# Jake Yeung

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## Research Plan

I am a computational biologist working at the interface of statistical/machine learning, functional genomics technologies, and genome biology. I develop computationally-driven experimental designs and biophysically interpretable analysis methods to understand how cells coordinate different genome regulatory layers in healthy states, and how they respond to curative therapy in diseased states.

## Professional Preparation

#### 2024 - Present Principal Scientist, Group Leader,

Genentech, Genentech Computational Sciences (gCS),

Analytical Methods.

#### Sep 2021-2023 **Independent Staff Scientist**,

Institute of Science and Technology (IST) Austria,

Computational biology and machine learning lead,

Two parental leaves 2022-01 and 2023-12.

#### Mar-Sep 2021 Machine Learning and Artificial Intelligence Team Leader,

Wellcome Sanger Institute, University of Cambridge,

Single-cell and spatial genomics.

## 2020 Human Frontier Science Program Fellow,

Hubrecht Institute for Developmental Biology and Stem Cell Research,

Group of Prof. Alexander van Oudenaarden, Single-cell genomics.

#### January 2019 PhD in Biotechnology and Bioengineering,

École Polytechnique Fédérale de Lausanne (EPFL). School of Biotechnology and Bioengineering, Group of Prof. Felix Naef, Computational Systems Biology.

Committee members: Prof. John Hogenesch, Prof. Erik van Nimwegen, Prof. Joachim Lingner EPFL Outstanding PhD Thesis Distinction (top 8%)

# April 2014 MSc in Genome Science and Technology,

University of British Columbia.

## June 2012 BSc in Chemical Engineering,

University of Waterloo,

Distinction and Dean's Honours.

Manuscripts (\* denotes co-first author, bold denotes a top-5 publication)

# 2023 J. Yeung\*, M. Florescu\*, P. Zeller\*, B. A. d. Barbanson, Max D. Wellenstein,

A. v. Oudenaarden

scChIX-seq infers dynamic relationships between histone modifications in single cells, Nature Biotechnology, 10.1038/s41587-022-01560-3.

"Behind the paper" post: https://tinyurl.com/y35yusxp

# 2022 P. Zeller\*, J. Yeung\*, H. V. Gaza, B. A. d. Barbanson, V. Bhardwaj, M. Florescu, A. v. Oudenaarden

Single-cell sortChIC identifies hierarchical chromatin dynamics during hematopoiesis,

*Nature Genetics*, https://doi.org/10.1038/s41588-022-01260-3.

Research briefing: https://www.nature.com/articles/s41588-022-01259-w

- 2021 J. Mermet\*, <u>J. Yeung</u>\*, F. Naef, *Oscillating and stable genome topologies underlie hepatic physiological rhythms during the circadian cycle*, *PLOS Genetics*, 17(2), e1009350.
- 2021 N. Phillips, A. Hugues, <u>J. Yeung</u>, E. Durandau, D. Nicolas, F. Naef, *The circadian oscillator analysed at the single-transcript level*, *Molecular Systems Biology*, 17(3), e10135.
- 2019 C. Hor\*, J. Yeung\*, M. Jan, Y. Emmenegger, J. Hubbard, I. Xenarios, F. Naef, P. Franken,

  Sleep-wake-driven and circadian contributions to daily rhythms in gene expression and chromatin accessibility in the murine cortex, *Proceedings of the National Academy of Sciences of the United States of America*, 116(51):25773–25783, 2018.
- 2019 B. Weger, C. Gobet, <u>J. Yeung</u>, E. Martin, S. Jimenez, A. Balvay, A. Foussier, A. Charpagne, C. Jason Chou, F. Naef, <u>F. Gachon</u>, The gut microbiome is required for sex-specific diurnal rhythms of gene expression and metabolism, *Cell Metabolism*, 29(2):362–382.e8, 2019.
- 2018 F. Gachon, <u>J. Yeung</u>, F. Naef, Cross-regulatory circuits linking inflammation, high-fat diet, and the circadian clock, *Genes & Development*, 32:347–358, 2018.
- 2018 <u>J. Yeung</u>, F. Naef, Rhythms of the Genome: Circadian Dynamics from Chromatin Topology, Tissue-Specific Gene Expression, to Behavior, *Trends in Genetics*, 34(12):915–926, 2018.
- 2018 J. Mermet\*, J. Yeung\*, C. Hurni, D. Mauvoisin, K. Gustafson, C. Jouffe, D. Nicolas, Y. Emmenegger, C. Gobet, P. Franken, F. Gachon, F. Naef,

