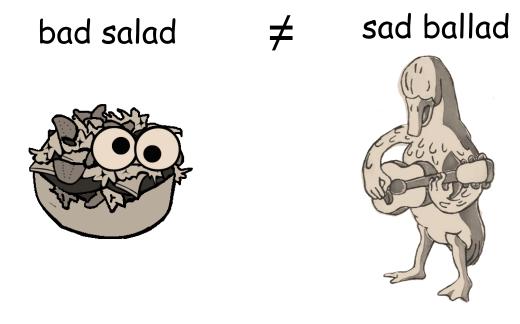
Algorithms on Strings, Trees, and Sequences

Introduction

by Marina Barsky

Sequences (Strings)

Strings are natural groupings of symbols into sequences, where the order has a special significance



Symbols: a b d l s

The course is about:

- String Algorithms (pattern matching, indexing, grouping, prediction): ideas, pseudocode, complexity
- Strings of interest are long and not broken into tokens (words)
- Trees: derived from strings

The area is called **Stringology**

The goal:

- To become familiar with the problems of modern Stringology
- To be able to identify which of these problems are efficiently computable
- Acquire algorithmic tools to solve these problems

Required Background:

- Algorithms
- Data structures
- Probability

Deliverables

- Assignments 40 %
- Class work* 30 %
- Final Project **- 30 %

- * Consists of:
- In-class quizzes to monitor comprehension
- In-class activities to learn how to communicate your ideas

** Term paper or implementation

Long strings

Some texts

Information Retrieval

Collection of documents:

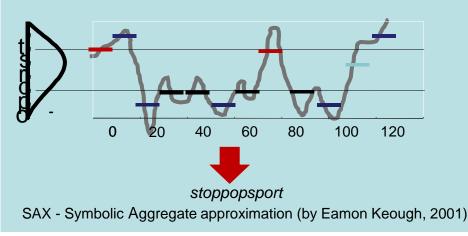
- mouse eats cheese
- cat eats mouse
- snake eats mouse
- Query: who eats mouse?

Strings of words

Music scales



Time series



Very long strings



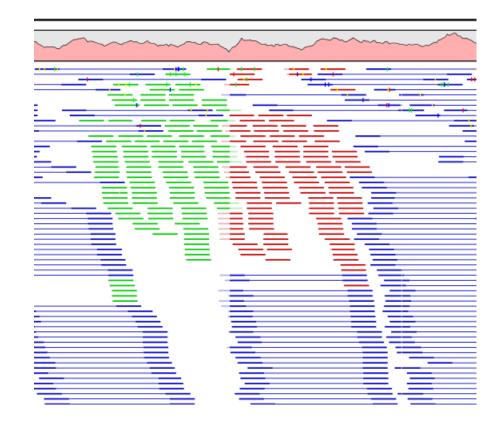
Sequences of molecules in different biological polymers: *DNA*, *RNA*, *proteins*

Digitalization of the molecular code \rightarrow new type of data:

- no clear separation into tokens
- large token size (30,000 and more)
- long "texts" (247 MB in chromosome I)
- virtually unlimited number of different substrings (2x10¹⁷ in Human genome)

Experiment with large strings: https://barsky.ca/marina/UTOR/experiments/bio_example/index.html

Input Dataset: 20 TB of short DNA reads from 232 individuals



Molecular Biology:

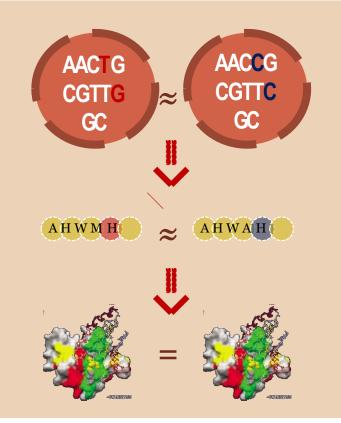
- Describes living things in terms of chemical matter (molecules) and chemical and physical mechanisms
- Studies macromolecules DNA, RNA, protein – and the mechanisms of their interaction

Bioinformatics:

 Applies concepts of Informatics and Computer Science to the field of Molecular Biology – to extract new knowledge from the information embedded in genetic code

Bioinformatics: assumptions

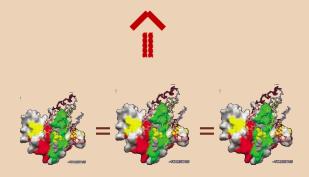
Similar sequences – similar function



If function is unknown – look at similar sequences with known functions

Similar function – partly similar sequences

ATWMH LMGPKPS SHWMH LPDGKPP AHVMH LSMDKPQ



If same function – look at similar substrings which may be responsible for it

Protocol of converting a biological problem into CS problem

- 1. Biological question (find *similar* sequences)
- 2. Formalization (how to measure *similarity*)
- 3. Design an *efficient* algorithm to solve the *formalized* problem
- 4. Model + learning learn parameters of an algorithm from real data
- 5. Evaluation of results distinguish (statistically) significant results from artifacts
- 6. Presentation of the results

Example 1: find similar sequence

- Input:
 - Query: sequence of DNA bases:
 AACCCTTAG
 - The set of sequences of known genes (with their functions):
 ACCTAG
 AGCCCGTA
 AAGCCGCTTA
- Question: which one is the most similar to the query sequence?

