DNA Computers

Dr. Shyam Kamal

Department of Systems Design and Informatics Kyushu Institute of Technology Japan

kamal@ces.kyutech.ac.jp

13 June 2016

Molecular computing + Molecular programming (Image courtesy: Internet)

Kyutech Japan (Dr. Shyam Kamal)

Э

イロト 人間ト イヨト イ

Genetic Information	DNA Based Computation	DNA-Based Logic Design	Concluding Remarks
Overview			

Genetic Information

DNA Based Computation

DNA-Based Logic Design

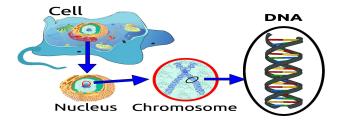
Concluding Remarks

3

Sac

A B + A B +
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 B
 A
 A
 A

Genetic Information



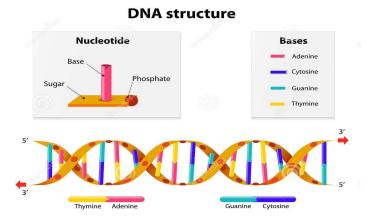
Chromosomes

- composed of protein and deoxyribonucleic acid (DNA)
- both parents contribute material that determines the characteristics of their offspring
- DNA carries the genetic information

A D > A D > A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A

-

DNA strands

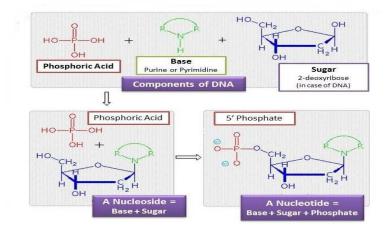


DNA consists of polymer chains, referred to as DNA strands. イロト イヨト イヨト イ

Э

990

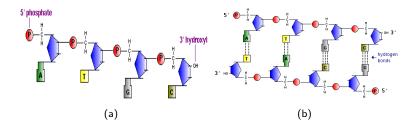
DNA strands in more detail



Any single strand has a natural orientation : a free 5' phosphate group, and the other end has a free 3' deoxyribose hydroxl group.

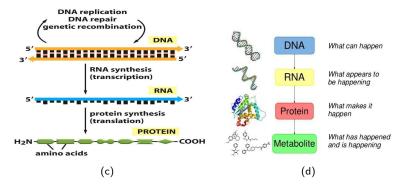
SOC

Single and Double Strand



- Bonding occurs by the pairwise attraction of bases; A bonds with T and G bonds with C.
- ► The pairs (A,T) and (G,C) are therefore known as complementary base pairs.
- ► The two pairs of bases form hydrogen bonds between each other

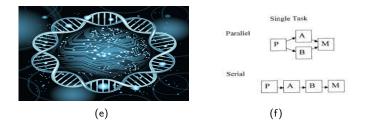
Transcription and Translation



DNA supports two key functions for life

- (1) Coding for the production of proteins which are the working molecules in organisms, (2) Self-replication.
- Remark: One of the most important functions proteins carry out is to act as enzymes.

Reasons for using DNA based Computation



- The information density of DNA is much greater than that of silicon: 1 bit can be stored in approximately one cubic nanometer. Others storage media, such as videotapes, can store 1 bit in 1,000,000,000 cubic nanometer.
- Operations on DNA are massively parallel: a test tube of DNA can contain trillions of strands. Each operation on a test tube of DNA is carried out on all strands in the tube in parallel.

A B A B A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A
 A

Principles of DNA Computing

• Duplication and Transmission of Information:

 DNA can either duplicate the information in DNA molecules or transmit this information to other DNA molecules.

• Representation of Information:

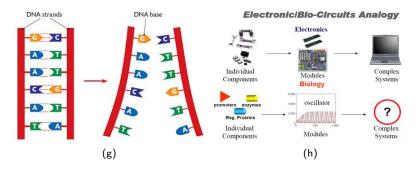
- Present computer using electrical impulses to represent bits of information,
- the DNA computer uses the chemical properties of these molecules by examining the patterns of combination or growth of the molecules or strings.

• Execution of the desired Calculation:

- DNA can do this through the manufacture of enzymes,
- which are biological catalysts that could be called the 'software', used to execute the desired calculation.

イロト イポト イヨト イヨト

Analogy of Electronic Computer



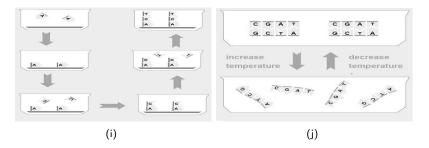
- Electronic computer encode information using {0 1}, similarly DNA using {A G C T}
- In a DNA computer, computation takes place in test tubes or on a glass slide coated in 24K gold.
- ► The input and output are both strands of DNA, whose genetic sequences encode certain information.

Why DNA is a convenient choice?

DNA is a convenient choice because

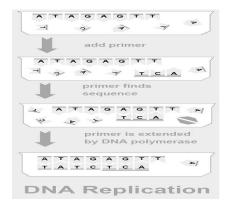
- ► it is both self-complementary, allowing single-stranded DNA to select its own complement
- can easily be copied
- Molecular biologists have already built a toolbox for manipulating DNA, including
 - restriction enzyme digestion
 - ► ligation
 - sequencing
 - amplification
 - fluorescent labelling

Bio-operations: Synthesizing, Annealing and Melting



- ► Synthesizing: a desired polynomial-length strand used in all models.
- Annealing: bond together two single-stranded complementary DNA sequences by cooling the solution. Annealing in vitro is known as hybridization.
- Melting: break apart a double-stranded DNA into its single-stranded complementary components by heating the solution. Melting in vitro is also known under the name of denaturation.

