## Ziqing (Winston) Zhao: Curriculum Vitae

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#### **RESEARCH INTERESTS**

My research interests intersect optical imaging, molecular biology and biophysical chemistry, with a focus on developing and applying advanced "imagenomic" approaches to quantitatively probe the biophysics of chromatin dynamics at the single-molecule/super-resolution level. In particular, we are interested in understanding how molecular processes that modulate genome organization, accessibility and expression are regulated in *space* and *time*, and how their misregulation underpin human diseases (particularly cancer and aging-associated disorders). More fundamentally, we also aim to illuminate the physico-chemical driving forces (*e.g.* biomolecular phase separation) that govern chromatin dynamics and cell nuclear architecture *in vivo*.

#### **PROFESSIONAL APPOINTMENTS**

#### National University of Singapore (NUS)

Assistant Professor (Presidential Young Professorship), Dept. of Chemistry	2019 - present
Principal Investigator, Centre for BioImaging Sciences (CBIS)	2019 - present
Co-Principal Investigator, Mechanobiology Institute (MBI)	2021 - present
Faculty Member, Integrative Sciences and Engineering Programme (ISEP)	2022 - present
Agency for Science, Technology and Research (A*STAR)	
Postdoctoral Fellow, Genome Institute of Singapore (GIS)	2018 - 2019
Research Fellow, Institute of Molecular and Cell Biology (IMCB)	2015 - 2018
Research Officer, Institute of Bioengineering and Nanotechnology $(\mathrm{IBN})$	2008 - 2009
EDUCATION	
Harvard University, Cambridge, MA	2009 - 2015
Ph.D. in Biophysics	
Thesis: "Probing the Spatio-Temporal Organizations and Dynamics of Gen DNA Replication in the Mammalian Cell Nucleus"	e Expression and
Advisor: X. Sunney Xie, Mallinckrodt Professor of Chemistry and Chemical	Biology
California Institute of Technology (Caltech), Pasadena, CA	2004 - 2008
B.S. (with honors), double major in Chemistry and Biology	
GPA: 4.12/4.0	
Raffles Junior College, Singapore	2002 - 2003
University of Cambridge GCE Advanced Level Examination Certificate	

## AWARDS AND HONORS

Invited to nominate candidates for the Kavli Prize, Norwegian Academy of		
Science and Letters		2023
Invited to nominate candidates for the Nobel Prize in Physiology or Medicine,		
Nobel Committee	2020,	2021
NUS Presidential Young Professorship, National University of Singapore		2019
GIS Super Team Award (Team Member), Genome Institute of Singapore		2018
Selected Delegate of $65^{th}$ Lindau Nobel Laureate Meeting, Lindau, Germany		2015
Certificate of Distinction in Teaching, Harvard University	2011,	2014
Cold Spring Harbor Asia Poster Award, Second Prize, Cold Spring Harbor Asia		2013
Student Research Achievement Award, The Biophysical Society One of the thirteen recipients selected internationally		2013
Dudley R. Herschbach Teaching Award, Harvard University Awarded to the best graduate student teaching fellow in Dept. of Chemistry & Chemical I	Biology	2012
National Science Scholarship (Ph.D.), A*STAR		2009
Richard P. Schuster Memorial Prize, Caltech		2008
Awarded to the best graduating senior in Division of Chemistry & Chemical Engineering		
Upper Class Merit Award (Carnation Scholarship), Caltech	2006,	2007
Chairman's Honors List, A*STAR	2005 -	2007
Phi Tau Phi Scholastic Honor Society of America Scholarship		2007
Summer Undergraduate Research Fellowship (Arthur R. Adams Fellow), Caltech		2007
University College London Scholars Program, Caltech		2006
Summer Undergraduate Research Fellowship (Samuel & Frances Krown Fellow), Ca	ltech	2006
National Science Scholarship (B.S.), A*STAR		2004
World 15 <sup>th</sup> Place, American Invitational Mathematics Examination		2003
Gold Medal and Team Champion, Singapore Chemistry Olympiad		2002
Gold Medals and Team Champion, Singapore Mathematical Olympiad	1999 -	2003

