# **CURRICULUM VITAE**

## PERSONAL INFORMATION

Name: Yaqiang Cao

Organization: System Biology Center, National Heart, Lung, and Blood Institute

(NHLBI), National Institutes of Health (NIH)

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Links: Google Scholar | GitHub

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## **EDUCATION**

2019 Ph.D. in Computational Biology

Chinese Academy of Sciences (CAS)- Max Planck Society (MPG) Partner Institute for Computational Biology, University of Chinese Academy of Sciences (UCAS),

Shanghai, China.

Mentor: Prof. Jing-Dong Jackie Han

2012 B.S. in Bioengineering

East China University of Science and Technology, Shanghai, China.

### RESEARCH EXPERIENCE

2024-Curr. Research Fellow

System Biology Center, National Heart, Lung, and Blood Institute (NHLBI),

National Institutes of Health (NIH), Bethesda, MD, USA.

Mentor: Dr. Keii Zhao

2019-2024 Visiting Fellow

NHLBI/NIH. Bethesda. MD. USA.

Mentor: Dr. Keji Zhao

2011-2018 Graduate Research Assistant

CAS-MPG Partner Institute for Computational Biology, Shanghai, China.

Mentor: Prof. Jing-Dong Jackie Han

### **RESEARCH SUMMARY**

**Focus:** Chromatin regulation in immunity, leukemia, and aging.

Methods: Created computational tools and experimental techniques to map high-

resolution enhancer-promoter interactions and single-cell nucleosomes.

**Discoveries**: Chromatin loops/histone modifications/transcription factors driving T-cell

differentiation, acute myeloid leukemia, B-cell aging.

Collaborations: Cross-disciplinary partnerships bridging basic science and clinical

applications, involving biologists, physicians, and data scientists.

**Impact**: User-friendly software for epigenomic and 3D genomic research;

mechanism understandings for chromatin and clinical translations.

