

# LECTURE 1: Synthetic Biology

Introduction to cellular system modelling  
Daniel Georgiev

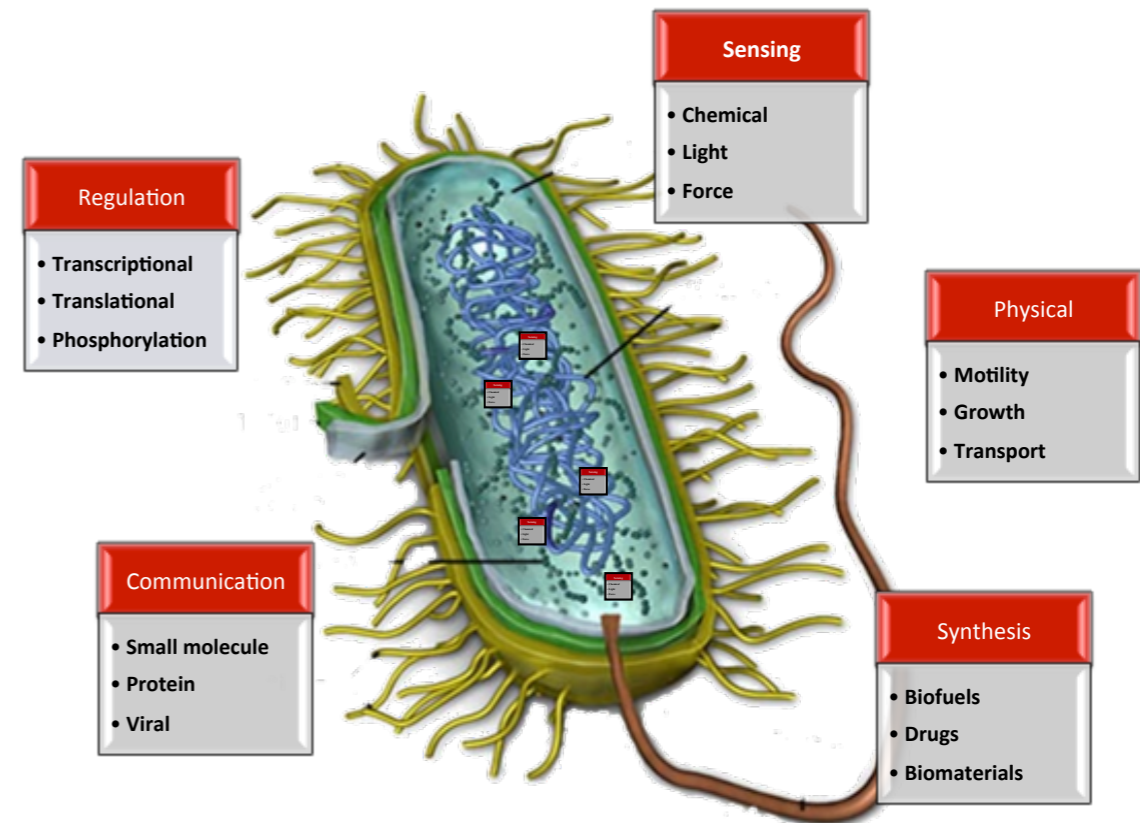
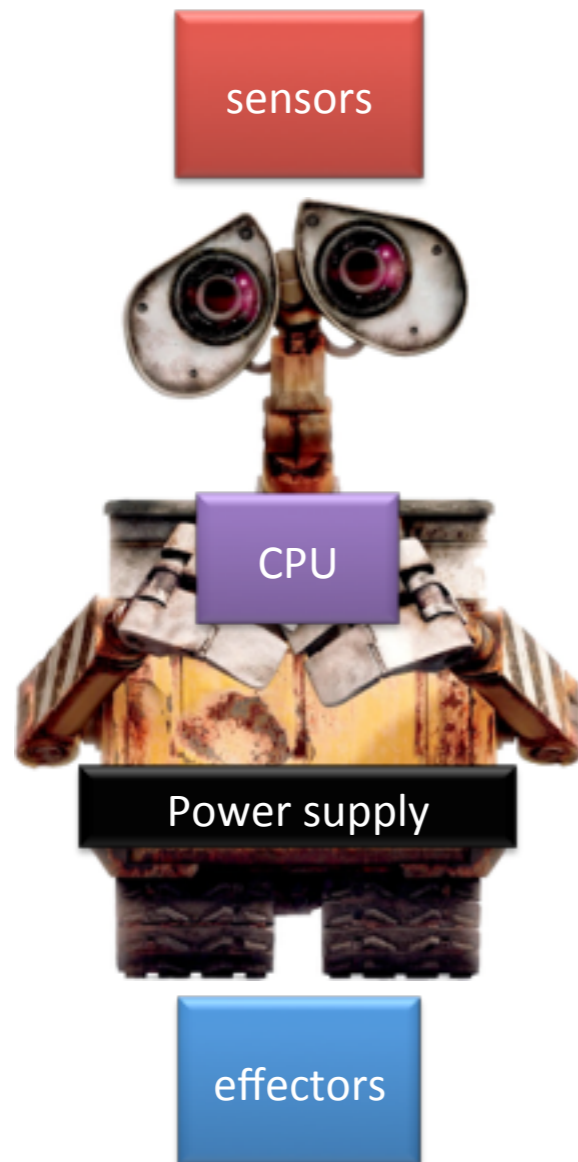
Summer 2015

# OUTLINE

- Robot cell
  - source
  - cpu
  - actuators
  - sensors
- Application
  - production
  - modern medicine
  - environment
- Basics
  - DNA
  - central dogma
  - metabolism
  - signalling
- Examples of simple synthetic devices
- Complex system design

