

Introduction: Experiments

Computational DNA nanotechnology

Damien Regnault <damien.regnault@univ-evry.fr>

Sergiu Ivanov <sergiu.ivanov@univ-evry.fr>

Fall 2022

Context

Unconventional computation

Explore unconventional computation.

Substrate : DNA.

Main goals:

- 1 Better understand the underlying processes.
- 2 Design nanoscale structures.
 - ▶ for biophysics, medicine, etc.

Current status

- Familiarized with the basic experimental protocol.
- Negotiated a lab bench.
- Negotiating AFM access : *in progress*.

Coming up next: Run our first experiment in Évry.

DNA computing

Unconventional computing

- 1 Compute with processes **different** from those which underlie conventional computers.
- 2 Get an **unusual take** on the real-world processes.

Adleman experiment

Hamiltonian path = a path going through all n vertices of a graph exactly once.

- 1 Generate DNA strands representing **random paths**.
- 2 Keep the strands representing paths containing exactly **n vertices**.
- 3 Keep the strands representing paths visiting **every single vertex**.

Adleman, L. M. (1994). **Molecular computation of solutions to combinatorial problems**. Science. 266 (5187): 1021–1024. *available on eCampus*

DNA self-assembly

Build nanoscale structures of DNA by abusing **Watson-Crick complementarity**.

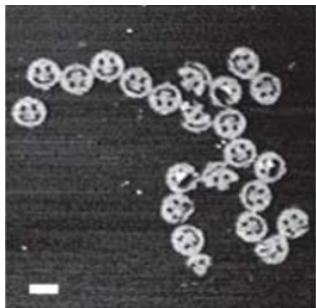
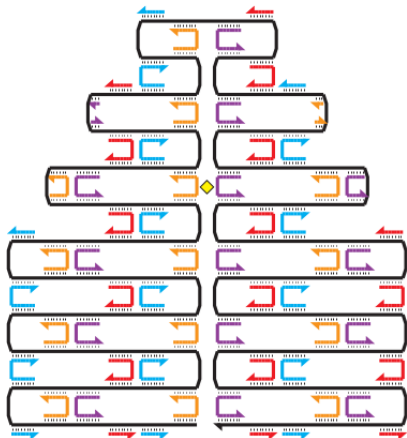
Generally shorter 40–60bp sequences are easier/cheaper to synthesize.



DNA origami

- 1 Take a long DNA sequence, e.g. a phage,
 - ▶ scaffold
- 2 Apply shorter synthetic sequences.
 - ▶ staples
- 3 Scaffold is bent into shape by the staples.

DNA origami experiments



Rothemund P.W. **Folding DNA to create nanoscale shapes and patterns.** *Nature*. 2006 Mar 16;440(7082):297-302.

Not biology yet

Forget about most of the DNA-related biological processes.

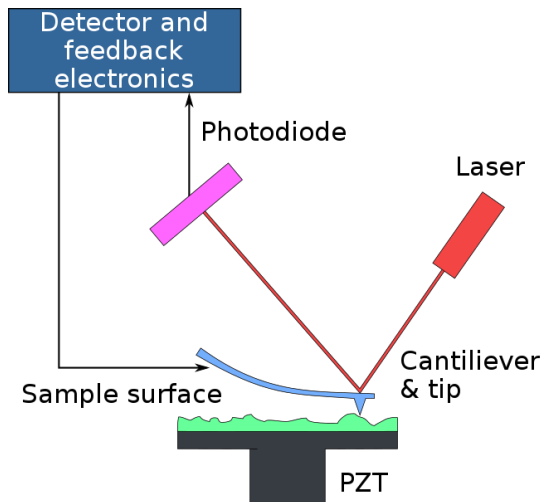
- no ribosomes
- no proteins
- no RNA
- ...

Experimental protocol

Overview

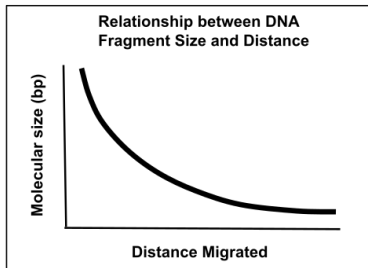
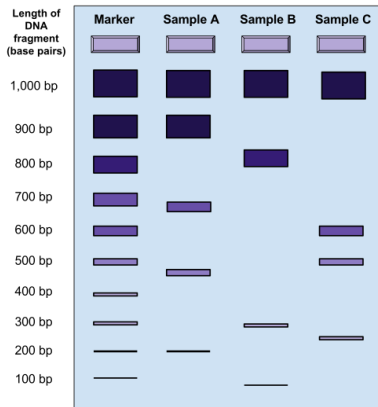
- 1 Grab a scaffold.
- 2 Design the staples.
 - ▶ hard
- 3 Order the staples.
 - ▶ expensive
- 4 Mix everything.
- 5 Heat to $\sim 95^{\circ}\text{C}$.
- 6 Cool down.
 - ▶ **Anneal**: first to 48°C , then slowly to 45°C , then to room temperature.
- 7 Imaging / gel electrophoresis / etc.

Atomic Force Microscopy



https://en.wikipedia.org/wiki/Atomic_force_microscopy

Gel electrophoresis



https://en.wikipedia.org/wiki/Gel_electrophoresis

Perspectives

Perspectives

- DNA breadboards
- Biosensors
- Molecular computers
- Intelligent biomachines

duh

Not biology?

DNA self-assembly can be used in non-biological contexts.

The DNA substrate makes it **look promising for biomedical applications.**

To make those work, we need biology ♥