

Shuaihua Gao, Ph.D.
Assistant Professor
Department of Chemical and Biomolecular Engineering
Tulane University

Office: Boggs Building 329 Lab: Boggs Building 343 Email: sgao11@tulane.edu

## **EDUCATION/PROFESSIONAL EXPERIENCE**

07/2023-present Assistant professor, Chemical and Biomolecular Engineering Department, Tulane University

07/2018 — 06/2023 Postdoctoral researcher at University of California, Berkeley

Department of Chemistry and QB3 Institute

New Orleans, LA, 70118.

Advisor: Prof. Judith P. Klinman

Project: Using biophysical probes of temperature dependent hydrogen deuterium exchange (TD-HDX) and Stokes shift fluorescence spectroscopy to investigate enzymatic thermal activation pathways and solvation dynamics for active site chemistry for a TIM barrel hydrolase adenosine deaminase. (Training in enzymology; protein dynamics; structural

biology, and biocatalysis)

11/2016 — 03/2018 PhD student researcher at University of California, Berkeley

Department of Chemistry and QB3 Institute

Supervisor: Prof. Judith P. Klinman

Project: Tool development in TD-HDX to map activity-related protein motions in murine

adenosine deaminase

09/2013—06/2018 Ph.D. in Chemical Engineering and Technology, Beijing University of Chemical

Technology, College of Life Science and Technology

Supervisor: Prof. Guojun Zheng

Project: Identification, protein engineering and origin of enantioselectivity of a  $(+)-\gamma$ -lactamase from Microbacterium hydrocarbonoxydans. (Training in biocatalysis, protein

engineering, structural biology, and enzyme discovery)

09/2009 — 06/2013 B.S. in Pharmaceutical Engineering, Beijing University of Chemical Technology

College of Life Science and Technology

Supervisor: Prof. Guojun Zheng

Project: Efficient synthesis of the intermediate of abacavir and carbovir using a novel lactamase as a catalyst. (Training in enzymology, pharmacology, and enzyme catalysis)

## RESEARCH EXPERIENCE

Postdoctoral Fellow, University of California at Berkeley (PI: Judith P. Klinman)

- Protein and structural dynamics studies using temperature dependent hydrogen deuterium exchange couple to mass spectrometry (TDHDX-MS)
- Biophysical basis of enzymatic reaction cycle via TDHDX-MS with bound ground and transition state analogs
- Heat dissipation from solvent to enzyme active site studies using temperature dependent stokes shift fluorescence spectroscopy

PhD student, Beijing University of chemical Technology (PI: Guojun Zheng)

- Enzyme discovery by environmental screening and in silico genome mining methods
- · High-throughput colorimetric screening method development for enzyme evolution and engineering
- Structural biology to investigate structural insight into the molecular mechanism of substrate recognition and origin of enantioselectivity
- Protein engineering in enhancing enantioselectivity and thermostability of enzymes by combined method of directed evolution and semi-rational design

### **AWARDS**

2013-2018	Outstanding graduate award, Beijing University of Chemical Technology
2016—2017	Overseas study program fellowship, China Scholarship Council
2013-2014	2 <sup>nd</sup> place prize in bio-separation design award
2011-2012	National Endeavor Fellowship for undergraduates, China
2012-2013	1 <sup>st</sup> tier undergraduate scholarship, Beijing University of Chemical Technology
2011-2012	1 <sup>st</sup> tier undergraduate scholarship, Beijing University of Chemical Technology
2010 — 2011	1st tier undergraduate scholarship, Beijing University of Chemical Technology
2009-2010	2 <sup>nd</sup> tier undergraduate scholarship, Beijing University of Chemical Technology

# **TEACHING EXPERIENCE**

_	
2017—present	Research mentor to 5 undergraduate students in Klinman lab, UC Berkeley
2018—2019	Teacher, Splash at Berkeley, Class: How enzymes achieve enormous rate
	accelerations and how to rationally engineer enzymes
2014—2018	Research mentor to 2 undergraduate and 3 graduate students, BUCT
2016	Teaching assistant, class of Molecular Biology
2015	Teaching assistant, class of Enzymology and Biocatalysis
2010-2013	Teacher in the teaching agency of Excel at Learning

