

Jake Yeung

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Research Statement

I am a computational and systems biologist combining single-cell genomics technologies, experimental design, and machine learning to understand how physiology is coordinated across multiple biological scales: from single cells, organs, to whole-animal behavior. I develop ways to measure, infer, and model epigenetic marks to quantify relationships between histone modifications in single cells and how they change as cells make decisions.

Professional Preparation

- 2021–Present **Independent Staff Scientist**,
Institute of Science and Technology (IST) Austria,
Computational biology and machine learning,
Parental leave between 2022-01 and 2022-04.
- 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)

- Accepted J. Yeung*, M. Florescu*, P. Zeller*, B. A. d. Barbanson, Max D. Wellenstein, A. v. Oudenaarden**
scChIX infers relationships between histone modifications in single cells,
Accepted in *Nature Biotechnology*,
bioRxiv <https://doi.org/10.1101/2021.04.26.440629>.
Accepted version: <https://tinyurl.com/yysva34t>
- Accepted P. Zeller*, J. Yeung*, H. V. Gaza, B. A. d. Barbanson, V. Bhardwaj, M. Florescu, A. v. Oudenaarden**
Hierarchical chromatin regulation during blood formation uncovered by single-cell sortChIC,
Accepted in *Nature Genetics*,
bioRxiv <https://www.biorxiv.org/content/10.1101/2021.04.26.440606v1>.
Accepted version: <https://tinyurl.com/4w66zhj4>

- 2021 J. Mermet*, J. Yeung*, 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, J. Yeung, 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, J. Yeung, E. Martin, S. Jimenez, A. Balvay, A. Foussier, A. Charpagne, C. Jason Chou, F. Naef, F. Gachon, *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, J. Yeung, F. Naef, *Cross-regulatory circuits linking inflammation, high-fat diet, and the circadian clock*, *Genes & Development*, 32:347–358, 2018.
- 2018 J. Yeung, 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. Westermarck, 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, J. Yeung, 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, J. Yeung, 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. Brahmabhatt, L. Jong, D. M. Mitchell, R. L. Johnston, A. Haegert, E. Li, J. Liew, J. Yeung, ..., 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*, J. Yeung*, 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 (upcoming) LMRL. Virtual
- 2022 Inferring relationships between histone modifications in single cells. Single-cell Omics Germany (upcoming) Epigenome 2022. Überherrn Germany
- 2022 Models to infer gene regulation from single-cell chromatin modification data: 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

Seminars

- 2021 Department seminar. IST Austria (virtual). Host: Prof. Mario de Bono
- 2021 Department seminar. University of California Los Angeles (virtual). Host: Prof. Roy Wollman
- 2020 Life sciences seminar. University of Hong Kong (virtual). Host: Prof. Danny Chan
- 2020 Candidate seminar. Genentech (virtual). Host: Dr. Bill Forrest
- 2020 Department seminar. Genome Institute of Singapore. Host: Dr. Shyam Prabhakar
- 2019 Bioengineering seminar. EPFL. Host: Prof. Felix Naef
- 2019 Faculty of Medicine seminar. University of Hong Kong. Host: Prof. Suet Yi Leung
- 2018 EMBO seminar. University of Edinburgh. Host: Prof. Adrian Bird
- 2018 Seminar. EMBL European Bioinformatics Institute. Host: Dr. John Marioni
- 2018 Seminar. Harvard Medical School. Host: Prof. Stirling Churchman
- 2018 Seminar. Lewis-Sigler Institute. Host: Prof. Joshua Rabinowitz
- 2018 Seminar. Wellcome Sanger Institute. Host: Dr. Sarah Teichmann

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>
- 2014-2018 **Computational and Mathematical Modeling in Biology**, *EPFL*,
Developed and supervised practical exercises for the third-year undergraduate 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 *scChIX* (<https://github.com/jakeyeung/scChIX>). 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 *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.