

EMERGENCY PREPAREDNESS – A MESSAGE FROM PURDUE

To report an emergency, **call 911**. To obtain updates regarding an ongoing emergency, sign up for Purdue Alert text messages, view www.purdue.edu/ea.

There are nearly 300 **Emergency Telephones** outdoors across campus and in parking garages that connect directly to the PUPD. If you feel threatened or need help, push the button and you will be connected immediately.

If we hear a **fire alarm** during class we will immediately suspend class, evacuate the building, and **proceed outdoors to the Grassy Area in front of SMITH HALL (Northeast)**. If inclement weather, **meet inside SMITH Hall**. Do not use the elevator.

If we are notified during class of a **Shelter in Place requirement for a tornado** warning, we will suspend class and **move to the Basement corridor**.

If we are notified during class of a **Shelter in Place requirement for a hazardous materials release, or a civil disturbance**, including a shooting or other use of weapons, we will suspend class and shelter in the classroom, shutting the door and turning off the lights.

Please review the Emergency Preparedness website for additional information.
http://www.purdue.edu/ehps/emergency_preparedness/index.html

ABE 591- Principles of Systems & Synthetic Biology

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Assistant Professor
Agricultural & Biological Engineering
Laboratory for Renewable
Resources Engineering

Fall 2018





**How do you engineer
a cell?**

DNA is the programming language of life

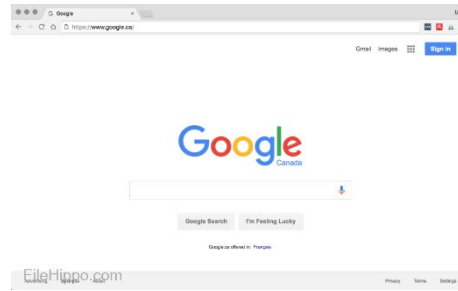
DNA → RNA → Protein → Function

Successes of recombinant technology:

- Insulin production
- Vaccine development
- Improved crop production

Paradigm shift: Biology as modular systems

DNA → RNA → Protein → Function



We can abstract or re-use modules with a specific functions

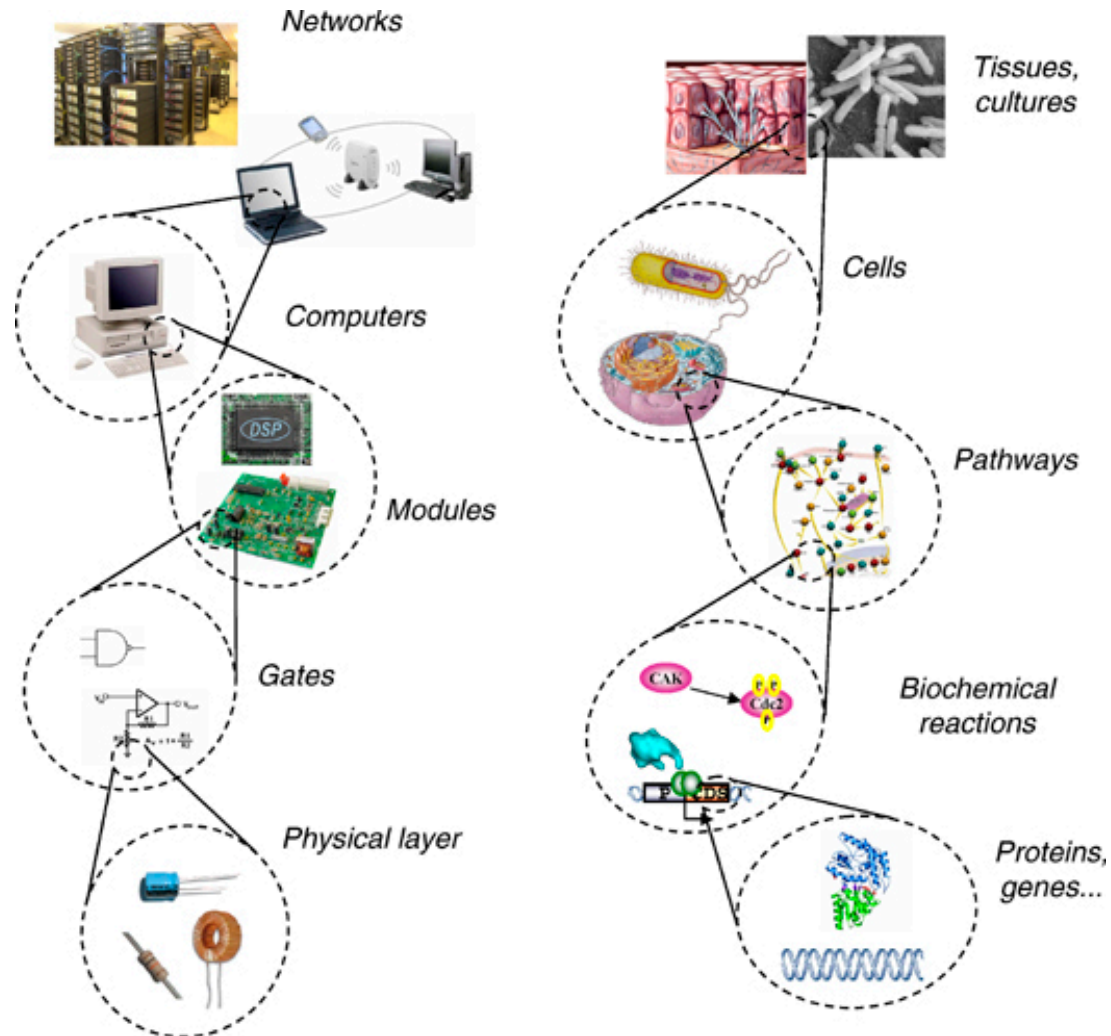
What is Synthetic Biology?