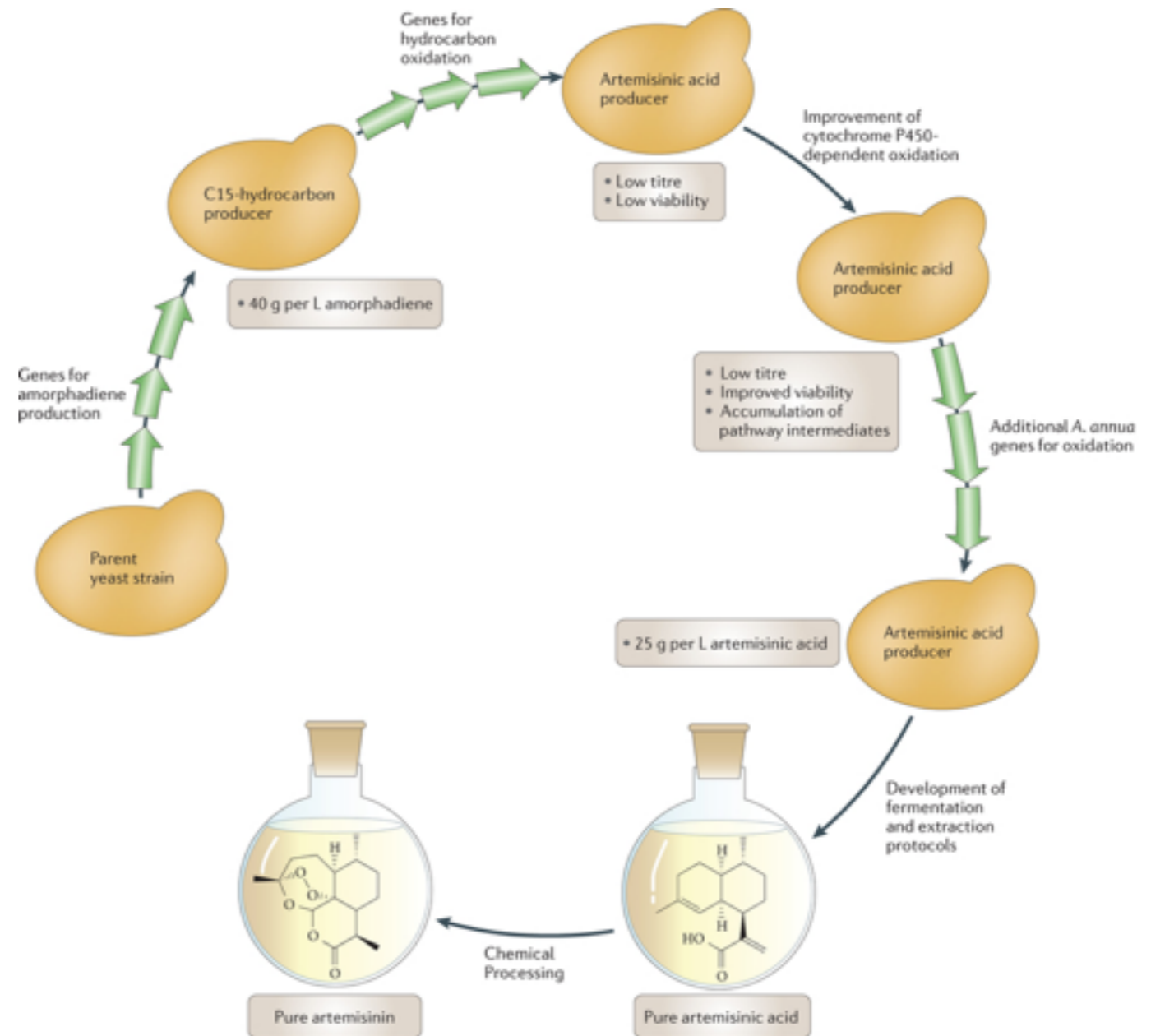
# SYNTHETIC BIOLOGY

building up from parts to complex systems



# SYNTHETIC BIOLOGY

applications - bioproduction



## MALARIA

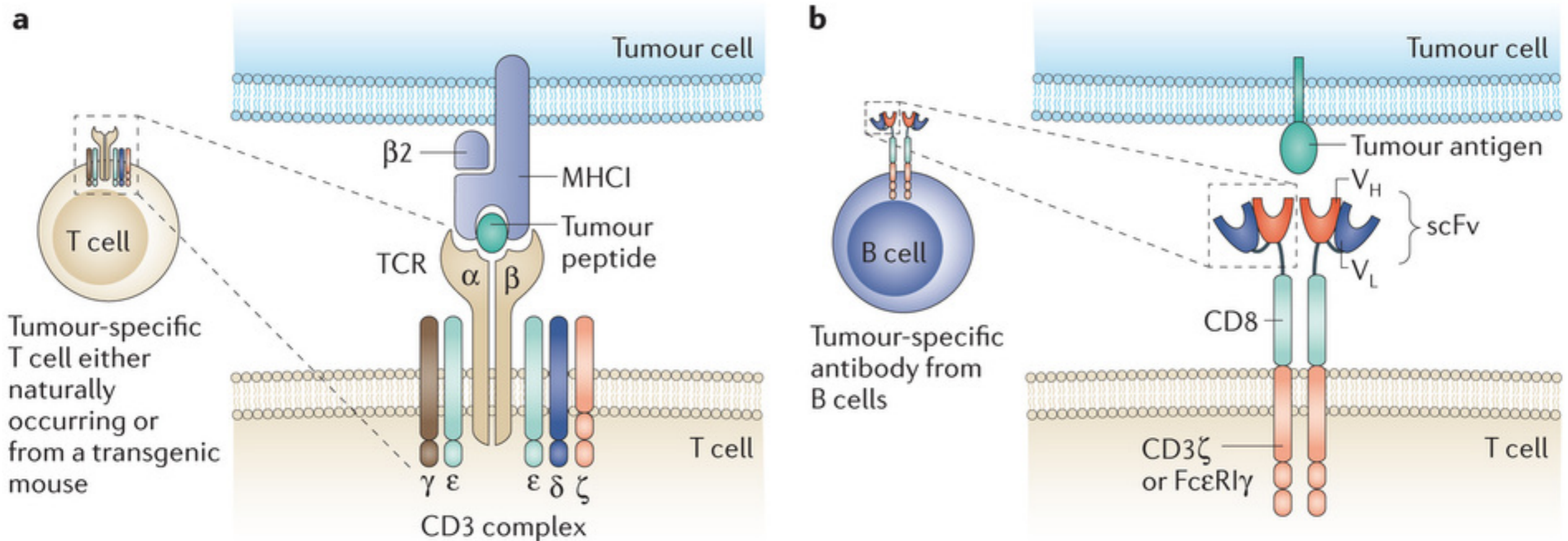
12 billion USD

198 million cases per year

drug artemisinin = \$1100/kg

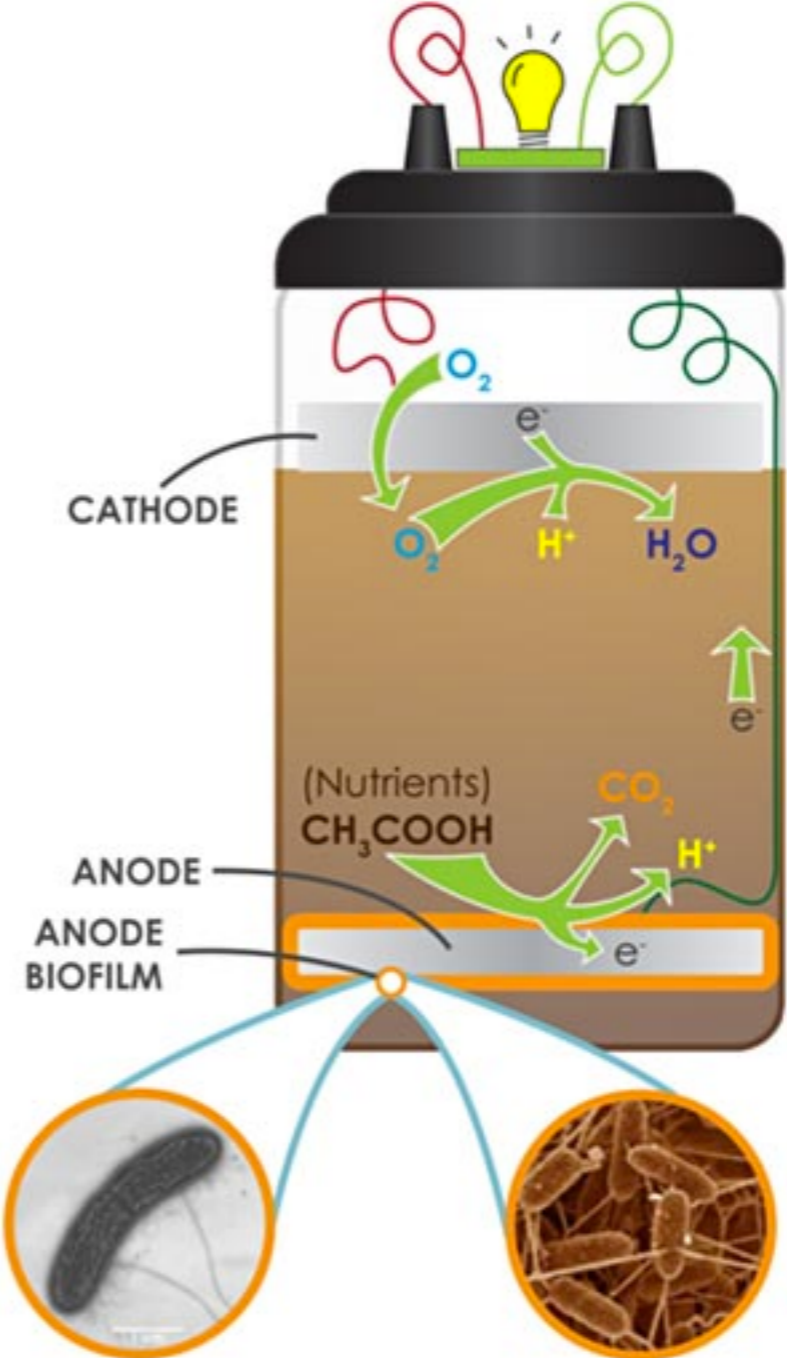
# SYNTHETIC BIOLOGY

applications - medicine



# SYNTHETIC BIOLOGY

applications - ecology



# MOLECULAR BIOLOGY

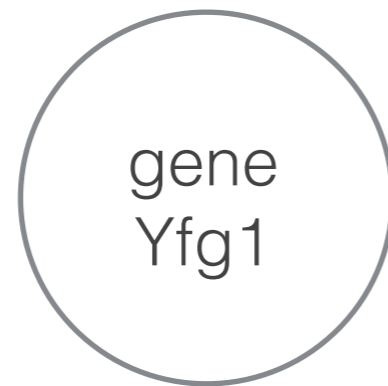