# GRANTS AND FUNDING

Academic Research Fund (AcRF) Tier 2 Grant, Ministry of Education, Singapore MOE-T2EP30222-0010; Role: PI; Amount: S\$1,317,987.00	2023 - 2027
Academic Research Fund (AcRF) Tier 3 Grant, Ministry of Education, Singapore MOET32020-0001; Role: Team PI; Amount: S\$8,424,000.00 (My share: S\$787,356.00	2021 - 2026 0)
Academic Research Fund (AcRF) Tier 1 Grant, Ministry of Education, Singapore Role: PI; Amount: S\$82,500.00	2024 - 2025
Competitive Research Programme (CRP), National Research Foundation, Singapore NRF-CRP25-2020-0001; Role: Team PI; Amount: S\$5,164,900.00 (My share: S\$561,57)	2021 - 2025 340.00)
Academic Research Fund (AcRF) Tier 1 Grant, Ministry of Education, Singapore Role: PI; Amount: S\$250,000.00	2022 - 2024
Young Individual Research Grant, National Medical Research Council, Singapore MOH-000227; Role: PI; Amount: S\$300,000.00	2019 - 2023

NUS Presidential Young Professorship start-up funding, NUS Role: PI; Amount: S\$1,250,000.00

#### PUBLICATIONS

#### Book/book chapter

- **Zhao, Z. W.**, Xie, X. S. Problems and Solutions to Life at the Single-Molecule Level: A Physical Chemistry Perspective (under contract with Oxford University Press).
- Ng, W. S., Sielaff, H., Zhao, Z. W. "Phase Separation in Chromatin-based Intranuclear Processes." In Droplets of Life: Membrane-less Organelles, Biomolecular Condensates, and Biological Liquid-Liquid Phase Separation, ed. Vladimir Uversky, pp. 461–483. London/San Diego/Cambridge/Oxford: Academic Press (an imprint of Elsevier), 2023.

<u>Papers</u> (total citations > 1,850 as of Oct 2024, according to Google Scholar) \*: co-first authorship; <sup>§</sup>: corresponding/co-corresponding authorship

#### At NUS

- Longo, M., Ahmed, S. M., Chen, Y., Tsai, C.-L., Namjoshi, S., Wang, X., Perera, R. L., Arvai, A., Lee, M., Kong, L. R., Engl, W., Ng., W. S., **Zhao, Z. W.**, Venkitaraman, A. R.<sup>§</sup>, Schlacher, K.<sup>§</sup>, Tainer. J. A.<sup>§</sup> BRCA2 C-terminal clamp restructures RAD51 dimers to bind B-DNA for replication fork stability. Under review.
- Sielaff, H.<sup>§</sup>, **Zhao**, **Z. W.**<sup>§</sup> Visualizing, quantifying and mapping chromatin remodelers at work with single-molecule and single-cell imaging. *Intl. J. Biochem. Cell Biol.* **176**, 106667 (2024).
- Engl, W.\*, Kunstar, A.\*, Chen, S.\*, Ng, W. S., Sielaff, H., Zhao, Z. W.<sup>§</sup> Single-molecule imaging of SWI/SNF chromatin remodelers reveals bromodomain-mediated and cancer-mutantsspecific landscape of multi-modal DNA-binding dynamics. *Nature Commun.* 15, 7646 (2024).
- Pu, R.\*, Zhan, Q.\*.§, Peng, X., Liu, S., Guo, X., Liang, L., Qin, X., Zhao, Z. W., Liu, X.§ Superresolution microscopy enabled by high-efficiency surface-migration emission depletion. *Nature Commun.* 13, 6636 (2022).
- Ng, W. S., Sielaff, H., Zhao, Z. W.<sup>§</sup> Phase separation-mediated chromatin organization and dynamics: From imaging-based quantitative characterizations to functional implications. *Int.* J. Mol. Sci. 23, 8039 (2022).
- Sielaff, H.<sup>§</sup>, Basu, S.<sup>§</sup>, Zhao, Z. W.<sup>§</sup> Imaging approaches to unravel chromatin organization and nuclear dynamics. *Front. Mol. Biosci.* 9, 929370 (2022) (editorial commentary).
- Liu, H., Peck, X. Y., Chong, Y. K., Ng, W. S., Engl, W., Raghuvamsi, P. V., Zhao, Z. W., Anand, G. S., Zhou, Y., Sivaraman, J., Xu, Q.<sup>§</sup>, Wong, S.-M.<sup>§</sup> Identification of putative binding interface of PI(3,5)P<sub>2</sub> lipid on rice black-streaked dwarf virus (RBSDV) P10 protein. *Virology* 570, 81–95 (2022).
- Goh, J. J. L.\*, Chou, N.\*, Seow, W. Y., Ha, N., Cheng, C. P. P., Chang, Y.-C., Zhao, Z. W., Chen, K. H.<sup>§</sup> Highly specific multiplexed RNA imaging in tissues with split-FISH. *Nature Methods* 17, 689–693 (2020).

