# CSCI2950-C: Topics in Computational Biology: Genomes, Networks, and Cancer, Fall 2009

Meeting time: Tu/Th 10:30-11:50 CIT 241

Instructor: Ben Raphael, CIT 505, braphael@brown.edu Website: http://cs.brown.edu/courses/csci2950-c/

#### Description

We will examine new and classic problems in computational biology including: genome assembly, genome rearrangements, phylogeny, and cellular interaction networks. We will introduce algorithmic, modeling, and machine learning challenges that emerge in these topics, and explore applications to The Cancer Genome Atlas, an ongoing genome project for cancer.

The course will be organized in seminar style. Following introductory lectures on each of the topics listed below, students will read and present recent research papers, and will undertake a research project.

#### **Prerequisites**

An undergraduate-level background in algorithms and/or probability/statistics.

## Grading

Paper Reviews 30%
Presentations 25%
Project 40% total
10% Written proposal.
20% Final written report.
10% Final presentation.
Participation 5%

**Paper Reviews**: You will submit a written review for each discussion paper, with the exception of those that you present. Reviews are due via email (to braphael@cs.brown.edu) BEFORE the start of class on the day that the paper is presented. Reviews will be graded on a 3-point scale: 2 = mastered the key issues introduced in the paper 1 = read the paper and understood the basics; 0 otherwise.

**Presentations**: Each student will present at least two papers from the reading list. The exact number will depend on class size. At least three days before the presentation, the student will email the instructor a detailed outline of the presentation. The talk slides or a PDF summary (if a chalkboard talk is given) will be submitted on the day of the presentation to be posted on the web page. A presentation is expected to be about 60 minutes, with the remaining class time for questions and discussion.

<u>Project</u>: The project is a semester-long effort to further study one of the class topics. The project could range from theoretical (e.g. designing a new algorithm and proving its correctness), to the practical (a software implementation) depending on the interest of the student. Projects can be undertaken in groups of 1-3 students. A list of suggested projects will be distributed in the third week. For more details, see the instructor.

<u>Participation</u>: You are strongly encouraged to contribute to the discussion by asking questions, making observations, identifying strengths and weaknesses of the approaches under discussion. Critical discussion and analysis are key prerequisites for research.

#### **Course Credits**

PhD: Area B (Algorithms)
ScM: "Theory" or "Practice"\* course.
Significant Programming\*
\*With appropriate class project.

## **Course Outline (Subject to change)**

I. Introduction -- Biology review

Genomes. Central dogma: DNA, RNA, and proteins. Gene regulation.

#### **II. Genome Assembly** (~ 2 lectures, 3 papers)

Shortest common superstring.

Hamiltonian and Eulerian path formulations.

Next-generation sequencing technologies and assembly.

#### **III. Genome Rearrangements** (~ 3 lectures, 2 papers)

Sorting by reversals

Breakpoints and breakpoint graph

Hannenhalli-Pevzner theory for Inversions and Translocations

**Duplications** 

**Probabilistic Methods** 

#### **IV. Phylogeny and Ancestral Genome Reconstruction** (~ 3 lectures, 2 papers)

Phylogenetic Trees

Parsimony and Likelihood Models for Ancestral Reconstruction

Orthology and paralogy; Gene trees and species trees.

#### **V. Cancer Genomes** (~ 3 lectures, 3 papers)

Measuring Mutations in Cancer

Models of Cancer Progression and Evolutionary Dynamics

# **VI. Cellular Interaction Networks** (~ 3 lectures, 4 papers)

Regulatory and Protein Interaction Networks
Network Alignment and Motifs
Models for signaling networks
Network Perturbations in Cancer