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Synthetic Biology

Engineering a Novel Organism

- Living things are constructed by proteins scripted out by the genes in our DNA.
- A living organism can have anywhere from 50-100,000 genes
- · Numerous genomic maps have been constructed
- · However this is just simple code which forms really complex interactions
- We are trying create a simpler system of 12 genes

Designing the Circuit

- The circuit is a cyclic three gene oscillating system
- Each gene turns the following one off
- On/off oscillation
- This creates polysystronic eukaryotic repressillator system

C21-Steroid hormone

Results will be visually displayed with color

Constructing the Circuit



Dehydroepiandro-

Androst-4-ene-

3,17-dione

5α-Androstane-

3.17-dione

Dehydro-

epiandroste

7α-Hydn

androsten

19-Hydrox

4-ene-3,17

19-0 xo-an

4-ene-3,17

17α-Hydroxy-

21-Steroid hormone

3-0xo-13,17-

17.13*6*: -lactone

3.17-dione

1.1.1.50 1.1.1.152

e tabolism

randrost-

- The system was designed from basic genes, and it is without a natural counterpart
- The genes used for the circuit will come from several different purchased plasmids (circular pieces of DNA)
- The specific genes that are needed from each purchased plasmid will be individually amplified using PCR
- These will then be coupled together using fusion PCR

Sulfate Glucuronide

The 3 main pieces are the Lac gene, the Cumate gene, and the Tet gene

16-Glucuronide C

Sulfate Glucuronide



Math Modeling

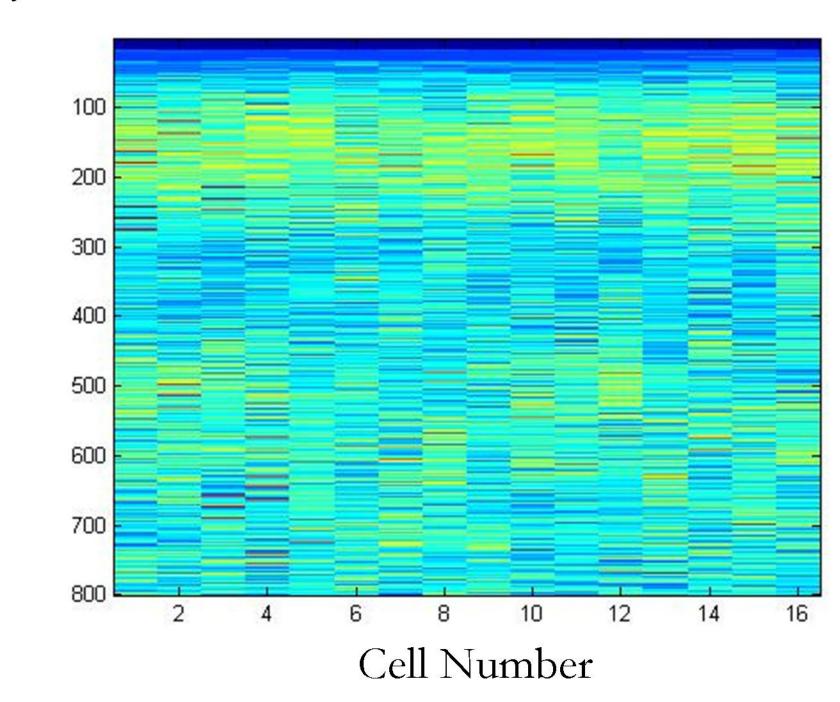
$\frac{dp}{dp} = -\beta(p_i - m_i)$

Problems

Nature is messy – the genes don't turn 'on' or 'off' as neatly as switches and randomness always plays a factor.

A fish is a huge multi cellular organism, made up of trillions of cells each with its own complete set of genes. We want them to all oscillate in phase.