Featured on *GenomeWeb*.

Su, Q. P.\*.§, Zhao, Z. W.\*.§, Meng, L., Ding, M., Zhang, W., Li, Y., Liu, M., Li, R., Gao, Y.-Q., Xie, X. S.<sup>§</sup>, Sun, Y.<sup>§</sup> Superresolution imaging reveals spatiotemporal propagation of human

2019 - 2023

replication foci mediated by CTCF-organized chromatin structures. *Proc. Natl. Acad. Sci.* U.S.A. **117**, 15036–15046 (2020).

Featured on BioArt; EurekAlert!/AAAS; Nanowerk; News Break; NUS News-In Focus; Peking University; Phys.org; Scienmag.

#### Prior to NUS

- Manning, S. A., Dent, L. G., Kondo, S., Zhao, Z. W., Plachta, N., Harvey, K. F.<sup>§</sup> Dynamic fluctuations in subcellular localization of the Hippo pathway effector Yorkie *in vivo. Curr. Biol.* 28, 1651–1660 (2018).
- White, M. D.\*, Zhao, Z. W.\*, Plachta, N.<sup>§</sup> In vivo imaging of single mammalian cells in development and disease. Trends Mol. Med. 24, 278–293 (2018) (cover article).
- Zhao, Z. W.\*, White, M. D.\*, Alverez, Y. D.\*, Zenker, J.\*, Bissiere, S., Plachta, N.<sup>§</sup> Quantifying transcription factor–DNA binding in single cells *in vivo* with photoactivatable fluorescence correlation spectroscopy. *Nature Protoc.* **12**, 1458–1471 (2017).
- Zhao, Z. W., White, M. D., Bissiere, S., Levi, V., Plachta, N.<sup>§</sup> Quantitative imaging of mammalian transcriptional dynamics: From single cells to whole embryos. *BMC Biol.* 14, 115 (2016).
- White, M. D.\*, Angiolini, J. F.\*, Alverez, Y. D.\*, Kaur, G.\*, Zhao, Z. W., Mocskos, E., Bruno, L., Bissiere, S., Levi, V.<sup>§</sup>, Plachta, N.<sup>§</sup> Long-lived binding of Sox2 to DNA predicts cell fate in the four-cell mouse embryo. *Cell* 165, 75–87 (2016) (cover article).

Featured on Cell cover, Medical Xpress; Straits Times; The Scientist.