### FIRST/CO-FIRST PUBLICATIONS (14 out of 31)

- \* co-first author # correspondence.
- Zhen, T. \*, *Cao, Y.* \*, Dou, T., Chen, Y., Lopez, G., Menezes, A.C., Wu, X., Hammer, J., Cheng, J., Garrett, Anderson, S., Kirby, M., Wincovitch, S., Sisay, B., Elkahloun, A. G., Wu, D., Castilla, L., Yang, W., Jiang, J., Zhao, K., Liu, P. P. \* (2025). CBFβ-SMMHC-driven leukemogenesis requires enhanced RUNX1-DNA binding affinity in mice. *J Clin Invest*. 10.1172/jci192923.
- Ma, F. \*, Cao, Y. \*, Du, H., Braikia, F., Zong, L., Ollikainen, N., Bayer, M., Qiu, X., Park, B., Roy, R., Nandi, S., Sarantopoulou, D., Ziman, A., Bianchi, A., Beerman, I., Zhao, K., Grosschedl, R., Sen, R. \*. (2024) Three-dimensional chromatin reorganization regulates B cell development during ageing. Nat Cell Biol. 1-12. Liu, S. \*, Cao, Y. \*, Cui, K\*., Ren, G., Zhao, T., Wang, X., Wei, D., Chen, Z., Gurram, R.K., Liu, C. and Wu, C., Zhu, J., & Zhao, K. \* (2024) Regulation of T helper cell differentiation by the interplay between histone modification and chromatin interaction. Immunity. 57(5), 987-1004.
- Cao, Y.\*, Liu, S.\*, Cui, K.\*, Tang, Q., & Zhao, K.\* (2023). Hi-TrAC detects active sub-TADs and reveals internal organizations of super-enhancers. Nucleic Acids Research, 51(12), 6172-6189.
   Cui, K.\*, Chen, Z.\*, Cao, Y.\*, Liu, S.\*, Ren, G., Hu, G., Fang, D., Wei, D., Liu, C., Zhu, J., Wu, C.\*, & Zhao, K.\* (2023). Restraint of IFN-γ expression through a distal silencer CNS–28 for tissue homeostasis. Immunity, 56(5), 944-958 (selected as cover, preview by Immunity)
- 2022 Cao, Y. \*, Liu, S. \*, Ren, G. \*, Tang, Q., & Zhao, K. # (2022). cLoops2: a full-stack comprehensive analytical tool for chromatin interactions. Nucleic Acids Research, 50(1), 57-71.
  Liu, S. \*, Cao, Y. \*, Cui, K., Tang, Q., & Zhao, K. # (2022). Hi-TrAC reveals division of labor of transcription factors in organizing chromatin loops. Nature communications, 13(1), 1-17.
- 2020 *Cao, Y.*\*, Chen, Z.\*, Chen, X.\*, Ai, D.\*, Chen, G., McDermott, J., Huang, Y., Guo, X., & Han, J. \* (2020). Accurate loop calling for 3D genomic data with cLoops. *Bioinformatics*, 36(3), 666-675.
  - Zhen, T. \*, *Cao*, *Y.* \*, Ren, G., Zhao, L., Hyde, R. K., Lopez, G., Feng, D., Alemu, L., Zhao, K., & Liu, P. P. \* (2020). RUNX1 and CBFβ-SMMHC transactivate target genes together in abnormal myeloid progenitors for leukemia development. *Blood*, The Journal of the American Society of Hematology, 136(21), 2373-2385. (selected as Plenary Paper)
  - Han, M. \*, Li, J. \*, *Cao, Y.* \*, Huang, Y., Li, W., Zhu, H., Zhao, Q., Han, J., Wu, Q., Li, J., Feng, J., & Wong, J. # (2020). A role for LSH in facilitating DNA methylation by DNMT1 through enhancing UHRF1 chromatin association. *Nucleic Acids Research*, 48(21), 12116-12134.
  - Zeng, Y. \*, *Cao, Y.* \*, Halevy, R. S. \*, Nguyen, P., Liu, D., Zhang, X., Ahituv, N. \*, & Han, J. \* (2020). Characterization of functional transposable element enhancers in acute myeloid leukemia. *Science China Life Sciences*, 1-13.

- Huang, F. \*\*, *Cao, Y.* \*, Wu, G., Chen, J., Lin, W., Lan, R., Wu, B., Xie, X., Hong, J., & Fu, L. (2020). BMP2 signaling activation enhances bone metastases of non-small cell lung cancer. *Journal of Cellular and Molecular Medicine*, 24(18), 10768-10784.
- 2019 *Cao, Y.* \*, Chen, G. \*, Wu, G. \*, Zhang, X. \*, McDermott, J., Chen, X., Xu, C., Jiang, Q., Chen, Z., Zeng, Y., Ai, D., Huang, Y., &Han, J. # (2019). Widespread roles of enhancer-like transposable elements in cell identity and long-range genomic interactions. *Genome research*, 29(1), 40-52.
- 2016 Wang, L. \*, Xu, X. \*, *Cao, Y.* \*, Li, Z., Cheng, H., Zhu, G., Duan, F., Na, J., Han, J., & Chen, Y. G. (2016). Activin/Smad2-induced H3K27me3 reduction is crucial to initiate mesendoderm differentiation of human embryonic stem cells. *Journal of Biological Chemistry*, jbc-M116.