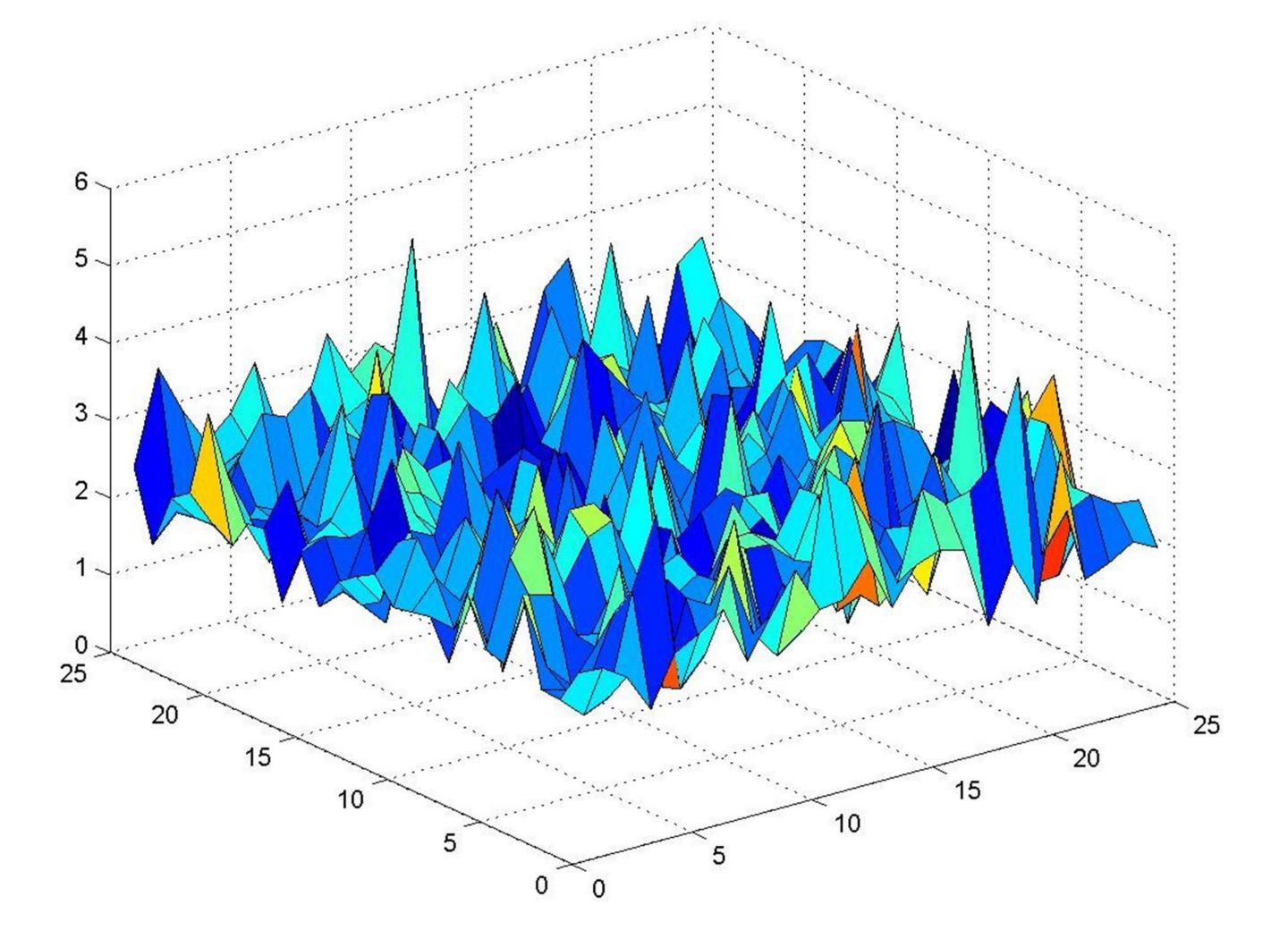
Unsynchronized

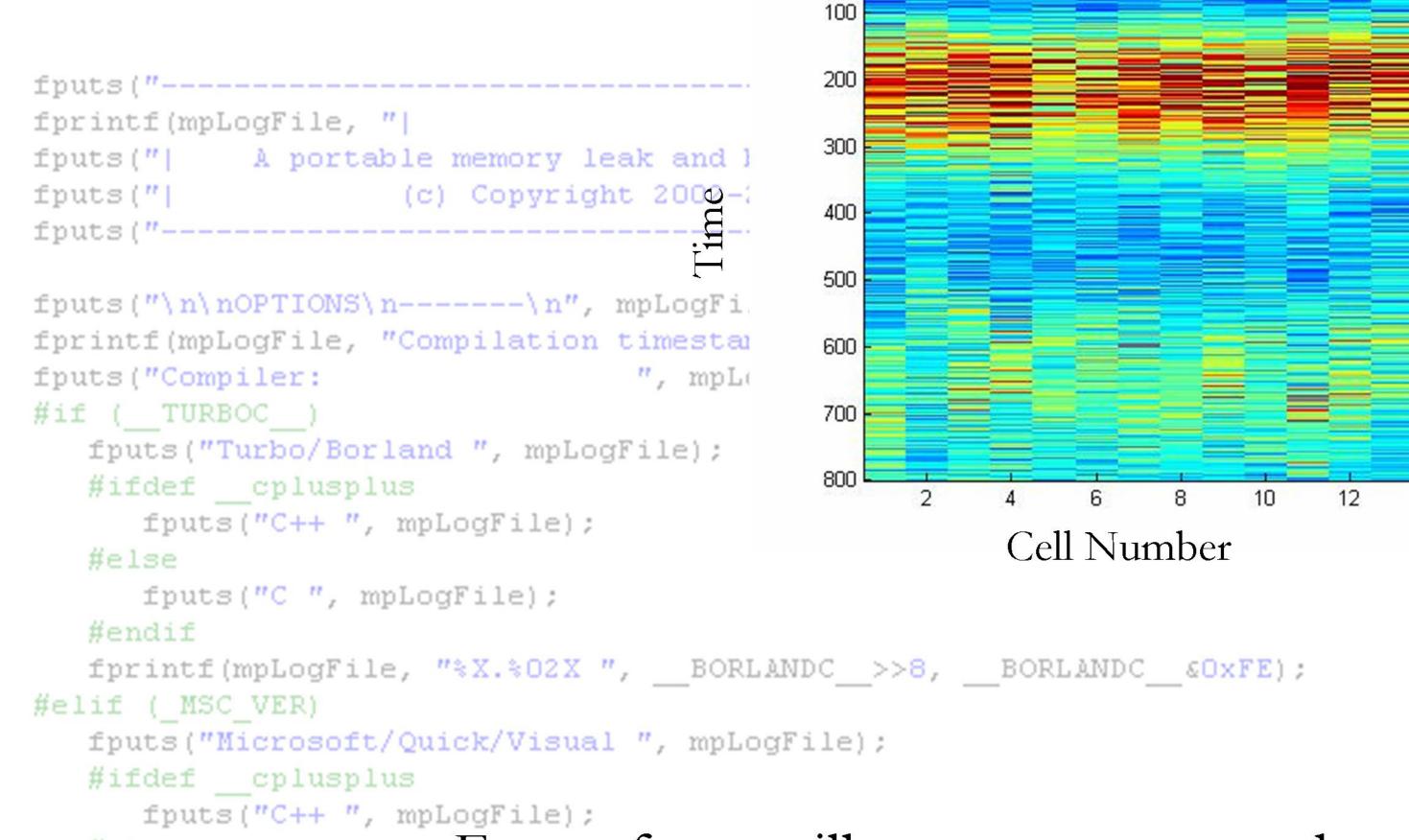


Synchronized

Solution

- Efficient computing, achieved by integrating C++ and MATLAB, provides the modeling capacity for multiple cells in a eukaryote.
- · A graphical user interface was developed to make simulation and analysis user-friendly.
 - · Added features in the interface allow quick control of multiple simulation parameters.
 - Development of different visual analysis tools for clear visual analysis of oscillator behavior.





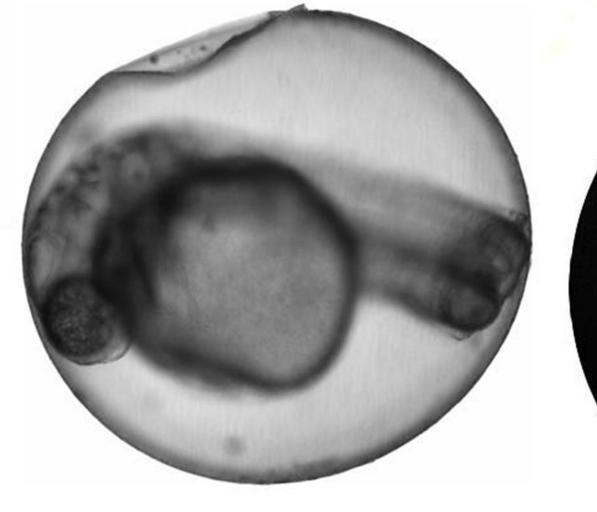
Future focus will concentrate toward quantitative analysis of the oscillator behavior using these efficient tools

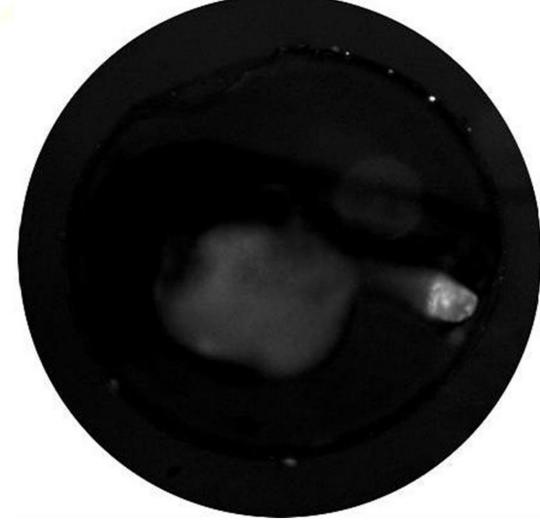
Zebrafish

Danio rerio



- · Easy to breed, and maintain
- Quick developmental cycle
- Transparent embryonic growth





Microinjection

- · Microinjection is used as a means of implementing DNA
- · Small glass needle pierces the embryo and inserts DNA
- · Using low pressure, DNA is pushed into the embryo
- Embryos are highly resilient and regenerate the membrane
- Main problem is low percentage of DNA incorporation into genome
- · Many trials must be attempted

Future and Design

Limitless possibilities are available with this technologies

- Improve life quality
- Cure diseases
- Create novel materials
- Bioremediation
- Designer life forms



Ethics

- Current regulations on the research and application of genetic engineering need to be improved to become more uniform between companies and more effective at preventing misuse
- The public needs to be made more knowledgeable of what this technology is doing and how its use is being controlled, and moreover why it is necessary at all
- The question of how far science is willing to go in modifying the human genome, either to prevent disease or to improve function, needs to be addressed