“engineering of living biological systems”

Goal: to design living systems using standard engineering practices to solve practical problems

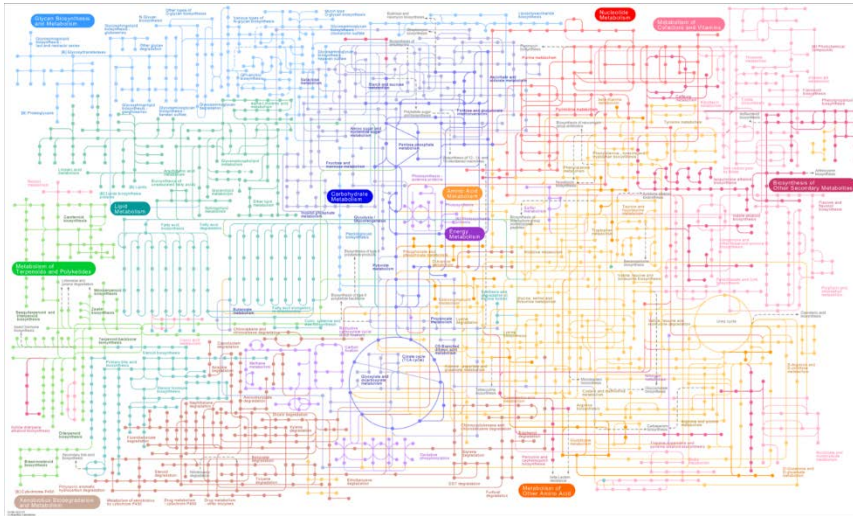


A possible hierarchy for synthetic biology is inspired by computer engineering.