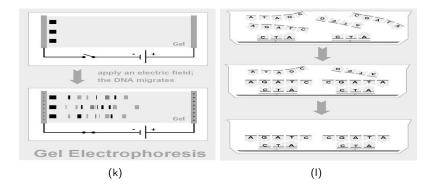
DNA Amplifying:



Amplifying (copying):

- ▶ make copies of DNA strands by using the Polymerase Chain Reaction.
- The replication reaction requires a guiding DNA single-strand called template, and a shorter oligonucleotide called a primer, that is annealed to it.

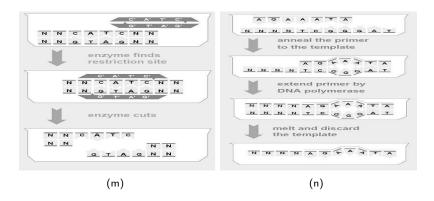
Bio-operations: Separating and Affinity purification



- Separating : the strands by length using a technique called gel electrophoresis that makes possible the separation of strands by length.
- Extracting : those strands that contain a given pattern as a substring by using affinity purification.

(日) (同) (三) (三)

Bio-operations: Cutting and Substituting



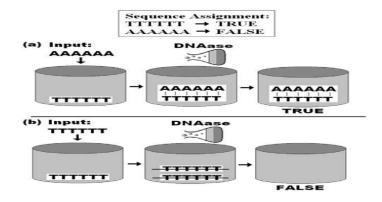
- Cutting : DNA double-strands at specific sites by using commercially available restriction enzymes.
- Substituting: substitute, insert or delete DNA sequences by using PCR site-specific oligonucleotide mutagenesis.

Some other Bio-operations

- Ligating:
 - paste DNA strands with compatible sticky ends by using DNA ligases.
 - ► Indeed, another enzyme called DNA ligase, will bond together, or "ligate", the end of a DNA strand to another strand.
- Marking single strands by hybridization:
 - complementary sequences are attached to the strands, making them double-stranded.
 - The reverse operation is unmarking of the double-strands by denaturing, that is, by detaching the complementary strands.
- Destroying the marked strands by using exonucleases:
 - ► By using enzymes called exonucleases, either double-stranded or single-stranded DNA molecules may be selectively destroyed.
- Detecting and Reading:
 - ▶ given the contents of a tube, say "yes" if it contains at least one DNA strand, and "no" otherwise.
 - PCR may be used to amplify the result and then a process called sequencing is used to actually read the solution.

SOR

DNA-Based NOT Gate



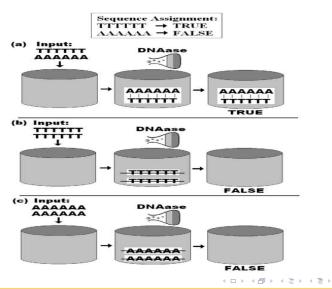
- (a) A false input results in a double stranded sequence representing a truth evaluation
- ► (b) a true input does not the input sequence is true, then TTTTTT will not bind with the base TTTTTT sequence. DNAase will destroy both sequences; no double stranded sequences will be observed. < => <=> =

Molecular computing + Molecular programming (Image courtesy: Internet)

Kyutech Japan (Dr. Shyam Kamal)

Sac

DNA-Based XOR Gate



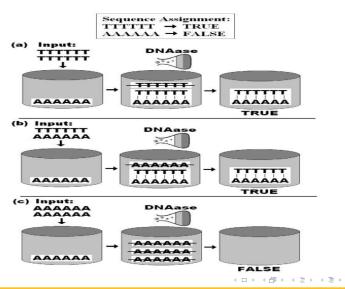
Molecular computing + Molecular programming (Image courtesy: Internet)

Kyutech Japan (Dr. Shyam Kamal)

3

SQA

DNA-Based OR Gate



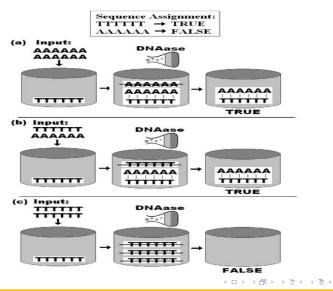
Molecular computing + Molecular programming (Image courtesy: Internet)

Kyutech Japan (Dr. Shyam Kamal)

3

SQA

DNA-Based NAND Gate



Molecular computing + Molecular programming (Image courtesy: Internet)

Kyutech Japan (Dr. Shyam Kamal)

Э

Sac

Concluding Remarks

Advantages

- Perform millions of operations simultaneously.
- Generate a complete set of potential solutions and conduct large parallel searches.
- Efficiently handle massive amounts of working memory.
- They are inexpensive to build, being made of common biological materials.
- ► The clear advantage is that we have a distinct memory block that encodes bits.
- Using one template strand as a memory block also allows us to use its compliment as another memory block, thus effectively doubling our capacity to store information.

1

イロト イポト イヨト イヨト

Concluding Remarks Continued...

Disadvantages:

- Generating solution sets, even for some relatively simple problems, may require impractically large amounts of memory (lots and lots of DNA strands are required).
- Many empirical uncertainties, including those involving: actual error rates, the generation of optimal encoding techniques, and the ability to perform necessary bio-operations conveniently in vitro.

DNA computers could not (at this point) replace traditional computers.

SOR

イロト イポト イヨト イヨト

The End

Molecular computing + Molecular programming (Image courtesy: Internet)

Kyutech Japan (Dr. Shyam Kamal)

2

990