Zhen Miao

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EDUCATION

Perelman School of Medicine, University of Pennsylvania, Philadelphia, PA, USA

2019-Present

Ph.D. in Genomics & Computational Biology

- Advisor: Dr. Junhyong Kim
- Thesis: Statistical modeling of single cell multiomics data

Wharton School, University of Pennsylvania, Philadelphia, PA, USA

2019-Present

Dual M.A. in Statistics

- Courses: Applied Regression & Analysis of Variance, Probability, Advanced Statistical Inference I & II, Bayesian Statistics, Stochastic Process, Nonparametric Statistics, Big Data in Genomics, Categorical Data Analysis & Survival Analysis
- Advisor: Dr. Nancy Zhang
- Thesis: Statistical modeling with spatial transcriptomics at single cell or sub-cellular resolution

Nankai University, Tianjin, China

2015-2019

B.S. in Biological Sciences with a minor in Applied Mathematics

- Program: Being selected to Poling Program, a research-training program for elite students
- *Ranking*: 1/27 in Poling Program
- Honors: Merit Student Title for all years, National Scholarship, and Outstanding Graduate title

RESEARCH INTERESTS

- Multi-omics data integration for analyzing tissue dynamics & complex disease traits
- Computational methods in single cell & spatial biology data
- Nephron development and renal diseases

SELECTED PUBLICATIONS

* indicates co-first authorship

First or co-first author:

- 1. **Miao Z**, Wang J, Park K, Kuang D, and Kim J. Model-based compound hypothesis testing for snATAC-seq data with PACS. *bioRxiv* (2023)
- 2. **Miao Z** and Kim J. Uniform quantification of single nucleus ATAC-seq data with Paired-Insertion Counting (PIC) and model-based insertion rate estimator. *Nature Methods* (In Press).
- 3. **Miao Z**, Humphreys BD, McMahon AP, Kim J. Multiomics integration in the age of million single cell data. *Nature Reviews Nephrology* (2021)
- 4. **Miao Z***, Balzer MS*, Ma Z*, et al., Single cell resolution regulatory landscape of the mouse kidney highlights cellular differentiation programs and renal disease targets. *Nature Communications* (2021)

Contributing author:

- 5. Liu H, Doke T, ..., **Miao Z**, et al., Epigenomic and transcriptomic analyses define core cell types, genes and targetable mechanisms for kidney disease. *Nature Genetics* (2022)
- 6. Govek KW*, Troisi EC*, **Miao Z**, et al., Single-Cell Transcriptomic Analysis of mIHC Images via Antigen Mapping. *Science Advances* (2021)
- 7. Zhao Q, Wang J, **Miao Z**, Zhang N, Hennessy S, Small D. S, & Rader D. J. "The role of lipoprotein subfractions in coronary artery disease: A Mendelian randomization study." *eLife* (2021)
- 8. Sheng X*, Guan Y*, Ma Z, Wu J, Liu H, Qiu C, Vitale S, **Miao Z**, et al., Mapping the genetic architecture of human traits to cell types in the kidney identifies mechanisms of disease and potential treatments. *Nature genetics* (2021)

- 9. Nguyen A*, Wang K*, Hu G*, Wang X, Miao Z, et al., APOE and TREM2 regulate amyloid-responsive microglia in Alzheimer's disease. Acta Neuropathologica (2020)
- 10. Zhong L, Yao L, Tower RJ, Wei Y, Miao Z, et al., Single cell transcriptomics identifies a unique adipose lineage cell population that regulates bone marrow environment. *eLife* (2020)
- 11. Gao J, Miao Z, Zhang Z, Wei H, and Kurgan L. "Prediction of Ion Channels and Their Types from Protein Sequences: Comprehensive Review and Comparative Assessment." Current Drug Targets (2018)
- 12. Tan T, Zhang X, Miao Z, Yu Y, Du S, Hou X, and Cai J. "A single point mutation in hmgA leads to melanin accumulation in Bacillus thuringiensis BMB181." Enzyme and Microbial Technology (2018)