- Zhao, Z. W., Xie, X. S.<sup>§</sup>, Ge, H.<sup>§</sup> Nonequilibrium relaxation of conformational dynamics facilitates catalytic reaction in an elastic network model of T7 DNA polymerase. J. Phys. Chem. B 120, 2869–2877 (2016).
- Zhao, Z. W.\*, Roy, R.\*, Gebhardt, J. C. M.\*, Suter, D. M.\*, Chapman, A. R., Xie, X. S.<sup>§</sup> Spatial organization of RNA polymerase II inside a mammalian cell nucleus revealed by reflected lightsheet superresolution microscopy. *Proc. Natl. Acad. Sci. U.S.A.* 111, 681–686 (2014).
- Zhao, Z. W., Gebhardt, J. C. M., Suter, D. M., Xie, X. S.<sup>§</sup> Reply to "Convergence of chromatin binding estimates in live cells". *Nature Methods* 10, 692 (2013).
- Gebhardt, J. C. M.\*, Suter, D. M.\*, Roy, R., Zhao, Z. W., Chapman, A. R., Basu, S., Maniatis, T., Xie, X. S.<sup>§</sup> Single-molecule imaging of transcription factor binding to DNA in live mammalian cells. *Nature Methods* 10, 421–426 (2013).
- Ong, S.-M., Zhao, Z., Arooz, T., Zhao, D., Zhang, S., Du, T., Wasser, M., van Noort, D., Yu. H.<sup>§</sup> Engineering a scaffold-free 3D tumor model for *in vitro* drug penetration studies. *Biomaterials* 31, 1180–1190 (2010).
- Zhang, C.\*, Zhao, Z.\*, Rahim, N. A. A., van Noort, D.<sup>§</sup>, Yu. H.<sup>§</sup> Towards a human-on-chip: Culturing multiple cell types on a chip with compartmentalized microenvironments. *Lab Chip* 9, 3185–3192 (2009) (inside cover article).
- Pletneva, E. V., Zhao, Z., Kimura, T., Petrova, K., Gray, H. B.<sup>§</sup>, Winkler. J. R.<sup>§</sup> Probing the cytochrome c' folding landscape. J. Inorg. Biochem. 101, 1768–1775 (2007).

#### PATENT

Chen, K. H., Goh, J. J. L., Chou, S. N., Seow, W. Y., Ha, N., Goh, C, **Zhao**, **Z. W.** Nucleic acid probes. Filed 24 Jun, 2020 (International application number: PCT/SG2020/050353).

AY2019/2020

## TEACHING

#### At NUS

 ${\bf CM3131} \mid {\rm Applications \ of \ Physical \ Chemistry}$ 

Student rating: 4.6/5.0

<u>Selected students feedback</u>:

- "Prof. Winston has shown to be one of the most competent teachers."
- "I could tell that he really cared for the learning and the well-being of the students... He also monitored our learning journey throughout the module well as he adjusted the concepts taught appropriately according to the students' learning abilities."
- "He was very effective in delivering the content of his lectures and ensured that the concepts learnt were easy to understand, despite the difficulty of the concepts taught. He also made multiple links to the previous concepts taught not only in the same module, but also from the previous 2k modules."
- "Prof Winston could explain his topics extremely well and was good as breaking down the problems given in the tutorial in smaller chunks to help us better understand what was actually going on."
- "Always explained the steps to tutorial questions in great detail and clarity, ensuring that we were able to follow his reasoning and would not hesitate to pause or go back to make sure we were still with him."
- "He provides very useful feedback and never fails to address questions."

## CM4236 | Spectroscopy and Imaging in Biophysical Chemistry AY2020/2021 – present (yearly) Student rating: 4.9/5.0 (twice); 4.8/5.0 (twice)

Selected students feedback:

- "He is definitely one of the most inspiring and effective lecturers I have ever met."
- "One of the best professors that I have encountered in my time in NUS. He makes tough content easier to understand and constantly checks in with us during lecture to ensure that the class is following him...Overall, a lecturer that I will remember fondly."
- "One of the few classes I genuinely looked forward to during my uni(versity) years."
- "Slides and module content were top–notch...Really amazing considering it's a 'new' module and ran alone by Prof. Zhao."
- "...I really appreciate his teaching style that has made this module challenging but manageable."
- "Prof. Zhao is really the nicest professor I have encountered in my time in NUS... He accommodates students' requests and (is) able to empathize with us and our workload and adjusts deadlines appropriately when needed, which I'm sure everyone in class appreciates."
- "...Very nice to know that there are teachers who care about us in this way."
- "He is a really caring and considerate prof. He tries his best to help us learn by adapting his homeworks and lectures. Also very encouraging and provides help whenever needed. Plus, he is like running one man show of a new mod(ule) which is pretty darn impressive given how comprehensive and excellent quality his content is."