basics - DNA





# MOLECULAR BIOLOGY

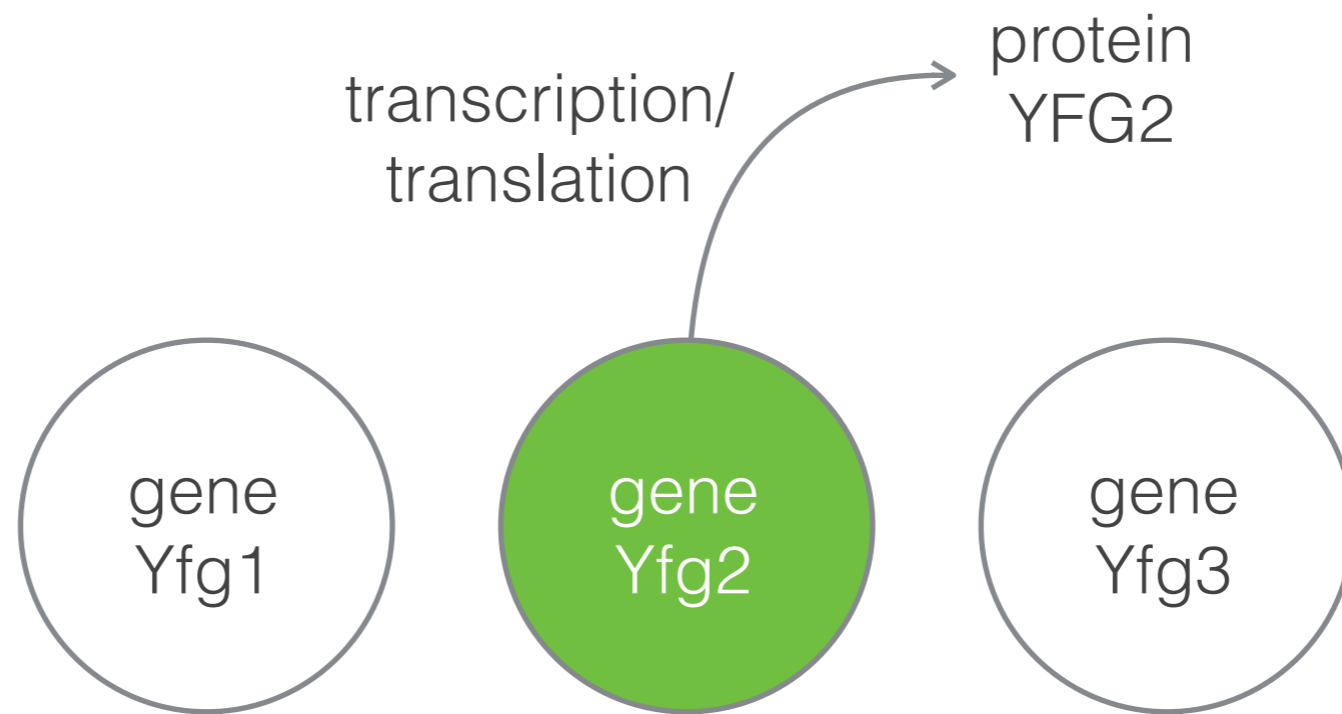
basics - DNA





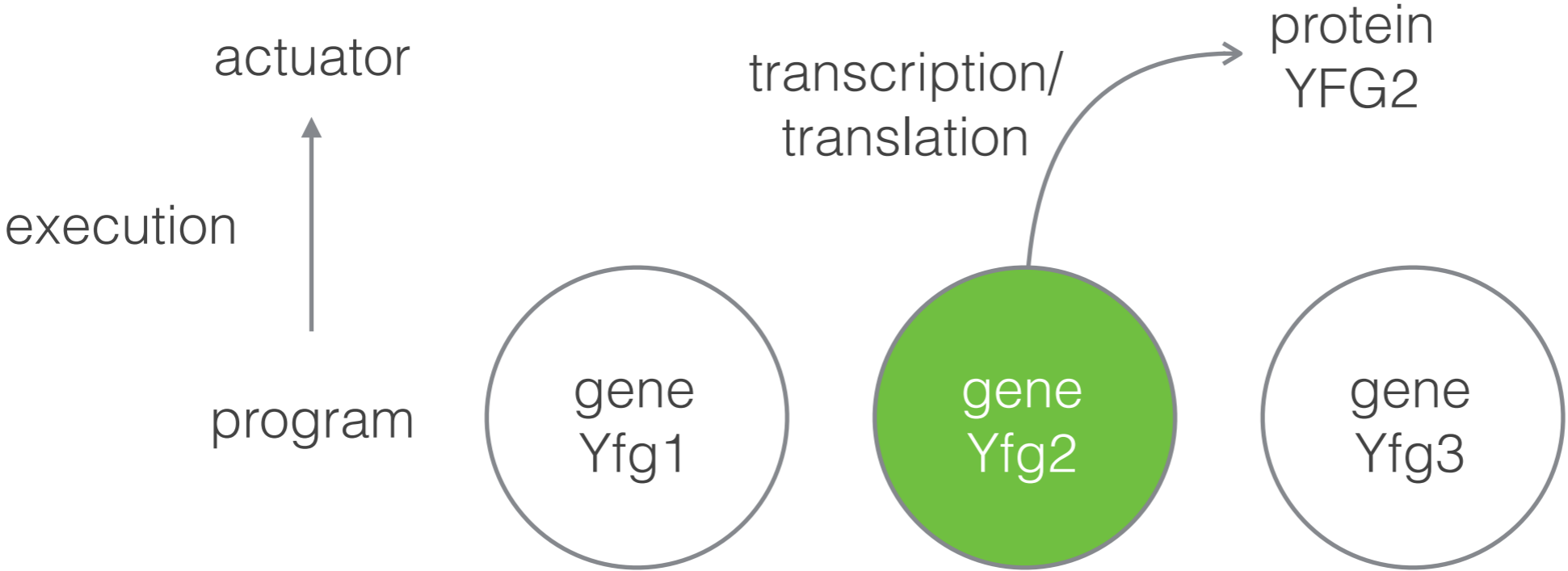
# MOLECULAR BIOLOGY

basics - DNA



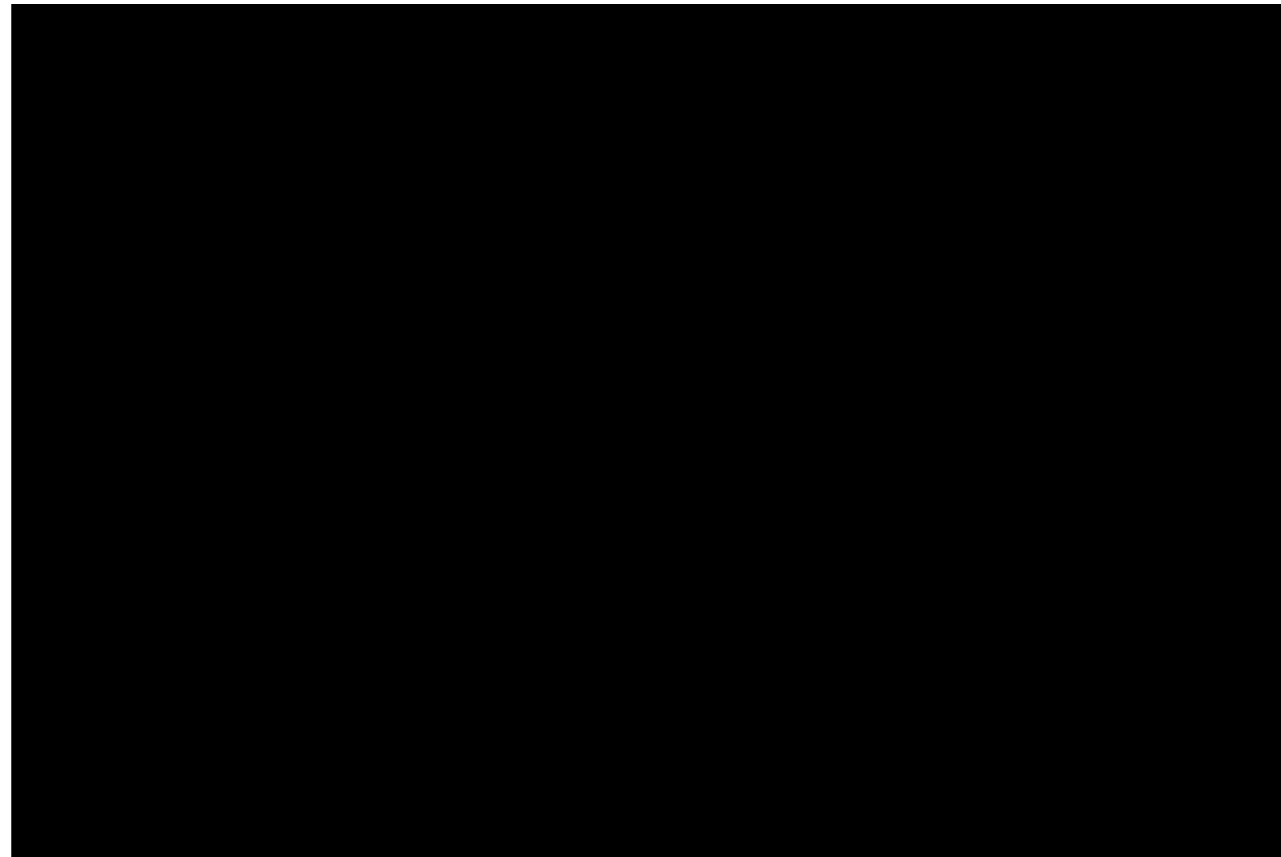
# MOLECULAR BIOLOGY

basics - DNA



# MOLECULAR BIOLOGY

basics - DNA



# MOLECULAR BIOLOGY

## basics - DNA

Engineering the Central Dogma of Molecular Biology  
 Daniel Georgiev, Faculty of Applied Science, UWB, Pilsen, 2011