Ernesto Andrianantoandro et al. *Mol Syst Biol*
2006;2:2006.0028

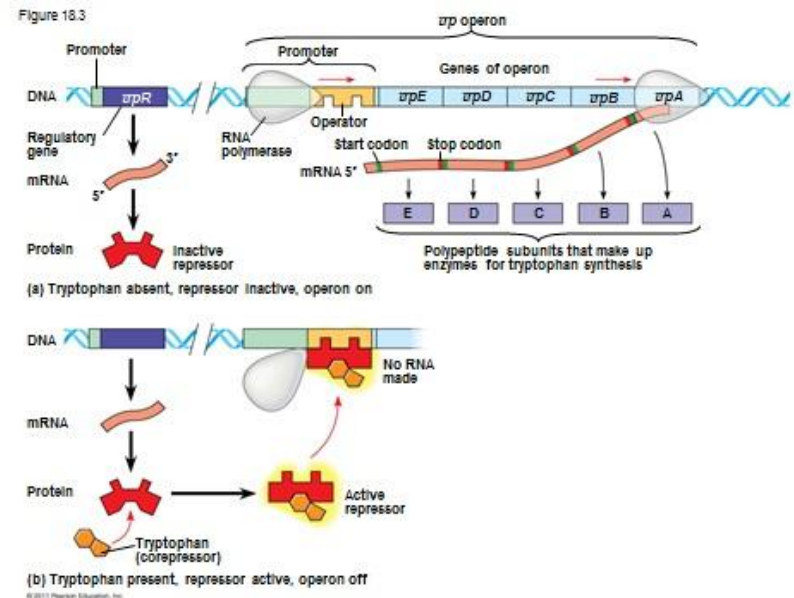
Why engineer biology?



KEGG

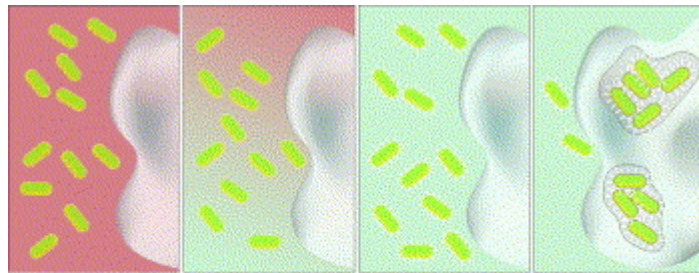
Living systems sense their environment and respond to stimuli