Which pair is most similar?



ACCTAG

AACCCTTAG

2

1

AGCCCGTA

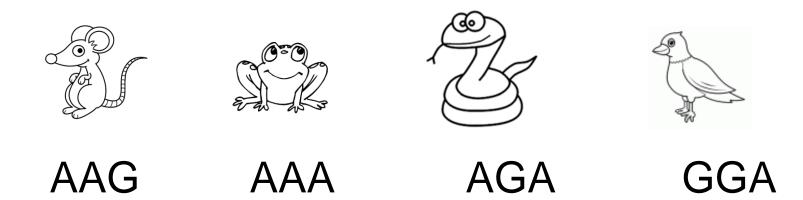
AACCCTTAG

AAGCCGCTTA

3

Example 2: evolutionary tree

Input: four DNA sequences taken from four species.



Following protocol

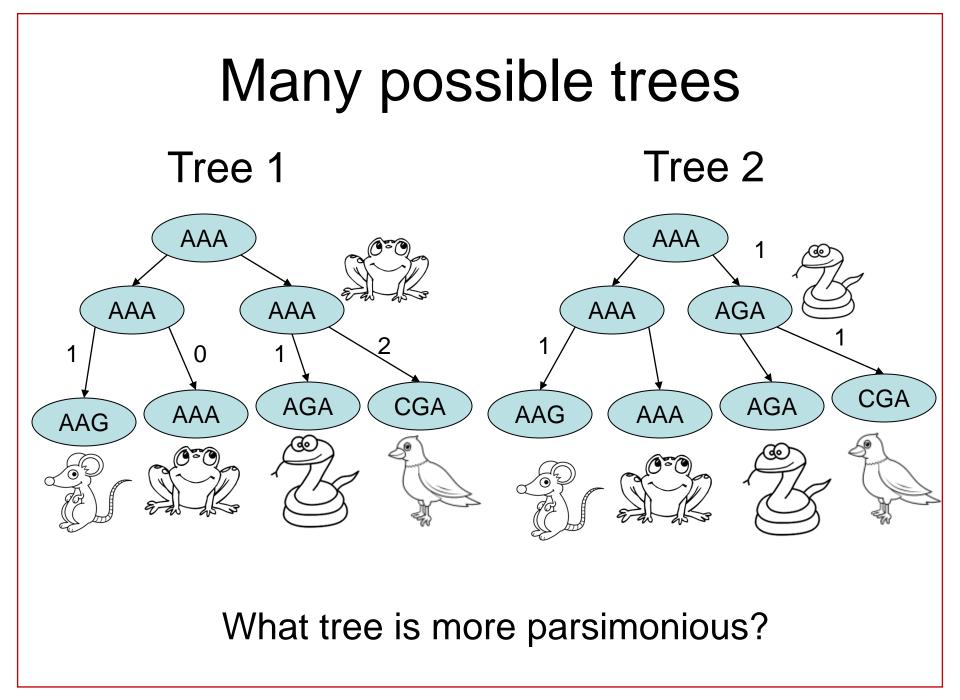
1. Biological question: which evolutionary tree *best* explains these sequences ?

2. Formalization: what is the metric for *the best* tree?

Let it be the parsimony principle

Parsimony principle

- In science, *parsimony* is preference for the least complex explanation. This is regarded as good when judging hypotheses.
- Occam's razor also states the "principle of parsimony": *entia non sunt multiplicanda praeter necessitatem*, is the principle that "entities must not be multiplied beyond necessity": the simplest explanation or strategy tends to be the best one
- Under maximum parsimony, <u>the preferred</u>
 <u>phylogenetic tree is the tree that requires the</u>
 <u>smallest number of evolutionary changes</u>.



Next steps

- 3. Efficient algorithm: how can we compute the best tree efficiently?
- 4. Adjusting parameters from the data: A is more likely to be replaced by G or by T?
- 5. Significance: is the best tree found significantly (statistically) better than others ?
- 6. Present results as a tree
- The main question remains: does the tree make biological sense ?

We will discuss solutions to the following sample problems

- Sequence comparison
- Pattern discovery
- Gene finding
- Sequence-based evolution

Algorithmic Tools: outline

- Discrete algorithms:
 - Combinatorial pattern matching
 - String indexing
 - Dynamic programming
- Probabilistic models:
 - Hidden Markov Models
 - Maximum likelihood
 - Bayesian inference
- Hard problems:
 - Heuristics
 - Approximation algorithms

'Strings' of life

- DNA
- RNA
- Proteins