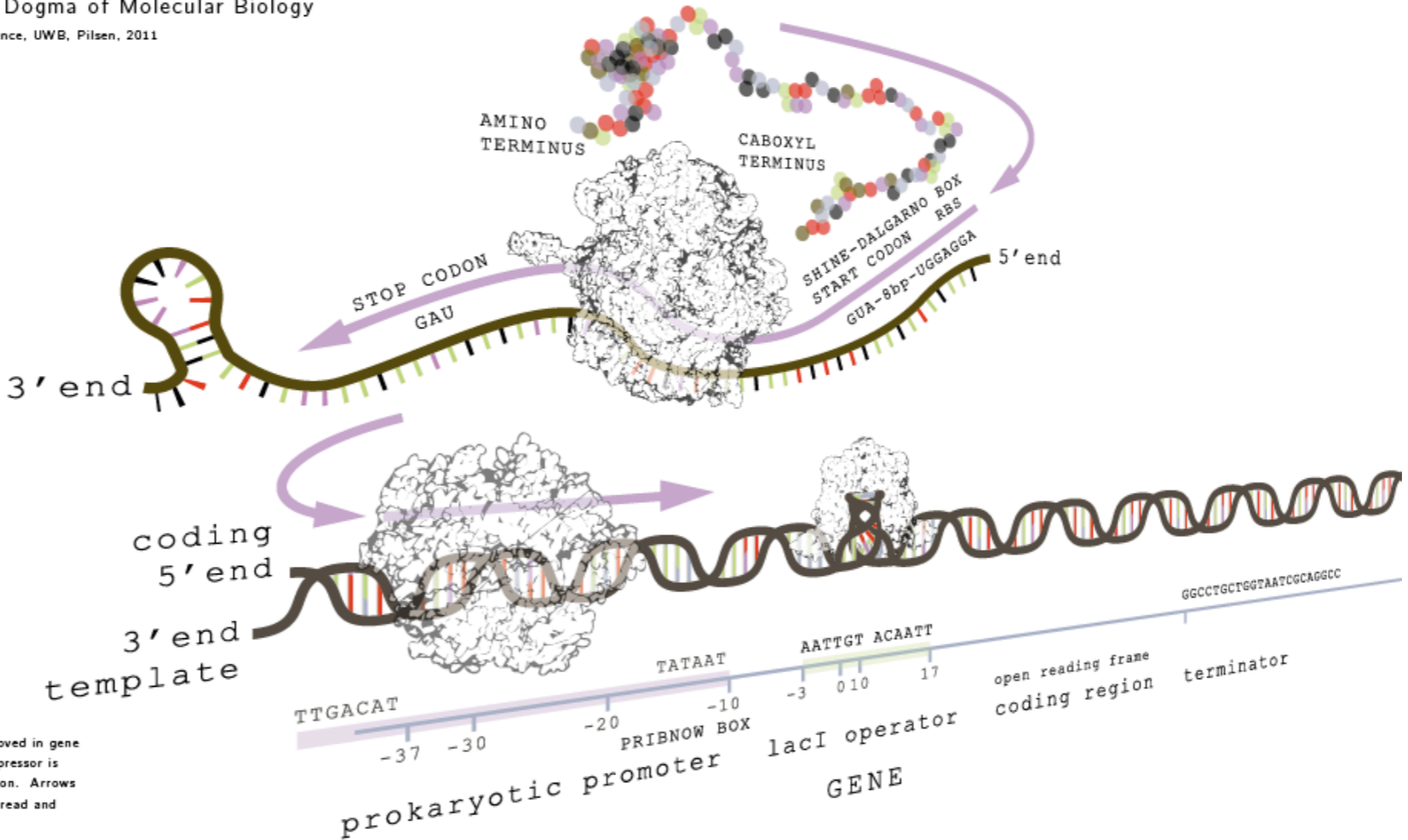


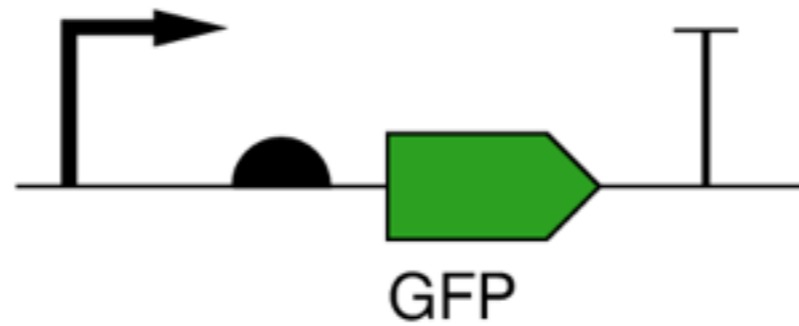
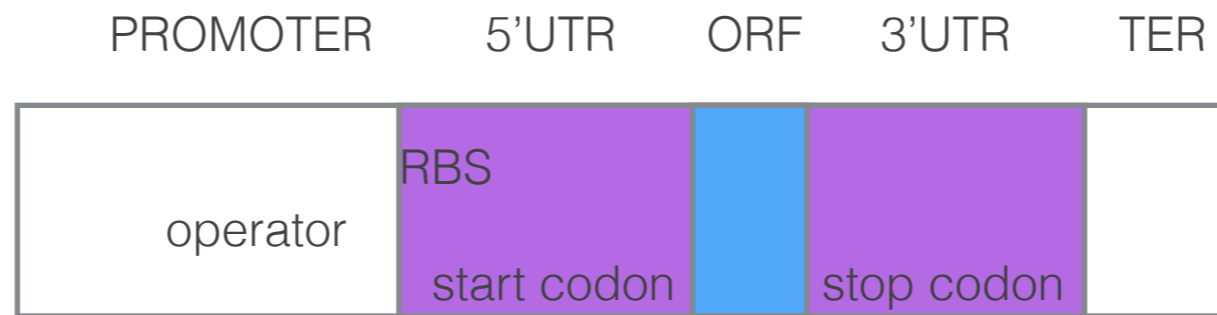
Illustration of the key components involved in gene expression and regulation. The lacI repressor is used to illustrate transcription regulation. Arrows indicate the direction in which DNA is read and RNA/proteins are built.



# MOLECULAR BIOLOGY

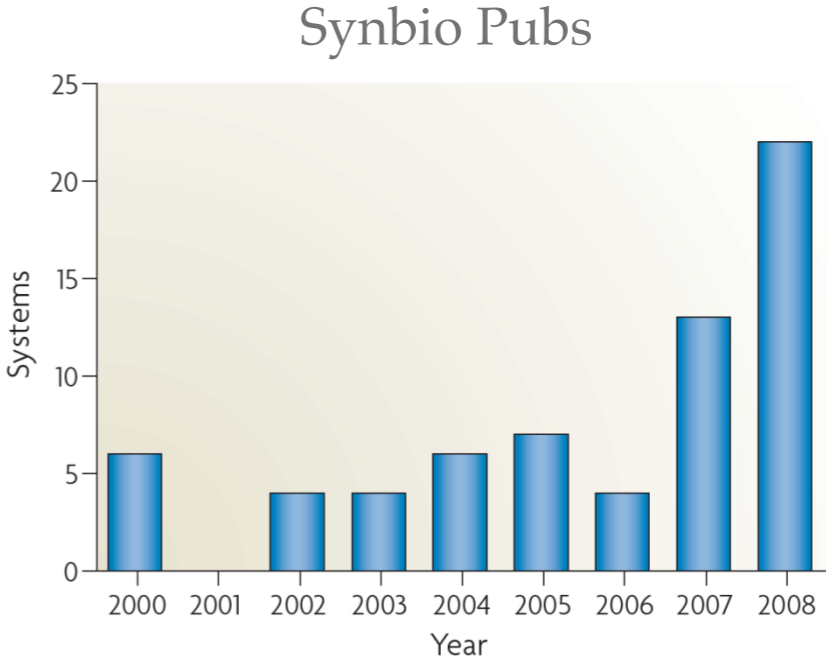
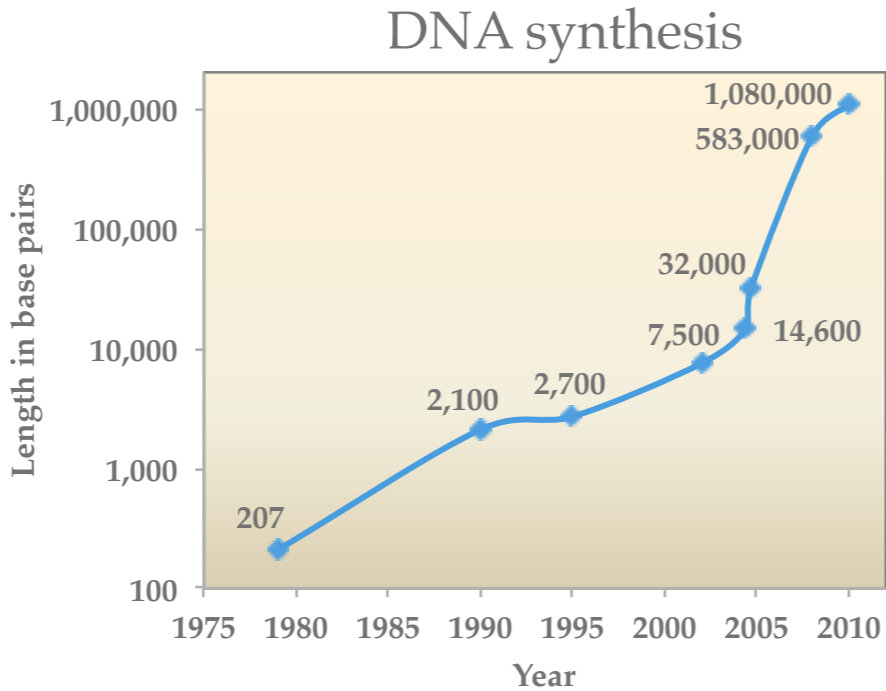
## basics - DNA