Cells are powerful chemical factories



Campbell Biology

(Some) Applications of synthetic biology



Aerobic Conditions
Low Cell Density
OFF

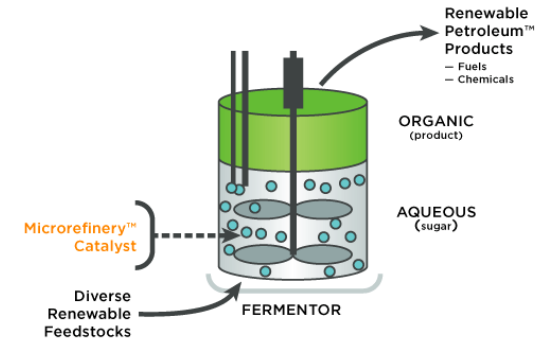
Hypoxia
High Cell Density
ON

→ *inv* Induction → Invasion

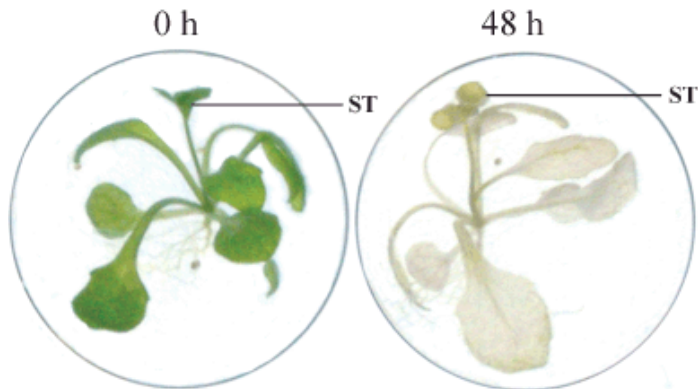
Cancer fighting bacteria



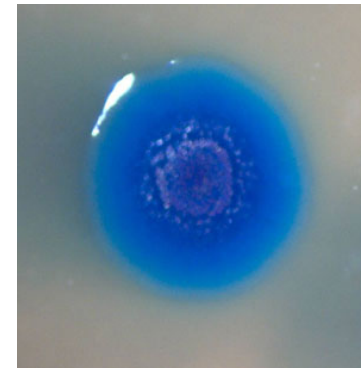
Bacterial photography



Nextgen biofuels



Bomb detecting plants



Study of biological systems

SynBio Industry



Global Companies: 290

Global Private Investment 2009-2015:
\$3.2B



Synbiobeta.com

 Dow AgroSciences

 Cargill®

 Agilent Technologies

 EVONIK
INDUSTRIES

 genomatica
sustainable chemicals

 TWIST
BIOSCIENCE

 AUTODESK

 GINKGO BIOWORKS™
THE ORGANISM COMPANY



OXITEC
GM mosquitoes to
control dengue fever



EVOLVA
High-value ingredients by
yeast fermentation



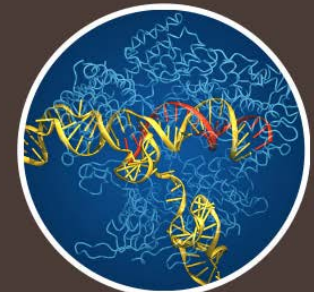
SYNTHACE
High-level programming
language for biology



INGENZA
Bioprocesses to manufacture chemicals,
pharmaceuticals and biofuels



ELIGO BIOSCIENCE
Microbiome precision
engineering

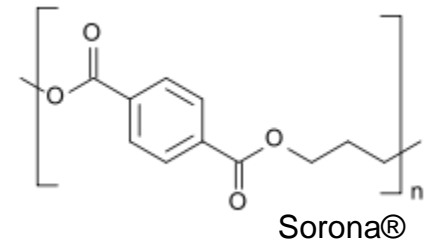


CRISPR Therapeutics
Gene editing technology
for medical therapies

Industrial applications: Sorona® & Hytrel® (Dupont)

Biopolymers from microbial 1,3-propanediol

- 100M pounds/yr
- \$100M plant in TN
- 25% of annual revenue



More sustainable

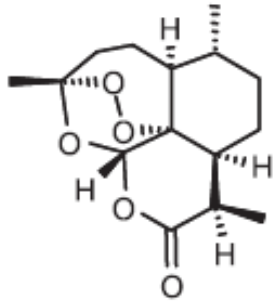
- 30% ↓ energy
- 63% ↓ greenhouse gas emission
- Renewable corn feedstock

More economical

- Less expensive than petroleum



Industrial applications: Artemisinin (Amyris)



Artemisinin

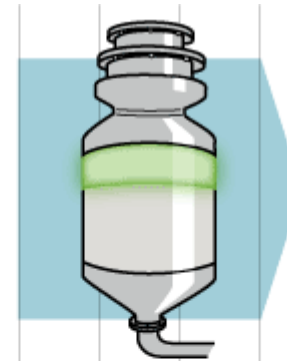
- Leading treatment of malaria
- 300 – 500 M malaria cases worldwide
- Difficult to make synthetically