CONFERENCES PRESENTATIONS

Platform Presentations:

- NIH Junior Atlas Builder 2023 meeting (3 min lightning talk)
 - o Title: Atlas-scale integrative analysis of snATAC-seq data with PACS
- ReBuilding A Kidney (RBK) 2021 semi-annual meeting (25 min platform presentation)
 - o Title: Probabilistic modeling of single cell epigenomics data
- Penn IRM Stem Cell Club 2021 (25 min platform presentation)
 - o Title: Charting kidney development through single cell multiomics study
- The American Society of Nephrology (ASN) 2020 annual meeting (8 min platform presentation)
 - o Title: Single-cell resolution regulatory landscape of the kidney highlights cellular differentiation programs and renal disease targets The 69th American Society of Human Genomics (ASHG) 2019 annual meeting (*15 min platform presentation*)
- - o Title: Characterizing cellular communication in human central nervous system by single-cell RNA-seq
- MidAtlantic Bioinformatics 2019 Conference (5 min lightning talk)
 - Title: Charting cellular communication network in Alzheimer's Disease

Chalk Talks:

GCB program 2022 and 2023 chalk talks (25 min platform presentation)

Poster Presentations:

- The 72nd American Society of Human Genomics (ASHG) annual meeting (2022)
- The American Society of Nephrology (ASN) annual meeting (kidney week, 2022)
- The 70th American Society of Human Genomics (ASHG) annual meeting (2020)
- Mid-Atlantic Bioinformatics Conference (2020)
- Fox Chase Epigenetics Symposium (2019)

AWARDS

- Blavatnik Family Fellowship in Biomedical Research, awarded to six students across Penn Medicine Ph.D. candidates (2022-2023) [link]
- Charles J. Epstein Trainee Awards, awarded to the top 10% of the presenters in Human Genetics Research for the 69th ASHG annual meeting (2019) [link]
- ACGA Trainee Awards, awarded to the top 3 predoctoral students' platform presentations (2019)
- UPenn BGS Travel Grant, awarded to selected students who present in scientific conferences (2019,2022)
- Outstanding Graduate of Nankai University, awarded to selected students with top academic performance (2019)
- National Scholarship, awarded to the top 0.2% of students for academic excellence (2017, 2019)
- Poling Scholarship, awarded to top 10 students in the Poling Program for achievement in Biological Sciences (2018)
- The Fifth Life Sciences Students' Experimental Skills Competition in Tianjin, the First Prize (2017)
- Biochemistry Experimental Skills Competition in Nankai University, the First Prize (2017)

RESEARCH EXPERIENCE

Perelman School of Medicine, University of Pennsylvania, Philadelphia, PA

July 2019-Present

Ph.D. candidate in Dr. Junhyong Kim's Lab, Department of Biology and Department of Computer Sciences Project 1: Statistical modeling of single nucleus ATAC-sequencing (snATAC-seq) data

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- Constructed a statistical framework for snATAC-seq data that adjust for the coverage disparities present in each cell and the problems with excessive zero
- Applied the statistical framework for real data application in cell label prediction, doublet identification, and differentially accessible region identification
- First-authored a paper that is on *bioRxiv*

Project 2: Reviewing existing computational models for single cell multiomics integration

- Summarized the existing approaches into two major categories: integration for matched data and for unmatched data, and discussed four high-level ideas of data integration and their pros and cons
- Reviewed the challenges and future directions for the field of single cell mulitomics
- First-authored a peer-reviewed paper that published in *Nature Reviews Nephrology*

Rotation student in Dr. Pablo Camara's Lab, Department of Genetics

- Contributed to the development of STvEA, a computational tool for integrating spatial proteomics data with CITE-seq data (single cell transcriptome and epitopes)
- Evaluated the performance of STvEA under scenarios with down-sampled cells or protein penal, and implemented a LASSO-based feature selection algorithm for STvEA
- Co-authored a peer-reviewed paper that published in Science Advances