1 atggtgaatg tgaaccagt aacgttatac gatgtcgcag agtatgccgg tgtctttat  
61 cagaccgttt cccgcgtggt gaaccaggcc agccacgttt ctgcgaaaac gcgggaaaaa ...



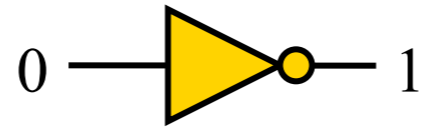
# MOLECULAR BIOLOGY

## basics - DNA



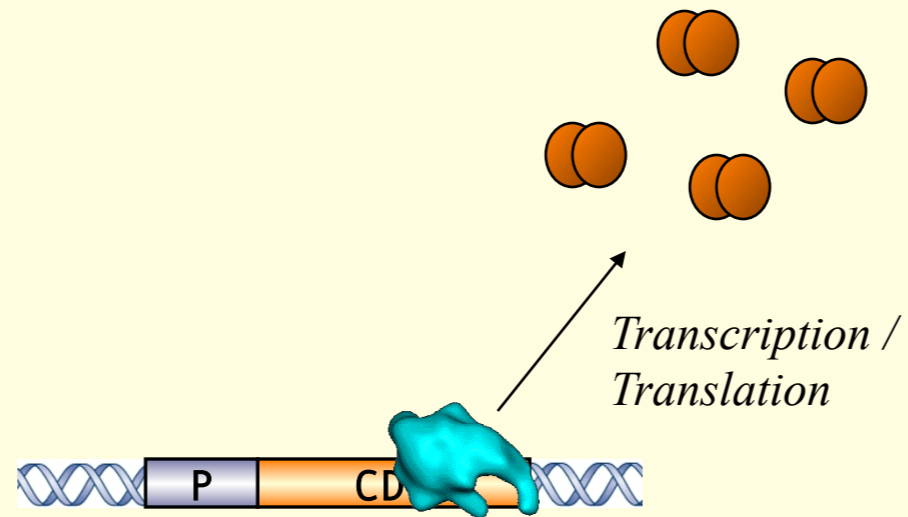
# SIMPLE PARTS

inverter



input protein  
(repressor)

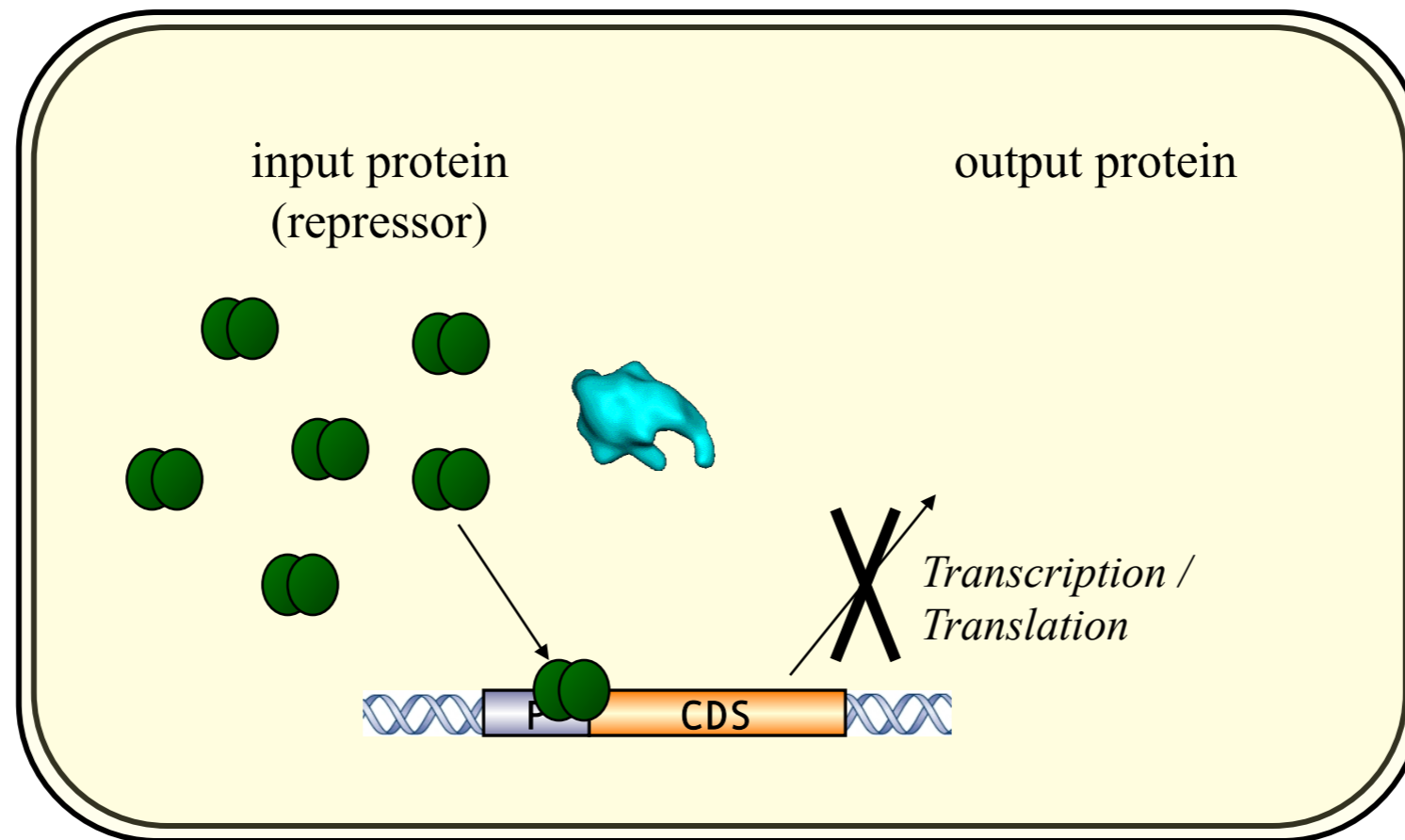
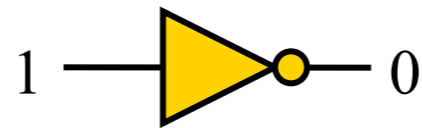
output protein