# PUBLICATIONS (\*denotes corresponding author)

Google scholar link for: my publications

- (1) **Shuaihua Gao**, Xin Ting Wu, Wenju Zhang, Tyre Richardson, Samuel L. Barrow, Christian A. Thompson-Kucera, Anthony T. Iavarone, and Judith P. Klinman. Temporal Resolution of Activity-Related Nanosecond Solvation Dynamics in the TIM Barrel Enzyme Murine Adenosine Deaminase. ACS Catalysis, 2024, 14, 4554–4567.
- (2) <u>Gao, S</u> and Klinman, J. P., Functional roles of enzyme dynamics in accelerating active site chemistry: emerging techniques and changing concepts. Current Opinion in Structural Biology, 2022, 75, 102434. (*Invited Review paper*).
- (3) <u>Gao, S.;</u> Zhang, W.; Barrow, S. L.; Iavarone, A. T.; Klinman, J. P., Temperature-dependent hydrogen deuterium exchange shows impact of analog binding on adenosine deaminase flexibility but not embedded thermal networks. Journal of Biological Chemistry, 2022, 298(9), 102350.
- (4) <u>Gao, S.;</u> Thompson, E. J.; Barrow, S. L.; Zhang, W.; Iavarone, A. T.; Klinman, J. P., Hydrogen–Deuterium Exchange within adenosine deaminase, a TIM barrel hydrolase, identifies networks for thermal activation of catalysis. Journal of the American Chemical Society 2020, 142 (47), 19936-19949.
- (5) Zhang, J.; Balsbaugh, J. L.; <u>Gao, S.;</u> Ahn, N. G.; Klinman, J. P., Hydrogen deuterium exchange defines catalytically linked regions of protein flexibility in the catechol O-methyltransferase reaction. Proceedings of the National Academy of Sciences 2020, 117 (20), 10797-10805.
- (6) Gao, S.; Lu, Y.; Li, Y.; Huang, R.; Zheng, G., Enhancement in the catalytic activity of Sulfolobus solfataricus
- P2 (+)-γ-lactamase by semi-rational design with the aid of a newly established high-throughput screening method. Applied microbiology and biotechnology 2019, 103 (1), 251-263.
- (7) **Gao, S.;** Zhu, S.; Huang, R.; Li, H.; Wang, H.; Zheng, G., Engineering the Enantioselectivity and Thermostability of a (+)-γ-Lactamase from *Microbacterium hydrocarbonoxydans* for Kinetic Resolution of Vince Lactam (2-Azabicyclo [2.2. 1] hept-5-en-3-one). Applied and environmental microbiology 2018, 84 (1).
- (8) <u>Gao, S.;</u> Zhou, Y.; Zhang, W.; Wang, W.; Yu, Y.; Mu, Y.; Wang, H.; Gong, X.; Zheng, G.; Feng, Y., Structural insights into the γ-lactamase activity and substrate enantioselectivity of an isochorismatase-like hydrolase from *Microbacterium hydrocarbonoxydans*. Scientific reports 2017, 7, 44542.

- (9) <u>Gao, S.</u>; Huang, R.; Zhu, S.; Li, H.; Zheng, G., Identification and characterization of a novel (+)-γ-lactamase from *Microbacterium hydrocarbonoxydans*. Applied microbiology and biotechnology 2016, 100 (22), 9543-9553.
- (10) <u>Gao, S.;</u> Su, Y.; Zhao, L.; Li, G.; Zheng, G., Characterization of a (*R*)-selective amine transaminase from *Fusarium* oxysporum. Process Biochemistry 2017, 63, 130-136.
- (11) <u>Gao, S.;</u> Zhu, S.; Huang, R.; Lu, Y.; Zheng, G., Efficient synthesis of the intermediate of abacavir and carbovir using a novel (+)-y-lactamase as a catalyst. Bioorganic & Medicinal Chemistry Letters 2015, 25 (18), 3878-3881.
- (12) Chen, Y.; Gao, F.; Zheng, G.; <u>Gao, S.,</u>\* Enantioselective synthesis of a chiral intermediate of himbacine analogs by Burkholderia cepacia lipase A. Biotechnology Letters 2020, 42 (12), 2643-2651.
- (13) Chen, Y.; Zhang, X.; Zheng, G.; <u>Gao, S.,</u>\* Preparation of the enantiomerically enriched precursor of lamivudine (3TC<sup>™</sup>) via asymmetric catalysis mediated by *Klebsiella oxytoca*. Process Biochemistry 2019, 81, 77-84.
- (14) Li, H.; <u>Gao, S.;</u> Qiu, Y.; Liang, C.; Zhu, S.; Zheng, G., Genome mining integrating semi-rational protein engineering and nanoreactor design: roadmap for a robust biocatalyst for industrial resolution of Vince lactam. Applied Microbiology and Biotechnology 2020, 104 (3), 1109-1123.
- (15) Shen, X.; Zhou, D.; Lin, Y.; Wang, J.; Gao, S.; Kandavelu, P.; Zhang, H.; Zhang, R.; Wang, B.-C.; Rose, J., Structural Insights into Catalytic Versatility of the Flavin-dependent Hydroxylase (HpaB) from Escherichia coli. Scientific reports 2019, 9 (1), 7087.
- (16) Su, Y.; <u>Gao, S.;</u> Li, H.; Zheng, G., Enantioselective resolution of  $\gamma$ -lactam utilizing a novel (+)- $\gamma$ -lactamase from *Bacillus* thuringiensis. Process Biochemistry 2018, 72, 96-104.
- (17) Zhu, S.; Huang, R.; <u>Gao, S.;</u> Li, X.; Zheng, G., Discovery and characterization of a second extremely thermostable (+)-γ-lactamase from *Sulfolobus solfataricus* P2. Journal of bioscience and bioengineering 2016, 121 (5), 484-490.
- (18) Ren, L.; Zhu, S.; Shi, Y.; <u>Gao, S.;</u> Zheng, G., Enantioselective resolution of  $\gamma$ -lactam by a novel thermostable type II (+)- $\gamma$ -lactamase from the hyperthermophilic archaeon *Aeropyrum pernix*. Applied biochemistry and biotechnology 2015, 176 (1), 170-184.
- (19) Zhu, S.; Gong, C.; Song, D.; <u>Gao, S.</u>; Zheng, G., Discovery of a novel (+)-γ-lactamase from *Bradyrhizobium japonicum* USDA 6 by rational genome mining. Applied and environmental microbiology 2012, 78 (20), 7492-7495.