Rotation student in Dr. Katalin Susztak's Lab, Department of Genetics

- Analyzed kidney snATAC-seq data in adult and P0 murine kidney, conducted manual cell type annotation and cell-type specific open chromatin peak calling
- Identified transcription factors (TFs) that are specific to major kidney cell types
- Investigated the open chromatin dynamics during kidney cell differentiation from nephron progenitors to proximal or distal tubules, and revealed potential genes that are involved to the process
- First-authored a peer-reviewed paper that published in *Nature Communications*

Perelman School of Medicine, University of Pennsylvania, Philadelphia, PA

July 2018-June 2019

Research Intern in Dr. Mingyao Li's Lab, Department of Biostatistics, Epidemiology & Informatics

- Built a ligand-receptor interaction network between microglial sub-clusters, based on single-nuclei RNA sequencing data from 16 human brains with or without Alzheimer's disease (AD)
- Analyzed expression patterns of GWAS identified genes in microglial sub-clusters, indicating a disease-related dysfunctional sub-cluster
- Investigated the relationship between APOE, TREM2 (the top two risk genes for AD), and the complement pathway in each microglial sub-cluster
- Studied the pseudo-time dynamics of transcription factors in microglial sub-clusters, and investigated TFs that trigger the change of microglial states

School of Mathematical Sciences, Nankai University, Tianjin, China

October 2017-July 2018

Undergraduate Researcher in Dr. Jianzhao Gao's Lab, Department of Informatics and Data Science

- Mapped m6A modification data onto GRCh38 genome, encoded features based on flanking sequence, nearby 3mer nucleotide frequency, and solvent accessibility computed in silico
- Conducted feature selection, trained Support Vector Machine (SVM) by selected features, and optimized SVM parameters by 5-fold cross-validation to build an m6A modification predictor

TEACHING EXPERIENCE

Graduate teaching assistant, University of Pennsylvania, Philadelphia, PA

2021 Fall

GCB-534 Statistics for genomics and biomedical informatics

Advising committee for GCB students, University of Pennsylvania, Philadelphia, PA

2021-2023

Role: offer advice on courses for first-year Ph.D. students tailored by their background and research interest, offer advice and guidance on candidacy exam for second-year Ph.D. students

Student Supervisory

Research Assistants: Ziyuan Ma, Kernyu Park

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SKILLS

- *Programming*: Proficient in Python, R, C++, Shell Scripting
- Statistical Modeling: categorical data analysis (GLM), Bayesian modeling (EM algorithm, MCMC), time-series data analysis, survival analysis, non-parametric statistics
- Bioinformatics packages: Proficient in Seurat, SnapATAC, ArchR, edgeR, limma, STAR, GenomicRanges

INTERNSHIP & SUMMER SCHOOL

Computational Biology group, 23andMe, Sunnyvale, CA, USA

June-Aug 2022

Graduate Student Intern, Supervisor: Robert Tunney, Ph.D.

- Engineered codes to enable parallel computing and more efficient data handling for identifying tissue-of-action (TACTICAL), so that the speed is ~200X faster
- Extended the TACTICAL framework to apply to single cell ATAC-seq data for fine-grained detection of disease-associated cell types

Institute for Interdisciplinary Information Sciences, Tsinghua University, Beijing, China

June 2018

Computer Science and Technology Summer School Participant

- Attended lectures on bioinformatics and learned interdisciplinary knowledge on science
- Initiated a project on *The Prediction of Novel Drug-Target Interaction* based on diverse drug-related information, and gave an oral presentation on the methods

Shenzhen Institutes of Advanced Technology, Chinese Academy of Sciences, Shenzhen, China

July 2017

Synthetic Biology Summer School Participant

- Attended lectures and seminars on the Current Development of Synthetic Biology & Synthetic Genomics
- Wrote a proposal on *Genetically Modified Microbes in Heavy Metal Control* with 5 other participants, and gave an oral presentation, ranked 1st among 10 groups