proactive Shaping of the Anthropocene**

**Technology Acceleration  
and Technology Convergence**

**Escalating Technical Complexity and  
Understanding Emergence in  
Complex Adaptive Systems**

# Complex Adaptive Systems: Increasing DDOF



- graded levels of autonomous behavior (components, system)
- escalating challenge of predicting system behavior and state shifts

# Understanding Complex Biological Systems and Directed Engineering of Biological Novelty



**“Oh, God help us!  
We’re in the hands of  
engineers.”**

**Dr. Ian Malcolm  
‘Chaotician’: Jurassic Park**