## **PATENTS**

Zheng, G., <u>Gao S.</u>, Huang, R., Zhu S., Li H., Applications of a *Microbacterium hydrocarbonoxydans* derived open reading frame in encoding (+)-γ-lactamase and the preparation of optically pure Vince-lactam. 201510051783.0, 2017, China.

### **ORAL PRESENTATIONS**

- 1. "Biophysical tools for the detection of protein dynamics across a variety of time scales", Mainland and Taiwan biotechnology symposium, 2016, Beijing.
- 2. "Mapping Activity-Related Protein Motions in Murine Adenosine Deaminase using Time-, Temperature-, and Mutation-Dependent Hydrogen-Deuterium Exchange Mass Spectrometry (HDX-MS)", Gordon Research, Seminar, Proteins, Holderness, NH, US, June 16-21, 2019.
- 3. "Hydrogen-Deuterium Exchange within Adenosine Deaminase, a TIM Barrel Hydrolase, Identifies Networks for Thermal Activation of Catalysis", ACS Fall 2020 Meeting & Expo, Biological Chemistry Division.
- 4. "Hydrogen-Deuterium Exchange within Adenosine Deaminase, a TIM Barrel Hydrolase, Identifies 5. Networks for Thermal Activation of Catalysis", ChemistLive, 2020.
- 5. "Identification of Thermal Networks for Catalysis in murine adenosine deaminase", Rutgers University symposium-Metalloproteins at the Crossroads of Design and Nature, January 27, 2021.
- 6. "Hydrogen deuterium exchange within adenosine deaminase provides long range protein networks and structural motifs for the thermal activation of active site chemistry", ACS Spring 2021 Meeting & Expo, Biological Chemistry Division. April 5-30, 2021.

### At Tulane:

- 7. "Identification, structural insights, and protein evolution of an enantioselective lactamase for antiviral drug synthesis", Tulane Center of Biomedical Informatics and Genomics, School of Medicine, Tulane University. Sep. 6, 2023.
- 8. "Protein engineering and structural insights into the lactamase activity and substrate enantioselectivity of a hydrolase from *Microbacterium hydrocarbonoxydans*", Department of Biochemistry and Molecular Biology, School of Medicine, Tulane University. Jan 22, 2024.
- 9. "Identification, structural insights, and protein evolution of an enantioselective lactamase for antiviral drug synthesis", ACS Spring 2024 Meeting, Biochemical Technology Division. March 17-21, 2024. New Orleans, Louisiana, USA.

## **POSTERS**

- 1. "Mapping Activity-Related Protein Motions in Murine Adenosine Deaminase using Time-, Temperature-, and Mutation-Dependent Hydrogen-Deuterium Exchange Mass Spectrometry (HDX-MS)" Gordon Research Conference, Proteins, Holderness, NH, US, June 15-16, 2019.
- 2. "Identification, Characterization, Protein Engineering and Structural Insights into the gamma-Lactamase Activity and Substrate Enantioselectivity of a Hydrolase from *Microbacterium Hydrocarbonoxydans*", ACS Fall 2020 Meeting & Expo Biochemical Technology Division.
- 3. "Discovery and engineering of novel enzymes for the preparation of antiviral drugs abacavir and carbovir", ACS Spring 2021 Meeting & Expo, Division of Medicinal Chemistry. April 5-30, 2021.
- 4. "Temperature Dependent Hydrogen Deuterium Exchange Within Adenosine Deaminase in Complex with Substrate and Transition State Analogs: Ligands Alter Protein Flexibility but Not Embedded Thermal Networks" Gordon Research Conference, Biocatalysis, Southern New Hampshire University, NH, US, July 10-15, 2022.