#### **OTHER PUBLICATIONS**

- Roy, S., Ren, M., Li, P., Cui, K., *Cao, Y.*, Fisk, B., Markowitz, T. E., Redekar, N., Sakamoto, K., Nagao, K., Oh, J., Spolski, R., Liao, W., Dubois, S. P., Kelsall, B. L., Zhao, K., Phelan, J. D., & Leonard, W. J. (2025). BLIMP1 negatively regulates IL-2 signaling in T cells. *Science Advances*, 11(29), eadx8105.
  Khateb, M., Jung, R., Leibou, S., Hadley, P., Yu, Z., Dinerman, A. J., Dulemba, V., Gasmi, B., Levin, N., Kim, P., Bhasin, A., Bhat, D., Sindiri, S., Gartner, J. J., Prickett, T. D., Benzine, T., Farid, S. S., Parkhurst, M. R., Halas, H., *Cao, Y., Zhao*, K., Yang, J. C., Robbins, P. F., Lowery, F., Krishna, S., Heller, T., McVicar, D., Rosenberg, S. A. & Klemen, N. D. (2025). Rapid enrichment of progenitor exhausted neoantigen-specific CD8 T cells from peripheral blood. *bioRxiv*, 2025-05.
- Zhu, X., Chen, X., *Cao, Y.*, Liu, C., Hu, G., Ganesan, S., Veres, T. Z., Fang, D., Liu, S., Chung, H., Germain, R. N., Schwartzberg, P. L., Zhao, K., & Zhu, J. (2024). Optimal CXCR5 Expression during Tfh Maturation Involves the Bhlhe40-Pou2af1 Axis. *bioRxiv*, 2024-05.
- Zhang, X., Jiang, Q., Li, J., Zhang, S., *Cao, Y.*, Xia, X., Cai, D., Tan, J., Chen, J., & Han, J. D. J. (2022). KCNQ1OT1 promotes genome-wide transposon repression by guiding RNA–DNA triplexes and HP1 binding. *Nature Cell Biology*, 1-13. Lin, X., Liu, Y., Liu, S., Zhu, X., Wu, L., Zhu, Y., Zhao, D., Xu, X., Chemparathy, A., Wang, H., *Cao, Y.*, Nakamura, M., Noordermeer, J., Russa, M., Wong, W., Zhao, K., & Qi, L. S. (2022). Nested epistasis enhancer networks for robust genome regulation. *Science*, 377(6610), 1077-1085.
  - Ren, G., Lai, B., Harly, C., Baek, S., Ding, Y., Zheng, M., *Cao, Y.*, Cui, K., Yang, Y., Zhu, J., Hager, G., Bhandoola, A., & Zhao, K. (2022). Transcription factors TCF-1 and GATA3 are key factors for the epigenetic priming of early innate lymphoid progenitors toward distinct cell fates. *Immunity*, 55(8), 1402-1413.
  - Pan, L., Ku, W. L., Tang, Q., *Cao, Y.*, & Zhao, K. (2022). scPCOR-seq enables coprofiling of chromatin occupancy and RNAs in single cells. *Communications Biology*, 5(1), 1-9.
  - Fang, D., Cui, K., *Cao, Y.*, Zheng, M., Kawabe, T., Hu, G., Khillan, J., Li, D., Zhong, C., Jankovic, D., Sher, A., Zhao, K., & Zhu, J. (2022). Differential regulation of transcription factor T-bet induction during NK cell development and T helper-1 cell differentiation. *Immunity*, 55(4), 639-655.

