# Introduction to Methods in Computational Biology and Genomics

#### C. Ryan Campbell

Duke University c.ryan.campbell@duke.edu

August 29, 2017

#### Overview

- Course
  - Course Intro
  - Teaching Philosophy
- In-Class Activity
- Computing and Genomics
  - Computer Requirements
  - Genomics: Why We're Here

## Today's Goals

- Get familiarized with course format
- Meet each other and myself
- Understand course expectations
- Know my teaching methods and reasons for offering this course

## Course Objectives

#### Objective 1

Learning programming basics and best practices within a biology framework (i.e. basic building blocks for students to use biological software)

## Course Objectives

#### Objective 1

Learning programming basics and best practices within a biology framework (i.e. basic building blocks for students to use biological software)

#### Objective 2

Learning the statistics that underlie these tools and the rapidly evolving suite of measures used in "genomics"

## Course Objectives

#### Objective 1

Learning programming basics and best practices within a biology framework (i.e. basic building blocks for students to use biological software)

#### Objective 2

Learning the statistics that underlie these tools and the rapidly evolving suite of measures used in "genomics"

#### Objective 3

Combine 1 and 2 to test biological hypotheses with RNAseq data and present those results in an "IMRD" format (pronounced EM-rod)

# Course Syllabus

## Course Grade

Honor Code

- Honor Code
- Group Work vs. Own Work

- Honor Code
- Group Work vs. Own Work
- No Direct Attendence Policy

- Honor Code
- Group Work vs. Own Work
- No Direct Attendence Policy
- Interact and Contribute in Class

#### Potential Course Policies - TBD

Latest pre-class email time (5pm? 9pm?)

#### Potential Course Policies - TBD

- Latest pre-class email time (5pm? 9pm?)
- Slack?

#### Potential Course Policies - TBD

- Latest pre-class email time (5pm? 9pm?)
- Slack?
- Office Hours?

Bring a laptop

- Bring a laptop
  - Yes, Every Day

- Bring a laptop
  - Yes, Every Day
- Tuesdays = Lecture

- Bring a laptop
  - Yes, Every Day
- Tuesdays = Lecture
- Thursdays = Lab

## Project

## **Teaching Philosophy**

- Clear Goals
- Active Learning
- Student Driven

#### Clear Goals

- Presented before each class
- Concepts or skills to focus on
- Call me on it if I forget them

## **Active Learning**

- Student Participation
- A different style than lecture-based courses (flipped courses fall into this category)
- Natural fit for smaller class size, advanced material, and learning skills

#### Student Driven

- Student Participation Required
- Work through examples in class and apply to your own question
- Many skills need to be practiced, not taught
- Grand Bargain

Pair up \*randomly\*

- Pair up \*randomly\*
- Fill out this Google form

- Pair up \*randomly\*
- Fill out this Google form
- Make a slide in the Google Slideshow for your partner with their answers, including (at minimum):

- Pair up \*randomly\*
- Fill out this Google form
- Make a slide in the Google Slideshow for your partner with their answers, including (at minimum):
  - Name
  - Picture
  - 'Three words' response
  - Two answers from "Whimsy"
  - Feel free to expand (see my slide for reference)

- Pair up \*randomly\*
- Fill out this Google form
- Make a slide in the Google Slideshow for your partner with their answers, including (at minimum):
  - Name
  - Picture
  - 'Three words' response
  - Two answers from "Whimsy"
  - Feel free to expand (see my slide for reference)
- Introduce partner to class



## Computer Requirements

- Bring your computer to class
- Run some software locally
  - How much storage do your computers have?
- Connect to cluster

#### Research Tools

- R
- Statistical software, data visualization and analysis

#### Research Tools

- R
- Statistical software, data visualization and analysis
- kallisto
  - Fast RNAseq analysis with a command line interface

#### Research Tools

- R
  - Statistical software, data visualization and analysis
- kallisto
  - Fast RNAseq analysis with a command line interface
- github
  - Website for version control as well as script and code storage

## **Programming Languages**

- R
- free statistical software
- bin/bash
  - unix language, automate many tasks, interact with cluster computers

## Software

● R -¿ R-Studio

## Software

- R-Studio ن R
- github -¿ SourceTree

### Software

- R -¿ R-Studio
- github -¿ SourceTree
- data analysis software -¿ fastqc/trimmomatic/kallisto

### **Cluster Computing**

- Duke Computing Cluster
- SLURM workload manager

## Hand Raising Request

Student Driven - Active Learning

## Hand Raising Request

- Student Driven Active Learning
- Raise your hand if you're confused

## Hand Raising Request

- Student Driven Active Learning
- Raise your hand if you're confused
- Provides me with helpful feedback



 Within the field of molecular biology, genomics is the study of genomes, the complete set of genetic material within an organism.

- Within the field of molecular biology, genomics is the study of genomes, the complete set of genetic material within an organism.
- Genomics involves the sequencing and analysis of genomes.