#### $\mathbf{CM3225} \mid \mathrm{Biomolecules}$

Student rating: 4.4/5.0

Selected students feedback:

- "Explanations are very clear.... let us see the big picture."
- "...This is especially hard to do over Zoom, and we really appreciate the efforts taken to replicate real life lectures as much as possible."

${f MB5104} \mid {f An}$ Integrative Approach to Understand Cell Function	AY2022/2023
Guest Lecturer	

#### Prior to NUS

Chem 161	Statistical Thermodynamics, Harvard University	AY2013/2014
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AY2023/2024 – present (yearly)

Teaching Fellow; Student rating: $4.8/5.0$	
Chem 163   Frontiers in Biophysics, Harvard University	AYs2010 - 2013
Teaching Fellow (taught three times); Student rating: $5.0/5.0$ (twice)	
Chem 24a & b   Introduction to Biophysical Chemistry, Caltech	AYs2006 - 2008
Teaching Assistant (taught twice)	

## MENTORING

# <u>At NUS</u>

<u>Research Fellows</u>	
Kaiyu Jin (Department of Chemistry)	2024 - present
Goran Biukovic (Department of Chemistry)	2022 - present
Wilfried Engl (Department of Chemistry)	2020 - present
Aliz Kunstar (Department of Chemistry)	2020 - present
Hendrik Sielaff (Department of Chemistry)	2020 - present
<u>Research Associates/Assistants/Apprentices</u>	
Hong Zi Mian (Research Assistant, Department of Chemistry)	2023 - 2024
Chen Siyi (Research Associate, Department of Chemistry)	2020 - 2023
Kuo Xuan (Research Apprentice, Department of Chemistry)	2020 - 2021
Nurul Diyana Bte Rosli (Research Apprentice, Department of Chemistry)	2020 - 2021
Ng Woei Shyuan (Research Assistant, Department of Chemistry)	2019 - 2020
<u>Ph.D. students</u>	
Kyna Khoo (Department of Chemistry)	2023 - present
Cui Yinuo (Department of Chemistry; Dean's Graduate Fellow)	2023 - present
Kuo Xuan (Mechanobiology Institute, co-supervised with Tony Kanchanawong)	2021 - present
Ng Woei Shyuan (Department of Chemistry)	2020 - present
<u>Master students</u>	
Xinyue Cui (M.Sc. in Chemical Sciences)	2024 - 2025
Zihui Lu (M.Sc. in Chemical Sciences)	2024 - 2025
Wentao Zeng (M.Sc. in Chemical Sciences)	2024 - 2025
Zhou Songsong (M.Sc. in Chemical Sciences)	2022 - 2023
<u>Undergraduate students</u>	
Clarissa Widyantoro (FYP student, Department of Chemistry)	2023 - 2024
Ryan Seow (FYP student, Department of Chemistry)	2023 - 2024
Hasna Firdaus Aryantha (NUS Amgen Scholars Program, Bandung Institute of	
Technology, Indonesia)	2023
Jasmine Kiley (NUS Amgen Scholars Program, Tulane University, U.S.A.)	2022
Nicole Sim Jiaxuan (FYP student, Department of Chemistry)	2021 - 2022
Serene Fong Siew Min (FYP student, Department of Chemistry)	2019 - 2020
Ph.D./M.Sc. thesis advisory/examination committees	
Yang Rongrong (Ph.D. thesis advisory committee, Department of Chemistry)	2024 - present