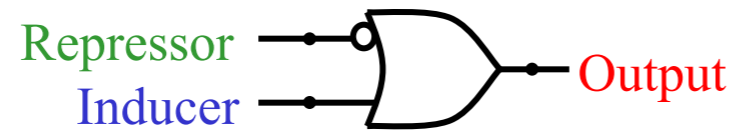
# SIMPLE PARTS

inverter

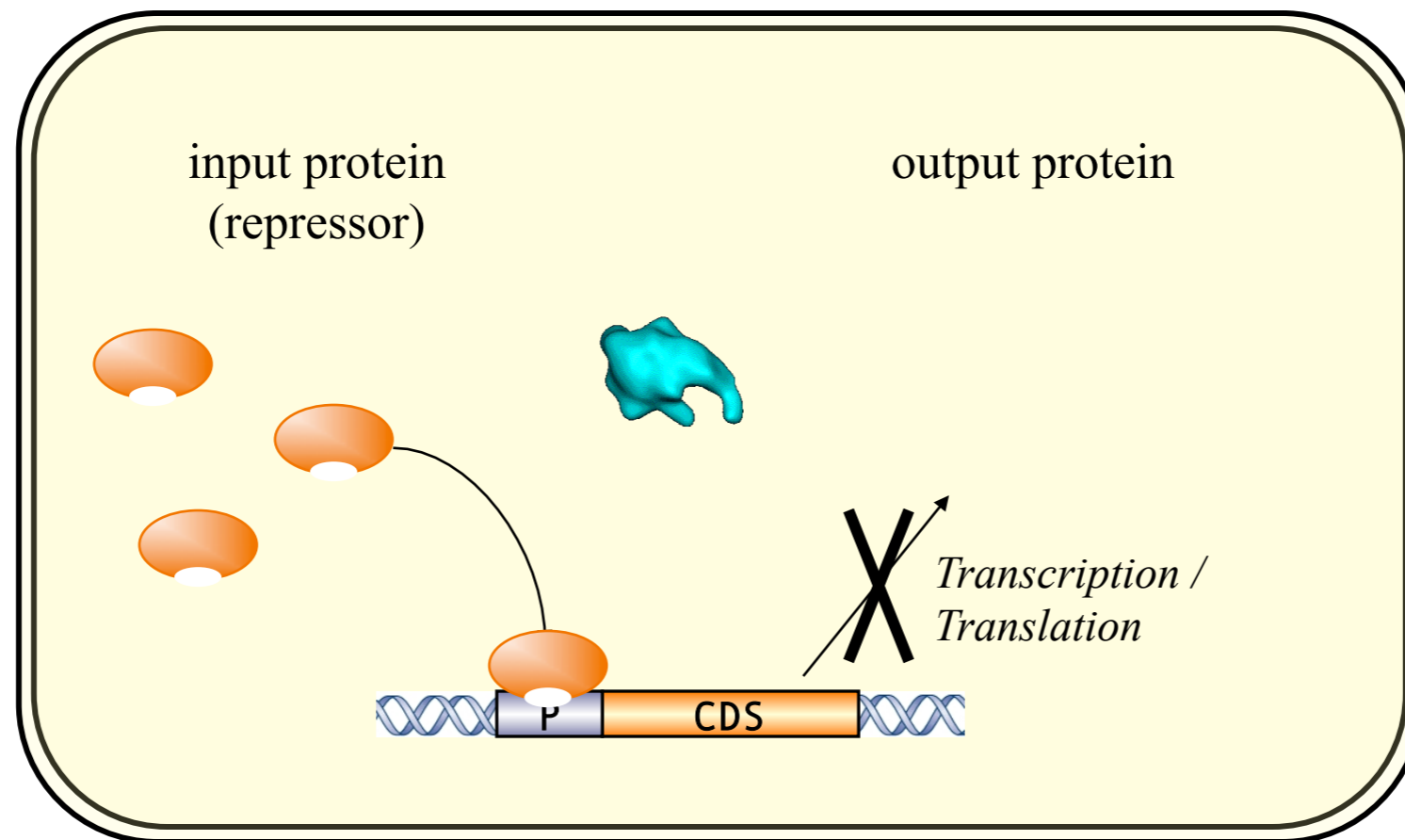


# SIMPLE PARTS

implies gate

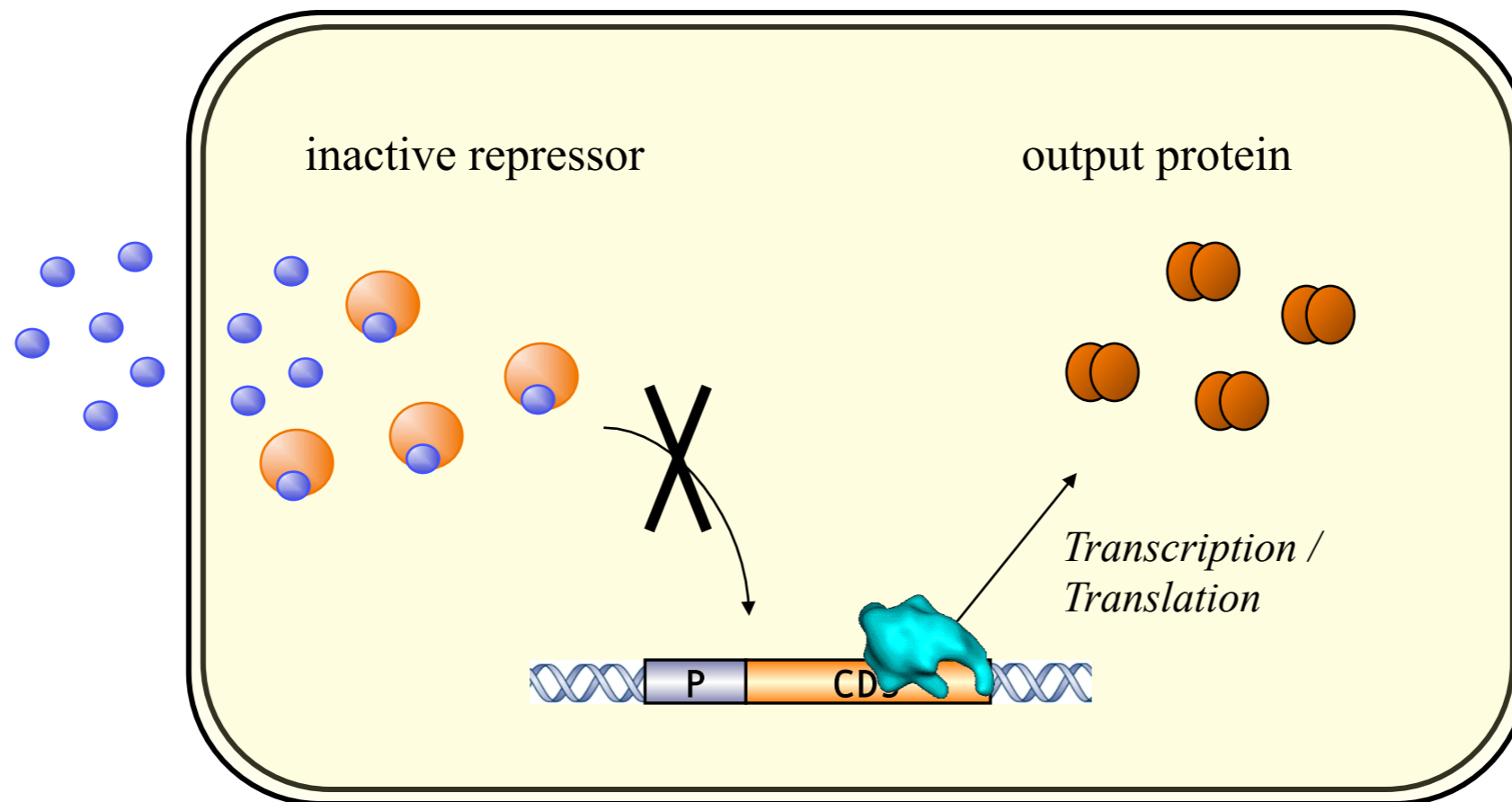
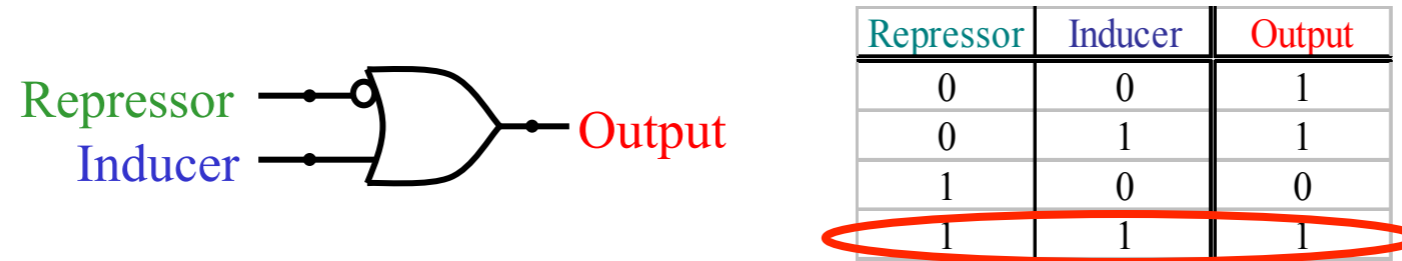


