# **BRIAN HIE**

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# **CURRENT POSITIONS**

Stanford University,	Palo	Alto,	CA
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# Assistant Professor of Chemical Engineering

#### The Dieter Schwarz Foundation SDS Faculty Fellow, Stanford Data Science 2024-Present

- Faculty Fellow, Sarafan ChEM-H
- Affiliate, Institute for Human Centered Artificial Intelligence (HAI)
- Member, Stanford Bio-X

Arc Institute, Palo Alto, CA

*Innovation Investigator* 2024-Present

# **EDUCATION**

Massachusetts Institute of Technology, Cambridge, MA

Electrical Engineering and Computer Science, Doctor of Philosophy 2017-2021

Electrical Engineering and Computer Science, Master of Science

• Areas of concentration: Computational biology, machine learning, statistics.

Stanford University, Palo Alto, CA

Computer Science, Bachelor of Science with Honors and Distinction 2012-2016

English Literature, Academic Minor

• Areas of concentration: Computational biology, computer systems, machine learning.

### PROFESSIONAL EXPERIENCE

# Stanford University School of Medicine, Palo Alto, CA

#### Stanford Science Fellow 2021-2023

• Machine learning at the host-pathogen interface.

Meta Platforms, Inc., Menlo Park, CA

#### Visiting Researcher, Meta AI FAIR 2022-2023

• Machine learning for protein biology.

Massachusetts Institute of Technology, Cambridge, MA

#### Graduate Researcher, Computer Science and Artificial Intelligence Laboratory 2017-2021

- Neural language modeling of viral evolution.
- Machine learning for biological discovery and design under uncertainty.
- Efficient geometric algorithms for single-cell biology.
- Cryptographically secure neural network training.

Google LLC, Mountain View, CA

#### Artificial Intelligence/Machine Learning Resident, X – The Moonshot Factory 2019

• Machine learning for early-pipeline moonshots.

Illumina, Inc., San Diego, CA

Machine Learning Intern, Bioinformatics 2018

• Statistical signal processing for genomics-based health monitoring.

Salesforce.com, Inc., San Francisco, CA

# Software Engineer, Cloud Infrastructure

2016-2017

• Robust performance monitoring of globally distributed core application infrastructure.

Stanford University, Palo Alto, CA

#### Undergraduate Researcher, Biology

2013-2016

• Statistics and machine learning for computational genomics.

Stanford University, Palo Alto, CA

#### Undergraduate Researcher, Digital Humanities, Stanford Humanities Center

2016

• Graph-theoretic analysis of the social network of early modern authors and publishers.

Microsoft Corporation, Redmond, WA

# Software Engineering Intern, Azure Compute and Microsoft Research

2015

· Distributed scheduling algorithms and their impact on data center utilization and availability.

Synaptics, Inc., San Jose, CA

#### Systems Architecture/Algorithms Intern

2014

• Algorithm design and implementation for embedded touchscreen firmware.

#### **PUBLICATIONS**

\*Equal contribution. †Co-corresponding author.

#### Journal articles

E. Nguyen\*, M. Poli\*, M. Durrant\*, B. Kang\*, D. Katrekar\*, D. Li\*, L. Bartie, A. Thomas, S.

King, G. Brixi, J. Sullivan, M. Ng, A. Lewis, A. Lou, S. Ermon, S. Baccus, T. Hernandez-

Boussard, C. Ré, P. Hsu<sup>†</sup>, and **B. Hie**<sup>†</sup>.

"Sequence modeling and design from molecular to genome scale with Evo."

Science, 386:6723 (cover article, featured with a Perspective by Theodoris).

2024

V. Shanker, T. Bruun, **B. Hie**<sup>†</sup>, and P. Kim<sup>†</sup>.

"Unsupervised evolution of protein and antibody complexes with a structure-informed language model."

Science, 385:6704. 2024

**B.** Hie<sup>†</sup>, S. Kim, T. Rando, B. Bryson<sup>†</sup>, and B. Berger<sup>†</sup>.

"Scanorama: integrating large and diverse single-cell transcriptomic datasets."

Nature Protocols, DOI: 10.1038/s41596-024-00991-3.

2024

S. Hendrix, Y. Mreyoud, M. McNelan, A. Smirnov, S. Chavez, B. Hie, M. Chamberland, T.

Bradstreet, A. Webber, D. Kreamalmeyer, R. Taneja, B. Bryson, B. Edelson, and C. Stallings.

"BHLHE40 regulates myeloid cell polarization through IL-10-dependent and -independent mechanisms."

Journal of Immunology, 212:11.

2024

A. Winnifrith, C. Outeiral<sup>†</sup>, and **B. Hie**<sup>†</sup>.

"Generative artificial intelligence for de novo protein design."