- Ku, W. L., Pan, L., Cao, Y., Gao, W., & Zhao, K. (2021). Profiling single-cell histone modifications using indexing chromatin immunocleavage sequencing. Genome Research, 31(10), 1831-1842.
  Huang, F., Cao, Y., Wang, C., Lan, R., Wu, B., Xie, X., Hong, J., Fu, L., & Wu, G. (2021). PNMA5 promotes bone metastasis of non-small-cell lung cancer as a target of BMP2 signaling. Frontiers in Cell and Developmental Biology, 1400.
- Qiu, X., Ma, F., Zhao, M., *Cao, Y.*, Shipp, L., Liu, A., Dutta, A., Singh, A., Braikia, F.Z., De, S., Wood, W. H., Becker, K.G., Zhou, W., Ji, H., Zhao, K., Atchison, M.L., & Sen, R. (2020). Altered 3D chromatin structure permits inversional recombination at the IgH locus. *Science Advances*, 6(33), eaaz8850.
  Xia, X., Chen, X., Wu, G., Li, F., Wang, Y., Chen, Y., Chen, M., Wang, X., Chen, W., Xian, B., Chen, W., *Cao, Y.*, Xu, C., Gong, W., Chen, G., Cai, D., Wei, W., Yan, Y., Liu, K., Qiao, N., Zhao, X., Jia, J., Wang, W., Kennedy, B., Zhang, K., Cannistraci, C., Zhou, Y., & Han, J. (2020). Three-dimensional facial-image analysis to predict heterogeneity of the human ageing rate and the impact of lifestyle. *Nature Metabolism*, 2(9), 946-957.
- 2019 Chen, X., Xu, C., Hong, S., Xia, X., *Cao, Y.*, McDermott, J., Mu, Y., & Han, J. (2019). Immune cell types and secreted factors contributing to inflammation-to-cancer transition and immune therapy response. *Cell Reports*, 26(7), 1965-1977.
- 2018 Xu, C., Ai, D., Shi, D., Suo, S., Chen, X., Yan, Y., *Cao, Y.*, Zhang, R., Sun, N., Chen, W., McDermott, J., Zhang, S., Zeng, Y. & Han, J. (2018). Accurate drug repositioning through non-tissue-specific core signatures from cancer transcriptomes. *Cell Reports*, 25(2), 523-535.
- Chen, W., Qian, W., Wu, G., Chen, W., Xian, B., Chen, X., Cao, Y., Green, C.D., Zhao, F., Tang, K., & Han, J. (2015). Three-dimensional human facial morphologies as robust aging markers. Cell Research, 25(5), 574-587.
  Huang, Y., Yu, X., Sun, N., Qiao, N., Cao, Y., Boyd-Kirkup, J. D., Shen, Q., & Han, J. (2015). Single-cell-level spatial gene expression in the embryonic neural differentiation niche. Genome Research, 25(4), 570-581.
- 2014 Hong, S., Huang, Y., **Cao, Y.**, Chen, X., & Han, J. D. J. (2014). Approaches to uncovering cancer diagnostic and prognostic molecular signatures. **Molecular & Cellular Oncology**, 1(2), e957981.

#### SOFTWARES

2022-Curr. astroBoy

An automated, Al-assisted platform for large-scale next-generation sequencing data analysis, supporting both in-house and publicly deposited datasets on a single-node server. Deployed at NHLBI/NIH, it has efficiently processed over 300 billion reads from more than 8,000 samples across 600+ analysis jobs since December 2022.

2023-Curr glitter

A custom-built, in-house web application for the analysis and visualization of single-cell RNA-seq data, developed to support collaborative research.

2023-Curr **VINCE** 

> A de novo Python package designed for nucleosome data analysis from singlecell MNase-seg, currently under active development and optimization.

2025 Rvder

> A Python package for epigenome data normalization with internal reference and variable feature detection (manuscript submitting).

2021 cLoops2

> A Python package for full stack analysis tool for chromatin interaction data such as Hi-TrAC, HiChIP and Hi-C (Nucleic Acids Research, 2022).

**TOWN** 2019

> A Python package built on Keras for high-throughput, deep learning-augmented bright-field microscopy to automatically count live C. elegans worms. Developed for in-house use in drug screening to identify compounds that extend worm survival curves.

2017 cLoops

> A Python package for accurate and flexible loops calling tool for 3D genomic data (Bioinformatics, 2020).

### **UNDER REVIEW/REVISE**

2025 1. Guangzhe Ge \*, Yaqiang Cao \*, Danping Wei \*, Mary Attaway, Barbara L Kee, Jinfang Zhu, and Keji Zhao #: Deciphering chromatin priming of T cell lineage by single-cell micrococcal nuclease sequencing

2. Yaqiang Cao \*#, Guangzhe Ge \*, and Keji Zhao #: Ryder: Epigenome Normalization and Variable Feature Identification

#### **AWARDS and HONORS**

2024 NHLBI Director's Award for Innovation

Discovery of a novel regulatory element for IFN-y expression (Immunity, 2023).

#### MENTORING

2024,2023 Hannah Zhu

University of Maryland; NIH Summer Internship Program.

Collaborated on STARR-seq and Hi-TrAC data analysis to study enhancer-

enhancer interactions.