Microbial fermentation

- Fast production
- Produced anywhere

Semisynthetic method
Fermentation & photochemistry

About 3 months from lab to ACT

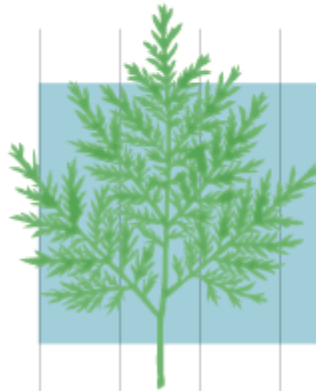


Traditional production via Chinese Sweet Wormwood plant

- Limited production
- Sensitive to climate
- Expensive

Agricultural method
Cultivation & extraction

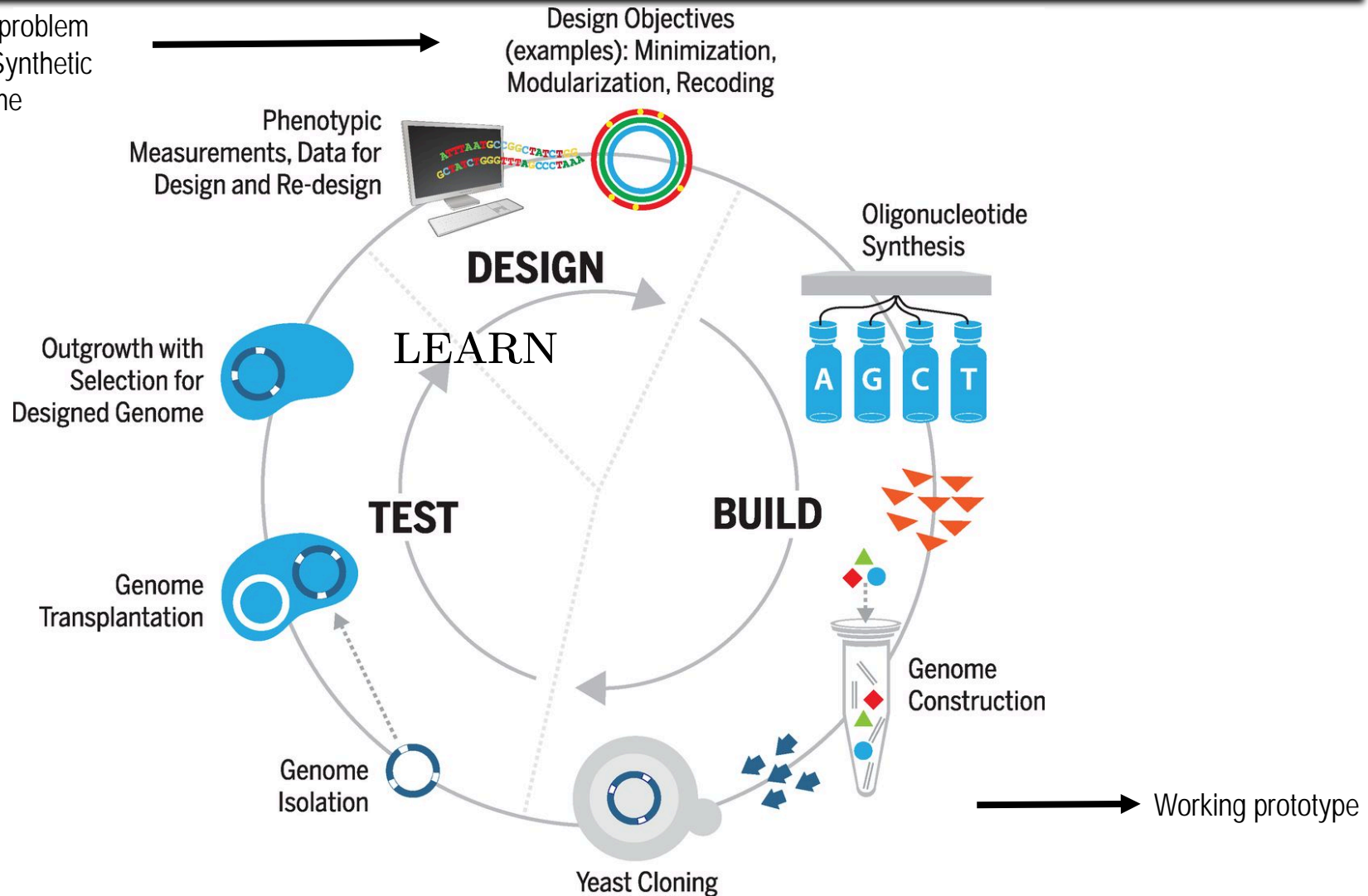
About 10 months from seed to ACT



**Sanofi-Pasteur &
Amyris** → 150M
treatments/a



SynBio follows an iterative Design-Build-Test-Learn Cycle



| | |
|-------------------|---|
| Week 1 | Engineering Biology Overview + Ethics |
| DESIGN | |
| Week 2 | Programming Biology |
| Week 3 | Properties of biological systems |
| Week 4 | Network Motifs |
| Week 5 | Nonlinear systems and emergent phenomena |
| Week 6 | BioCAD |
| BUILD | |
| Week 7 | Test 1 |
| Week 8 | DNA Assembly |
| Week 9 | DNA Assembly Cont'd |
| Week 10 | Genome Engineering |
| Week 11 | Genome Scale Assembly |
| Week 12 | Applications – Metabolic Engineering |
| Week 13 | SynBio Frontiers – Microbial communities and directed evolution |
| TEST/LEARN | |
| Week 14 | Test 2/Thanksgiving |
| Week 15 | Learning from Nature +-omics |
| | |
| Week 16 | Design Projects |