Li Hangfeng (Ph.D. thesis advisory committee, Mechanobiology Institute)	2023 - prese	ent
Lau Jun Wei (Ph.D. thesis advisory committee, Department of Chemistry)	2022 - prese	ent
Sui Mingyu (Ph.D. thesis advisory committee, Department of Chemistry)	2022 - preset	ent
Huang Zengxin (Ph.D. thesis advisory committee, Mechanobiology Institute)	2022 – prese	ent
Chen Jiaye (Ph.D. thesis advisory committee, Department of Chemistry)	2021 – prese	ent
Joel Ng (Ph.D. thesis examination panel, Department of Biological Sciences)	20	)24
Daniel Aik (Ph.D. thesis examination panel, Department of Chemistry)	20	)24
Kavitha Rajasekhar (M.Sc. thesis examination panel, Department of Biological Sc	iences) 20	)22
Saradha Venkatachalapathy (Ph.D. thesis examination panel, Mechanobiology Ins	titute) 20	)21
Zhou Yu (Ph.D. thesis examination panel, Department of Physics)	20	)21

## Prior to NUS

Xu Peihao (H3 Research Attachment student, Institute of Molecular and Cell Biolog	gy)	2017
Julie C. Chang (Undergraduate student, Harvard University)	2013 -	2014

# CONFERENCE AND SEMINAR PRESENTATIONS

Nov 2024
June 2024
June 2024
May 2024
Apr 2024
Feb 2024
Aug 2023
Apr 2023
Jan 2023
Oct 2022
op,
Feb 2022
Nov 2021
Jul 2021
Mar 2021
Mar 2021
Feb 2021
$\mathrm{Sep}\ 2020$
Dec 2019
Nov 2019
Feb 2019
Jun 2017

Cold Spring Harbor Laboratory Meeting: Nuclear Organization and Function,	
Cold Spring Harbor, U.S.A.	May 2016
Harvard Medical School Epigenetics Symposium, Boston, U.S.A.	Dec 2014
Cold Spring Harbor Asia Meeting: New Advances in Optical Imaging of Live Cells	
and Organisms, Suzhou, China	Aug 2013
Biophysical Society 57 <sup>th</sup> Annual Meeting, Philadelphia, U.S.A.	Feb 2013
EMBO   EMBL Symposium: The Complex Life of mRNA, Heidelberg, Germany	Oct 2012
4 <sup>th</sup> Combined Scientific Meeting of the Life Sciences, Singapore	Jan 2003

# JOURNAL/GRANT REVIEWING/EDITING

Review Editor, Frontiers in Chemical Biology	2022 - present
Invited Topic Editor, Frontiers in Molecular Biosciences	2020 - 2022
Ad hoc reviewer for: Analytical Chemistry, Biophysical Journal, Cell, Gen	nes, Journal of Physical
Chemistry, Micron, Nano Letters, Nanoscale, Nature, Nature Commun	ications, NPJ Imaging,
Science Advances, Trends in Genetics	
Invited grant reviewer for: Austrian Science Fund (FWF) (declined)	

#### ADMINISTRATIVE SERVICES

Sub-director, NUS Suzhou Research Institute "3+1+1" Program in Chemistry and TJU-NUS	
Double-Master's Degree Program in Chemistry, Department of Chemistry	2023 - present
Member, Committee on M.Sc. in Chemical Sciences, Department of Chemistry	2022 - present
Member, Committee on Graduate Education, Department of Chemistry	2021 - present
Member, Committee on Student Life, Department of Chemistry	2020 - present
Member, Committee on College of Humanities & Sciences, Department of Chemistry	2020 - 2021
Member, University Research Committee Expert Panel, NUS	2019

## PROFESSIONAL SERVICES/OUTREACH ACTIVITIES

Judge, Science Mentorship Programme (SMP), Faculty of Science, NUS	2023
Judge, 26 <sup>th</sup> Chemistry • Communication Challenge: Chemistry for Singapore 2030	
and Beyond, NUS Chemistry	2021
Speaker, Advancing the Frontiers of Science and Technology with Chemistry	
E-outreach, NUS Chemistry	2021 (twice)
Judge, A*STAR Talent Search (ATS), A*STAR	2020
Speaker, NUS-ACS Student Chapter Graduate Studies Talk	2020
Selection panelist for nominees to $70^{\text{th}}$ Lindau Nobel Laureate Meeting, National	
Research Foundation, Singapore	2019
Poster judge, 2 <sup>nd</sup> Chemistry National Meeting Singapore (ChnmSG 2019), Singapore	e
National Institute of Chemistry	2019