  Clock-dependent chromatin topology modulates circadian transcription and behavior,

  Genes & Development, 32:347–358, 2018, highlighted by a companion outlook article.
- 2018 J. Wang\*, L. Symul\*, J. Yeung, C. Gobet, J. Sobel, S. Luck, P. Westermark, N. Molina, F. Naef, Circadian clock –dependent and –independent post-transcriptional regulation underlies temporal mRNA accumulation in mouse liver, *Proceedings of the National Academy of Sciences of the United States of America*, 115(8):E1916–E1925, 2018.
- 2018 J. Yeung\*, J. Mermet\*, C. Jouffe, J. Marquis, A. Charpagne, F. Gachon, F. Naef, Transcription factor activity rhythms and tissue-specific chromatin interactions explain circadian gene expression across organs, *Genome Research*, 28(2):182–191, 2018, featured on the cover.
- 2017 R. Shrestha, E. Hodzic, T. Sauerwald, P. Dao, K. Wang, <u>J. Yeung</u>, S. Anderson, F. Vandin, G. Haffari, C. Collins, S. C. Sahinalp, HIT'nDRIVE: Patient-specific multi-driver gene prioritization for precision oncology, *Genome Research*, 27(9):1573–1588, 2017.
- 2016 J. Mermet, <u>J. Yeung</u>, F. Naef, Systems Chronobiology: Global Analysis of Gene Regulation in a 24-Hour Periodic World, *Cold Spring Harbor Perspectives in Biology*, 9(3) a028720, 2016.
- 2014 A. W. Wyatt, F. Mo, K. Wang, B. McConeghy, S. Brahmbhatt, L. Jong, D. M. Mitchell, R. L. Johnston, A. Haegert, E. Li, J. Liew, <u>J. Yeung</u>, ..., and C. C. Collins, The inter-tumor transcriptome heterogeneity of high-risk primary prostate cancer, *Genome Biology*, 15(8):426.
- 2014 R. Shrestha\*, E. Hodzic\*, <u>J. Yeung</u>\*, K. Wang, T. Sauerwald, P. Dao, S. Anderson, H. Beltran, M. Rubin, C. Collins, G. Haffari, S. C. Sahinalp, HIT'nDRIVE: Multi-Driver Gene Prioritization based on Hitting Time, *Research in Computational Molecular Biology*, 293–306, Cham, 2014.
  - Talks at Conferences, Workshops, and Discussion Groups
- 2022 Learning relationships between histone modifications in single cells. NeurIPS 2022 Workshop LMRL. Virtual, accepted paper
- 2022 Inferring relationships between histone modifications in single cells. Single-cell Omics Germany Epigenome 2022. Überherrn Germany
- 2022 Models to infer gene regulation from single-cell chromatin modification data: BIRS Mathematical models in biology. Banff Canada