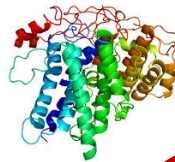
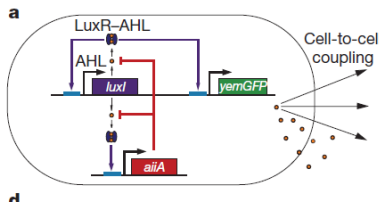
SYLLABUS

Course overview

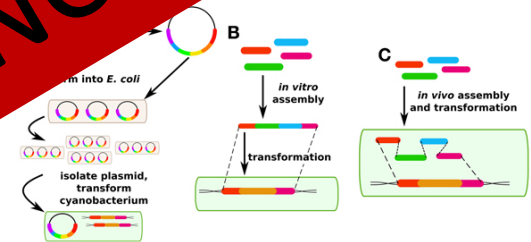
1. What are biological parts?

- How do we design/define new parts?

2. How do parts function together?



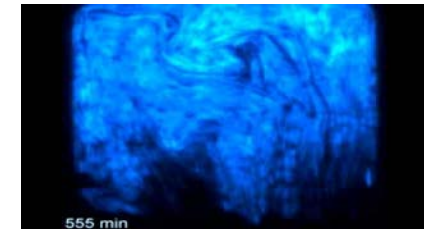
4. How do we synthesize these systems?



3. Describe the quantitative behavior of systems

5. Design & test systems for specific applications

$$\begin{aligned}\frac{\partial A}{\partial t} &= \gamma_A A - \frac{\gamma_A A}{1 + f(A + I)} \\ \frac{\partial I}{\partial t} &= \gamma_I I - \frac{\gamma_I I}{1 + f(A + I)} \\ \frac{\partial H_i}{\partial t} &= \frac{\gamma_I}{1 + kI} - \frac{\gamma_H A H_i}{1 + gA} + D(H_e - H_i) \\ \frac{\partial H_e}{\partial t} &= -\frac{d}{1-d} D(H_e - H_i) - \mu H_e + D_1 \frac{\partial^2 H_e}{\partial x^2}\end{aligned}$$



What should we engineer?

Best practices are to minimize risk and weigh potential good vs harm

Multiple perspectives:



**Scientists/
Experts**



Investors



Government



Public

Technical Risks & Ethical/Legal/Social Concerns

1. Biosafety

- Toxins? Ecosystem damage? Biological harm

2. Biosecurity

- Dual-use? Increased lethality? Proliferation?

3. Justice and fairness

- Who benefits?

4. Socioeconomic

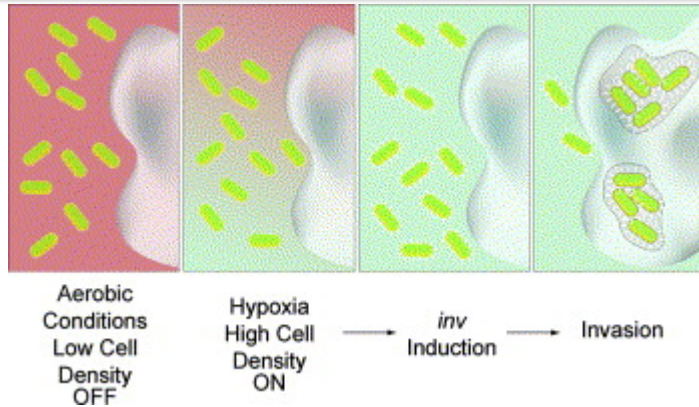
- IP? How disruptive is tech?

5. Eugenics/Playing God

- Should we be tampering with nature?

Which risks are most important to the different stakeholders?

Example



Cancer fighting bacteria

- To hunt down tumors, need to evade immune system → potential for “dual use”
- Will treatment be accessible?
- Safe? Accurate?

Scientists need:

- to be aware of risks (technical or not)
- address risks where possible in design
 - e.g. ‘kill’ switches to prevent survival in wild (nutrient auxotrophies, non standard amino acids)
- Engage public to effectively communicate risk