# Mathematical Frameworks for Integrative Analysis of Multi-omics Biological Data

This manuscript (<u>permalink</u>) was automatically generated from <u>BIRSBiointegration/whitePaper@6b895b4</u> on April 22, 2020.

# **Authors**

#### • Kim-Anh Lê Cao

D 0000-0003-3923-1116 · ☐ mixOmicsTeam · ☑ mixOmicsTeam

Melbourne Integrative Genomics, School of Mathematics and Statistics, University of Melbourne, Australia · Funded by Grant National Health and Medical Research Council Career Development fellowship (GNT1159458).

#### • Jane Roe

Department of Something, University of Whatever; Department of Whatever, University of Something

# **Abstract**

# Introduction to single cell and imaging multi-omics

Testing github pages.

# **Current multi-omic technologies**

# **Challenges for interpretation**

Need for technology-specific questions and analysis methods vs one size fits all data blender

# **Case studies**

# scNMT-seq as a case-study for epigenetic regulation

#### Overview and biological question

#### **Computational challenges**

- Identification of multi-omics signatures that characterise lineage, stage or both.
- Handling missing values
- Do epigenetic changes in some genomic contexts affect cell fate decision more than others? If so, how?

### Methods for stats/maths analyses and results summary

# scRNA-seq + FISH as a case study for spatial transcriptomics

# Overview and biological question

#### **Computational challenges**

- Can scRNA-seq data be overlaid onto seqFISH for resolution enhancement
- What is the minimal number of genes needed for data integration?
- Are there signatures of cellular co-localization or spatial coordinates in the non-spatial scRNA-seq data?

#### Methods for stats/maths analyses and results summary

#### Spatial proteomics and cross-study analysis

#### Overview and biological question

# **Computational challenges**

- Integrating partially-overlapping proteomic data collected on different patients with similar phenotypes
- Integration of spatial x-y coordinate co-location and co-expression
- Integration with other 'omics datasets (e.g., scRNA-seq) to support the results of these proteomic analyses
- Can we predict the spatial expression patterns of proteins measured on mass-tag but not measured in the MIBI-TOF data?

• What additional information can we learn about the different macrophage and immune populations in breast cancer by conducting integrated analyses of these datasets?

# Methods for stats/maths analyses and results summary

# Overview of common analytical methods spanning technologies / case studies

- matrix factorization
- neural network / autoencoders

# Data structures and software packages to enable analyses

- Bioc/multiAssayExperiment for single cell
- new classes for proteomics
- PyTorch

# Techniques and challenges for benchmarking methods

- realistic simulation studies
- cross-study validation
- benchmark datasets

# **Discussion**

**Emerging analytical methods and technologies** 

Community needs for data structures, analysis methods, etc

# References