- 2021 Uncovering multi-modal chromatin states in single cells. EMBO: Multiomics to Mechanisms: Challenges in Data Integration (virtual)
- 2021 Deconvolving multiplexed histone modifications in single cells. CSHL: Biology of Genomes (virtual)
- 2021 Statistical challenges in single-cell epigenomics. Hubrecht single-cell workshop (virtual)
- 2020 Using machine learning to make maps of stem cell differentiation. Machine learning in Physics, Chemistry and Materials discussion group (MLDG). Cambridge UK
- 2019 Profiling histone modifications in single cells reveals the dynamics and control of blood cell differentiation. EMBO Workshop: Single Cell Biology. Tokyo Japan
- 2018 Clock-dependent chromatin topology modulates circadian transcription and behavior. EMBL: From Functional Genomics to Systems Biology. Heidelberg Germany
- 2018 Dynamic gene expression and regulatory outputs of the mouse cortex. Society for Sleep medicine, Sleep Research and Circadian rhythms. Basel Switzerland
- 2018 Clock-dependent chromatin topology modulates circadian transcription and behavior. CSHL: Biology of Genomes. Cold Spring Harbor USA
- 2018 Transcription factor activity rhythms and tissue-specific chromatin interactions orchestrate circadian gene expression across organs. Genomic Regulation 2018. Haute-Nendaz, Switzerland
- 2017 Interaction of tissue-specific and circadian transcriptional regulatory networks. European Biological Rhythms Society. Amsterdam, Netherlands
- 2017 Tissue-specific circadian transcriptional regulation. Gordon Conference Seminars on Chronobiology. Stowe, USA
- 2016 Integrated analysis reveals distinct roles of promoters underlying rhythmic gene expression. Gene Regulation and Promoter Meeting: Eukaryotic Promoter Database 30th Anniversary Symposium. Lausanne, Switzerland
- 2016 Integrating functional genomics data reveals tissue-dependent mechanisms underlying circadian gene expression. Society for Research on Biological Rhythms. Tampa Bay, USA

#### — Grants and Awards

- 2022 Invitation to the 72nd Lindau Nobel Laureate Meeting 2023
- 2019 Human Frontier Science Program (HFSP) Fellowship (3-years)
- 2018 Swiss National Science Foundation Postdoctoral Fellowship (1.5-years)
- 2018 EPFL Outstanding PhD Thesis Distinction (top 8%)
- 2018 European Molecular Biology Organization (EMBO) Long-Term Fellowship Finalist (withdrawn after accepting HFSP fellowship)
- 2015 Natural Resources and Engineering Research Council of Canada Doctoral Fellowship (3 years)
- 2012 Genome Science and Technology Graduate Award (2 years)
- 2012 First Prize Walmart Green Student Challenge Competition (\$60000 award)

# Teaching and Mentoring Experience

- 2021 Present **Co-supervising PhD students**, *IST Austria*, Jessica Kirchner, Puzhen Xia.
  - 2021 **Supervising postdoctoral fellows**, *Wellcome Sanger Institute*, Fabiola Curion.
  - 2019 **Episystem Workshop: Single-Cell Epigenetics**, *Hubrecht Institute*,

    Developed and led practical sessions to teach analysis of single-cell epigenetics data.

    Link to course material: https://github.com/avolab/episystem-workshop

Curriculum Vitae: Jake Yeung

## 2014-2018 Computational and Mathematical Modeling in Biology, EPFL,

Developed and supervised exercises for the third-year undergraduate bioengineering course.

#### 2018 Supervising Master's Projects in Computational Biology, EPFL,

Moritz Durtschi: Gaussian process latent variable models for inferring gene expression pseudotime. Alice Hugues: Stochastic trajectory and noise characteristics in gene expression.

# Genomics and Machine Learning Software and Packages

Author scChlX (https://github.com/jakeyeung/scChlX). Statistical inference method for mapping multiple histone modifications in single cells.

sortChIC (https://github.com/jakeyeung/sortchicAnalysis). Statistical analyses of sortChIC data.

CircadianRNASeq (https://github.com/jakeyeung/TissueCiradianAnalysis). Complex-valued matrix decomposition method for extracting periodic signal from temporal gene expression data.

SleepDepAnalysis (https://github.com/jakeyeung/SleepDepAnalysis). Statistical model for integrating RNA-seq with electroencephalography (EEG) time-series data.

Circadian4Cseq (https://github.com/jakeyeung/Circadian4Cseq). Analysis of time-resolved chromatin conformation data.

Contributor gpjax (https://github.com/JaxGaussianProcesses/GPJax). JAX Gaussian Processes.

glmpca (https://cran.r-project.org/web/packages/glmpca/index.html). Generalized linear model extensions to principal component analysis.

SingleCellMultiOmics (https://pypi.org/project/singlecellmultiomics). General framework for processing single-cell genomics and epigenomics data.