Repressor	Inducer	Output
0	0	1
0	1	1
1	0	0
1	1	1



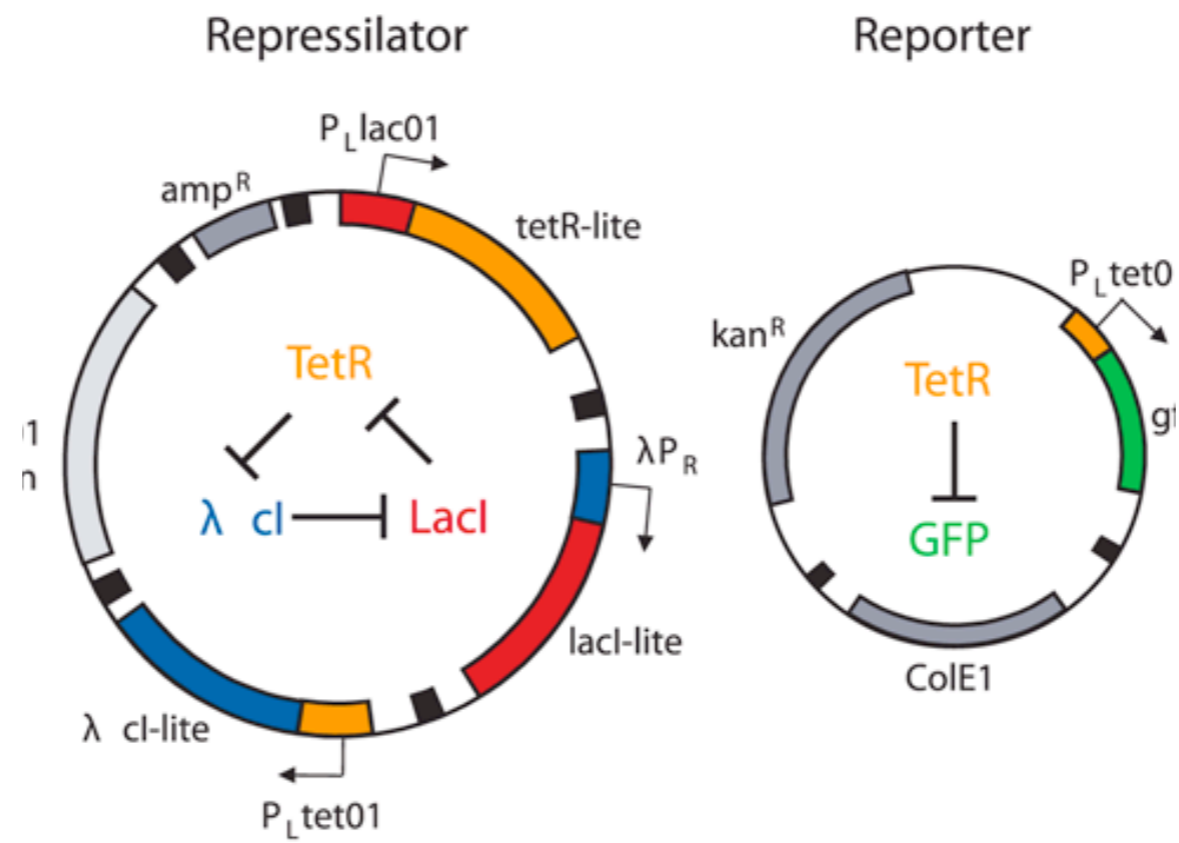
# SIMPLE PARTS

implies gate



# SIMPLE PARTS

oscillator



# SIMPLE PARTS

oscillator

VALIDATION OF TECHNOLOGY

proof of concept plus quantitative characterization

48 min

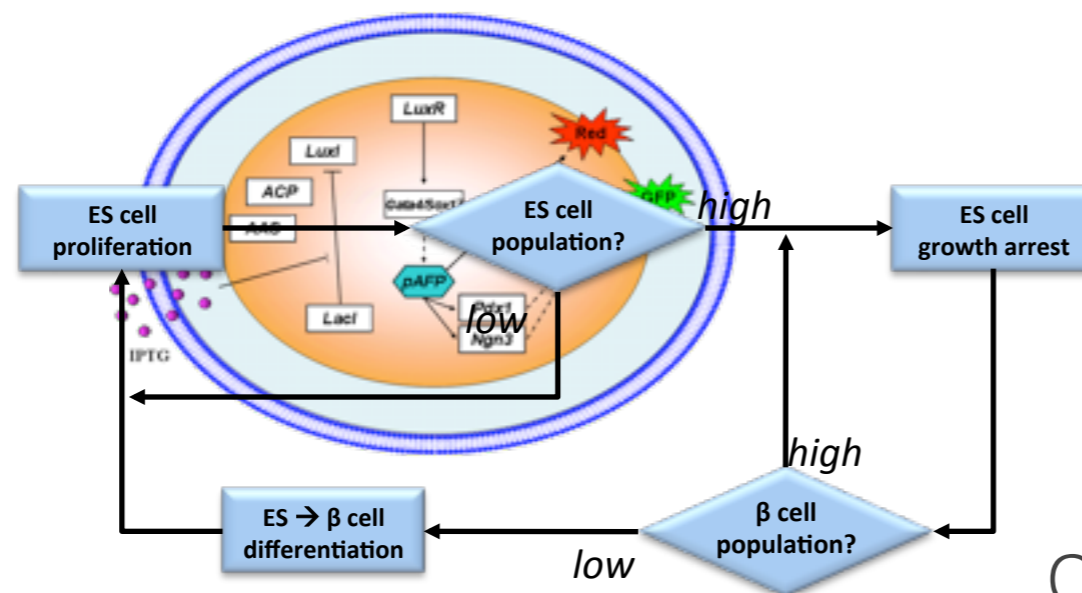
# COMPLEX DESIGN

beta cell homeostasis with regulated ESC differentiation

7.8% of the US population has diabetes  
In Diabetes Type I (10% of diabetics),  
auto-immune response (slowly)  
kills insulin-producing pancreatic  $\beta$  cells.

Goal:

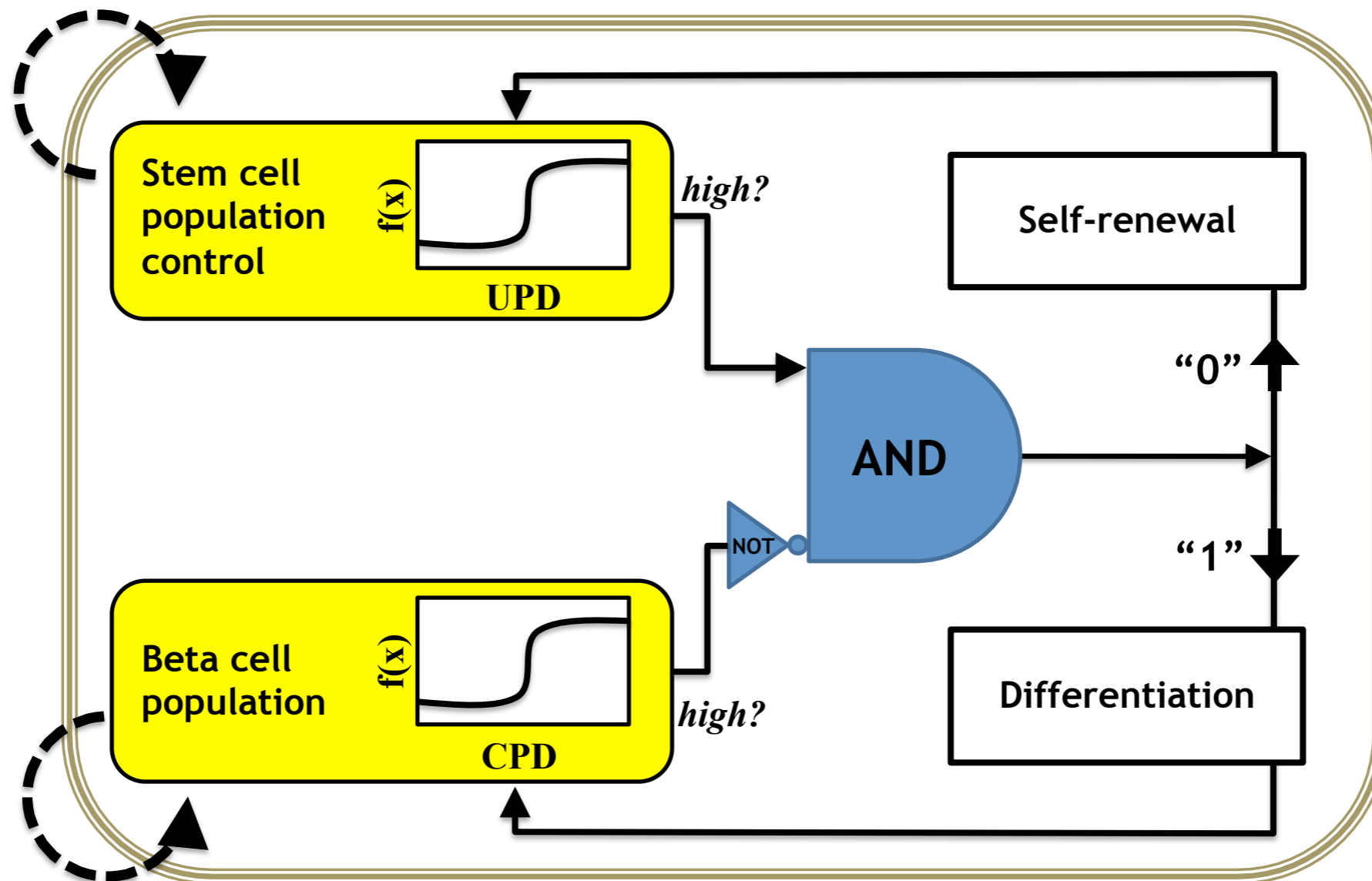
Maintain population level of  $\beta$  cells using  
auto-regulated differentiation of ES cells  
that counter-balances auto-immune attacks.