2024 Emily Yu

The Bryn Mawr School; High School Senior Project.

Currently Yale University; mentored on single-cell RNA-seg analysis.

2019-2020 Jonathan Perrie

NIH Postbac Program. Co-mentored in bioinformatics.

Successfully enrolled into the bioinformatics Ph.D. program at UCLA.

#### 2015-18 Yingying Zeng

Partner Institute for Computational Biology; Junior master's student. Co-mentored in bioinformatics and collaborated on research investigating the epigenetic features of MIR retrotransposons. Co-first-authored a publication and successfully enrolled into the Ph.D. program at Nanyang Technological University (Singapore).

#### 2016-18 Zhaoxiong Chen

Partner Institute for Computational Biology; Junior Ph.D. student. Co-mentored in bioinformatics and collaborated on research optimizing the cDBSCAN algorithm and cLoops. Co-first-authored a publication.

#### 2016-18 Daosheng Ai

Partner Institute for Computational Biology; intern. Co-mentored in bioinformatics.

Co-first-authored a publication. Successfully enrolled into the Ph.D. program at Peking University (China).

### **PEER REVIEWER**

Bioinformatics, Communications Biology, BMC Genomics, F1000Research, Frontiers in Immunology, Frontiers in Bioinformatics, Genes, Cells etc.

#### **TALKS**

2025 TriLab Bioinformatics Core, NIDDK, Bethesda, U	JSA
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- 2024 Bioinformatics and Scientific Programming Core, NICHD, Bethesda, USA
- 2024 Invited by Dr. Gordon L. Hager, NCI, Bethesda, USA
- 2023 Invited by Prof. Lei Hou, Boston University, Boston, USA
- 2023 DECODE Seminar Series, NIH, Bethesda, USA
- 2017 Epigenetics Retreat, Ningbo, China

### **CONFERENCE & POSTERS**

- 2025 NIH/FDA Immunology Interest Group Retreat, DC, USA
- 2024 NCI Histone Modifications and Chromatin Structure Symposium, Bethesda, USA
- 2024 Epigenetics & Chromatin, Cold Spring Harbor Laboratory, New York, USA
- 2024 NHLBI Fellow Festival, NIH, Bethesda, USA
- 2024 NIH/FDA Immunology Interest Group Retreat, DC, USA
- 2023 Genome Architecture in Cell Fate and Disease, Ventura, California, USA
- 2023 NHLBI Fellow Festival, Bethesda, USA
- 2022 SCBA DC-Baltimore Chapter Annual Scientific Symposium, University of Maryland, USA
- 2019 4D Nucleosome Annual Meeting, DC, USA
- 2018 Chromatin, Epigenetics & Transcription, Suzhou, China
- 2017 First Chromatin Biology Conference, Shenzhen, China

- 2016 3rd International Symposium on 3D Genomics, Huazhong Agricultural University, Wuhan, China
- 2015 Otto Warburg Summer School, Max Planck Institute for Molecular Genetics, Berlin, Germany

## **TECHNICAL SKILLS**

- Epigenomic, 3D Genome, and single-cell data analysis (Hi-C, ATAC-seq, ChIP-seq, MNase-seq, Hi-TrAC, RNA-seq, Micro-C, 10x single-cell RNA-seq, STARR-seq), with expertise in customized and sophisticated data mining approaches
- Proficient in Python programming and package development
- Experienced CPU/GPU servers setup and maintenance
- Web-application development
- Application and development of deep-learning and machine-learning algorithms

### <u>REFERENCE</u>

Dr. Keji Zhao	Distinguished Investigator, NHLBI/NIH	zhaok@nhlbi.nih.gov
Dr. Paul P. Liu	Senior Investigator, NHGRI/NIH	pliu@nhgri.nih.gov
Dr. Chuan Wu	Senior Investigator, NCI/NIH	chuan.wu@nih.gov
Dr. Jingfang Zhu	Senior Investigator, NIAID/NIH	jfzhu@niaid.nih.gov