## 36. Analysis of single-cell CRISPR screens1 unit, Thomas Norman, November 17, 2025

- 1. CRISPR screens: design principles, readout
  - a. Quantification of fitness CRISPR screens (MAGECK)
- 2. High-content readouts for inferring gene function
  - a. Genome-scale Perturb-seq
  - b. Optical screens
- 3. Guide assignment and incomplete penetrance
  - a. Molecular strategies for barcode capture
  - b. Guide assignment (crispat)
  - c. Incomplete penetrance (Mixscape)
- 4. Achieving greater capacity through multiplexing
  - a. "CRISPR-QTL" approaches to studying enhancers
  - b. Differential expression (SCEPTRE)
- 5. Predictive models of perturbations
  - a. Predicting across context (Compositional Perturbation Autoencoder)
  - b. Genetic interactions (GEARS)

## Paper discussion:

Mapping information-rich genotype-phenotype landscapes with genome-scale Perturb-seq

https://www.sciencedirect.com/science/article/pii/S0092867422005979?via%3Dihub

The datasets from this paper will form the basis of the practical project.

## **Optional background paper:**

High-content CRISPR screening

This is a reasonably up-to-date review covering basics. We will discuss some elements of it in the lecture.

https://www.nature.com/articles/s43586-021-00093-4