Complex system with 22 components  
Design with 'known' modules!

# COMPLEX DESIGN

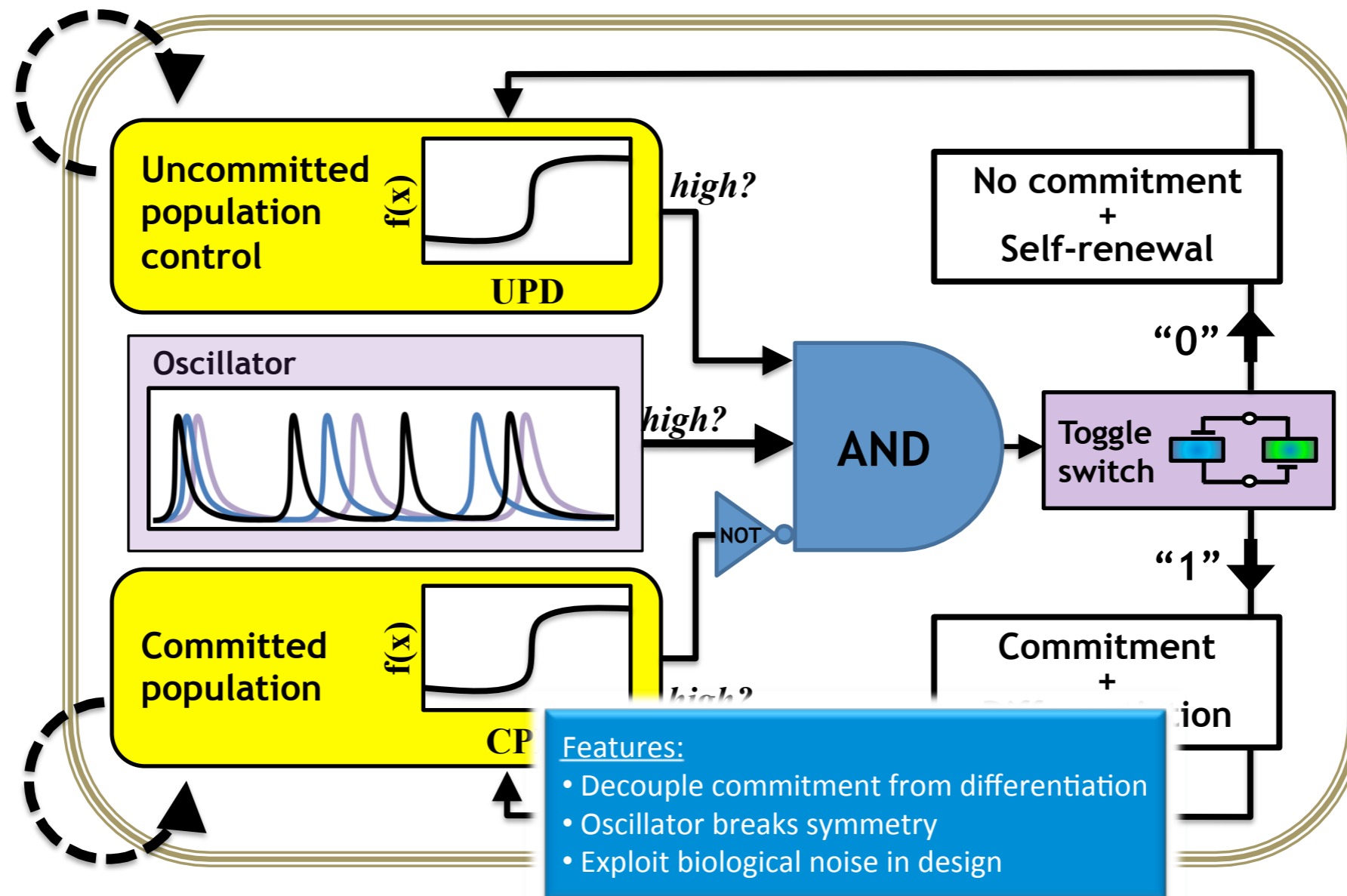
beta cell homeostasis with regulated ESC differentiation

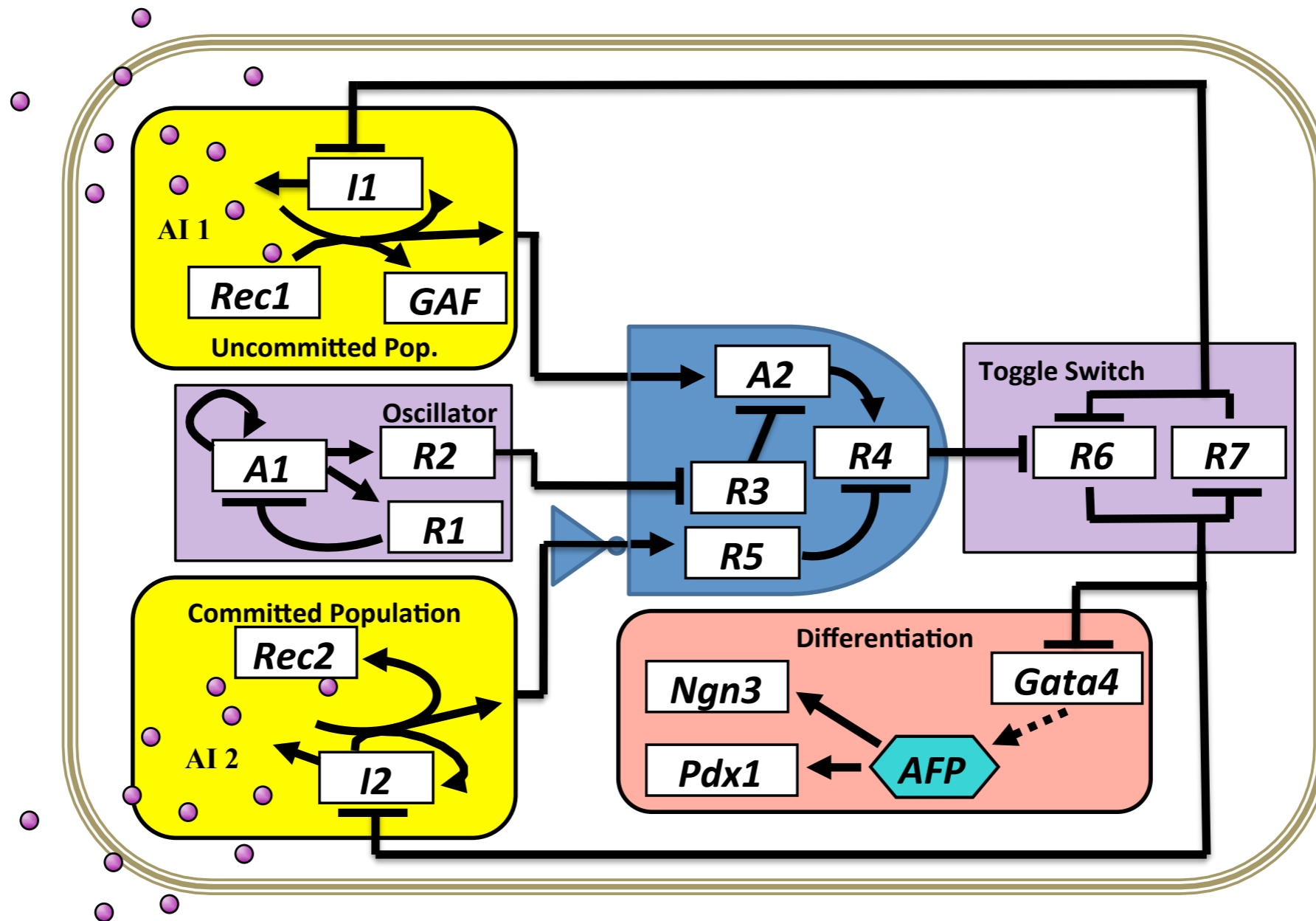




# COMPLEX DESIGN

beta cell homeostasis with regulated ESC differentiation





# THE BIG PROBLEM

COMPLEX DESIGNS ARE NOT FEASIBLE WITH USING CURRENT ENGINEERING PRINCIPLES

# SIMPLE PARTS

design and engineering approach is not scalable

