

Francesca-Zhoufan Li



AI for Science & Engineering

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US Citizen, Shanghainese American

Goal

To advance AI for science with a protein engineering focus, a computational skill set grounded in experimental experience, and an interdisciplinary adaptable collaborative mindset

Skills & Tools

Python Matlab LaTeX Bash Git

HPC PyTorch CUDA Scikit-Learn

Pandas Numpy SciPy Biopython

BLAST ClustalW ESM CARP

Data analysis Machine learning

Biotechnology Protein engineering

Analytical chemistry Collaboration

Research Proposal & paper writing

Leadership Workshop organization

Teaching Mentoring Outreach

Selected Talks & Posters

- ICML, 2024
- ML Protein Engineering Seminar, 2024
- SynBioBeta, 2023
- Caltech Bioscience Futures Day, 2023
- Seagate-Minnesota AI/ML Virtual Distinguished Speaker Series, 2023
- Google Research invited talk, 2022

Other Experiences

- Co-organized the GEM Bio workshop at ICLR 2024 with over 108 paper submissions and 300 attendances
- Code with Young Legends: led intro to coding workshop
- i-STEM: mentored under-invested high school students on research projects
- Bioengineering Honor Society: mentored high school bioengineering research competitions, with one of the teams winning a 2nd place
- Biotech Connection Los Angeles: to grow the local biotech landscape
- Bioengineering Scholars Program: to challenge who can do STEM

Education

- 09/20-Present **Ph.D. in Bioengineering, GPA: 4.0** California Institute of Technology
• NSF Graduate Research Fellowships Program • Amazon AI4Science Fellowship • Biotechnology Leadership Training Program
- 08/15-05/19 **B.S. in Bioengineering, GPA: 3.96** University of California, Berkeley
B.S. in Chemical Biology
• Highest Honors • Jack & Birthe Kirsch Prize • Tau Beta Pi Scholarship
• John Gorton Davis Scholarship • T. Dale Stewart Scholarship
• Genentech Outstanding Student Award Runner-Up

Industry & Academic Experience

- 06/22-09/22 **BioML Research Intern** Microsoft Research
• Performed a systematic analysis of protein language model transfer learning via 370 experiments across downstream tasks, architectures, model sizes, model depths, and pretraining time (ICML 2024)
- 01/21-Present **Machine Learning for Proteins** Arnold Lab & Yue Group, Caltech
• Developing zero-shot predictors for non-native enzyme activities
• Systematically analyzed multiple machine learning-assisted directed evolution strategies, including active learning and focused training using six distinct zero-shot predictors, across 16 diverse protein fitness landscapes
• Facilitated the rapid generation of sequence-function data and tool development for constructing protein mutant libraries
• Led and facilitated 3 grant writing and cross-group collaborations
- 09/20-01/21 **Extremophile Genetic Component Discovery** Murray Lab, Caltech
• Constructed an RNA-seq analysis pipeline in R to discover novel genetic circuit components in non-canonical cell-free extracts
• Delivered results to groups at Caltech, the U.S. Army Chemical Biological Center, and the Imperial College London
- 06/20-08/20 **RNA-Seq Sample Preparation Pipeline Optimization** Zymergen
• Developed a Python package to design DNA oligos for RNaseH-based ribosomal RNA depletion for 8 strains in 7 programs
• Wrote R scripts to quality control and pre-process RNA extraction data from industry-standard electrophoresis instruments
• Delivered talks, collaborated across and outside of the company
- 06/19-05/20 **Bioinformatics Tool Development** Koide Lab, NYU Langone Health
• Developed Matlab software for SARS-CoV-2 mutation analysis from GISAID database covering 25k global sequence entries
• Wrote easy-to-use Matlab scripts to identify monobody and antibody complementary-determined region mutations for protein engineering
• Automated chromatogram visualization with user-chosen features
- 05/18-07/18 **Cell-Free Platform Streamline** Tierra Biosciences, QB3 Program
• Optimized non-standard protein production in cell-free expression systems with Design Of Experiments methodology
- 01/16-05/19 **Independent Bioengineering Researcher** Dueber Lab, UC Berkeley
• Automated time-course betaxanthin production analysis in Matlab
• Engineered yeast to increase benzyloquinoline alkaloids yield
• Improved beta-glucosidase stability & activity in a basic solution for indigo bio-production in *E. coli* via error-prone PCR libraries
• Honor thesis: A "Microbial Factory" Toolkit: Yeast Spheroplast Transformation Method Development for CRISPR-Cas9 Multiplexing

Featured Publications

- 2024 **Li F-Z, et al.** Evaluation of Machine Learning-Assisted Directed Evolution Across Diverse Combinatorial Landscapes. *bioRxiv*.
- 2024 **Li F-Z, et al.** Feature Reuse and Scaling: Understanding Transfer Learning with Protein Language Models. *PMLR*. **235**, 27351-27375.
- 2024 Yang J, **Li F-Z**, & Arnold FH. Opportunities and Challenges for Machine Learning-Assisted Enzyme Engineering. *ACS Cent. Sci.* **10**, 226-241.
- 2024 Long Y, Mora A, Gürsoy E, Johnston KE, **Li F-Z**, & Arnold FH. LevSeq: Rapid Generation of Sequence-Function Data for Directed Evolution and Machine Learning. *bioRxiv*.
- 2023 Yang J, Ducharme J, Johnston KE, **Li F-Z**, et al. DeCOIL: Optimization of Degenerate Codon Libraries for Machine Learning-Assisted Protein Engineering. *ACS Synth. Biol.* **12**, 2444-2454.