- Within the field of molecular biology, genomics is the study of genomes, the complete set of genetic material within an organism.
- Genomics involves the sequencing and analysis of genomes.
- Genomics is also concerned with the structure, function, comparison, and evolution of genomes.

- Within the field of molecular biology, genomics is the study of genomes, the complete set of genetic material within an organism.
- Genomics involves the sequencing and analysis of genomes.
- Genomics is also concerned with the structure, function, comparison, and evolution of genomes.
- The field also includes studies of intragenomic (within the genome) phenomena such as heterosis (hybrid vigour), epistasis (effect of one gene on another), pleiotropy (one gene affecting more than one trait) and other interactions between loci and alleles within the genome.

- Within the field of molecular biology, genomics is the study of genomes, the complete set of genetic material within an organism.
- Genomics involves the sequencing and analysis of genomes.
- Genomics is also concerned with the structure, function, comparison, and evolution of genomes.
- The field also includes studies of intragenomic (within the genome) phenomena such as heterosis (hybrid vigour), epistasis (effect of one gene on another), pleiotropy (one gene affecting more than one trait) and other interactions between loci and alleles within the genome.
- In contrast to genetics, which refers to the study of individual genes and their roles in inheritance, genomics uses high throughput DNA sequencing and bioinformatics to assemble, and analyze the function and structure of entire genomes.

## **Brief History of Sequencing**

- Allozymes
  - Electrophoresis separates different proteins by amino acid makeup
- Sanger Sequencing
  - Determines the sequences a single piece of DNA up to 500bp
- NGS Next Generation Sequencing
  - Reads sequence of many pieces of DNA many billions of times

## **Brief History of Sequencing**

- Allozymes
  - 1960's
- Sanger Sequencing
  - 1977
- NGS Next Generation Sequencing
  - 2000

## Next Generation Sequencing vs. Sanger

- Output for Quality Tradeoff
  - NGS = High Output / Lower Quality
  - Sanger = Low Output / High Quality

# Next Generation Sequencing vs. Sanger

- Output for Quality Tradeoff
  - NGS = High Output / Lower Quality
  - Sanger = Low Output / High Quality
- NGS Methods and Machines
  - IonTorrent
  - Illumina
  - PacBio

# Next Generation Sequencing vs. Sanger

- Output for Quality Tradeoff
  - NGS = High Output / Lower Quality
  - Sanger = Low Output / High Quality
- NGS Methods and Machines
  - PyroSeq
  - IonTorrent
  - Illumina
  - PacBio



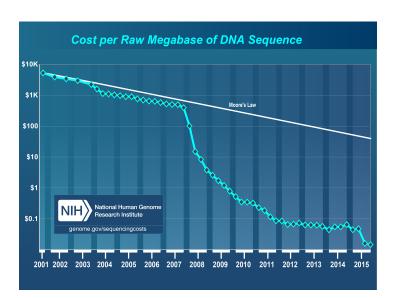


This meme is everywhere, so I thought I'd add a biology twist to it.

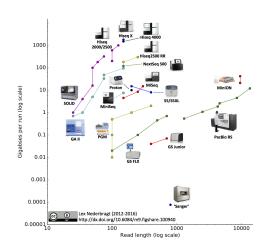


7:11 PM - 25 Aug 2017

## Sequencing Cost



### **Sequencing Output**



More and more data can be generated cheaper

- More and more data can be generated cheaper
- Data length and quality are both improving

- More and more data can be generated cheaper
- Data length and quality are both improving
- How does this change the scope of research?

- More and more data can be generated cheaper
- Data length and quality are both improving
- How does this change the scope of research?
  - (hint: Sanger is good for studying what?)

### **Course Motivations**

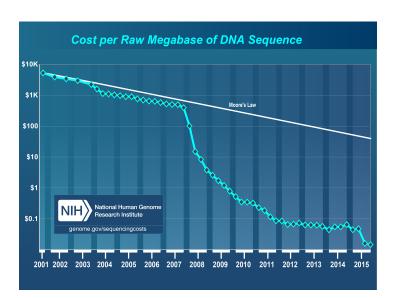
- Biologist increasingly need to be programmers
- Topics often aren't introduced to undergraduates
- Cheap and Free Data

## Computational Need

- Biologist increasingly need to be programmers
- Many hypotheses are tested by generating piles of data
- Regardless of your future plans, programming and hypothesis testing are skills that all STEM students should have

## Cheap/Free Data

### Cheap/Free Data



## Cheap/Free Data

- Data is cheap and often free
- Computation is getting faster
- Combined, this means student projects are feasible

### Personal Experience

- Worked in Duke IGSP/CHGV Sequencing Core for 4 years
  - Ran Illumina GA, GA2, HiSeq2000
- PhD Thesis on Mouse Lemur Genomics
  - Whole Genome Sequencing and Mutation Rate
  - Rates of Sperm Gene Evolution

## The End