B. Hie, B. Bryson, and B. Berger.

B. Hie<sup>†</sup>, D. Xu, V. Shanker, T. Bruun, P. Weidenbacher, S. Tang, W. Wu., and J. Pak, and P. Kim<sup>†</sup>. "Efficient evolution of human antibodies from general protein language models and sequence information alone." Nature Biotechnology, 42:275-283 (featured with a News and Views by Outeiral and 2023 Deane). Z. Lin\*, H. Akin\*, R. Rao\*, B. Hie\*, Z. Zhu, W. Lu, N. Smetanin, R. Verkuil, O. Kabeli, Y. Shmueli, A. dos Santos Costa, M. Fazel-Zarandi, T. Sercu, S. Candido, and A. Rives. "Evolutionary-scale prediction of atomic level protein structure with a language model." Science, 379:6637. 2023 **B.** Hie<sup>†</sup>, K. Yang, and P. Kim<sup>†</sup>. "Evolutionary velocity with protein language models predicts evolutionary dynamics of diverse proteins." 2022 Cell Systems, 13:4 (featured with a Preview by Sandhu et al.). M. Maher, I. Bartha, S. Weaver, J. di Iulio, E. Ferri, L. Soriaga, F. Lempp, B. Hie, B. Bryson, B. Berger, D. Robertson, G. Snell, D. Corti, H. Virgin, S. Kosakovsky Pond, and A. Telenti. "Predicting the mutational drivers of future SARS-CoV-2 variants of concern." Science Translational Medicine, 14:633. 2022 B. Hie and K. Yang. "Adaptive machine learning for protein engineering." Current Opinion in Structural Biology, 72: February 2022. 2022 R. Singh\*, **B. Hie**\*, A. Narayan, and B. Berger. "Schema: Metric learning enables interpretable synthesis of heterogeneous single-cell modalities." Genome Biology, 22:131. 2021 B. Hie, E. Zhong, B. Berger, and B. Bryson. "Learning the language of viral evolution and escape." Science, 371:6526 (featured with a Perspective by Kim and Przytycka). 2021 **B.** Hie, B. Bryson, and B. Berger. "Leveraging uncertainty in machine learning accelerates biological discovery and design." 2020 Cell Systems, 11:5. B. Hie\*, J. Peters\*, S. Nyquist\*, A. Shalek, B. Berger, and B. Bryson. "Computational methods for single-cell RNA sequencing." 2020 Annual Review of Biomedical Data Science, 3:1. B. Hie\*, H. Cho\*, B. DeMeo, B. Bryson, and B. Berger. "Geometric sketching compactly summarizes the single-cell transcriptomic landscape." Cell Systems, 8:6 (cover article). 2019

"Efficient integration of heterogeneous single-cell transcriptomes using Scanorama." Nature Biotechnology, 37:6.	2019		
A. Tehranchi, <b>B. Hie</b> , M. Dacre, I. Kaplow, K. Pettie, P. Combs, and H. Fraser. "Fine-mapping cis-regulatory variants in diverse human populations." eLife, 8:e39595.	2019		
<b>B. Hie*</b> , H. Cho*, and B. Berger.  "Realizing private and practical pharmacological collaboration."  Science, 362:6417.	2018		
A. Tehranchi, M. Myrthil, T. Martin, <b>B. Hie</b> , D. Golan, and H. Fraser. "Pooled ChIP-seq links variation in transcription factor binding to complex disease risk." Cell, 165:3.	2016		
Conference papers			
T. Widatalla*, R. Shuai*, <b>B. Hie</b> <sup>†</sup> , and P. Huang <sup>†</sup> .  "Sidechain conditioning and modeling for full-atom protein sequence design with FAMPNN."			
International Conference on Machine Learning (ICML).	2025		
M. Poli, A. Thomas, E. Nguyen, S. Massaroli, P. Ponnusamy, B. Deiseroth, K. Kersting, T. Suzuki, <b>B. Hie</b> , S. Ermon, C. Ré, and C. Zhang. "Mechanistic design and scaling of hybrid architectures."  International Conference on Machine Learning (ICML).	2024		
C. Hsu, R. Verkuil, J. Liu, Z. Lin, <b>B. Hie</b> , T. Sercu, A. Lerer, and A. Rives.			
"Learning inverse folding from millions of predicted structures."  International Conference on Machine Learning (ICML).	2022		
B. Hie, E. Zhong, B. Bryson, and B. Berger.  "Learning mutational semantics."  Neural Information Processing Systems (NeurIPS)	2020		
Neural Information Processing Systems (NeurIPS).	2020		
<b>B. Hie</b> , H. Cho, B. DeMeo, B. Bryson, and B. Berger.  "Geometric sketching of single-cell data preserves transcriptional structure."  Research in Computational Molecular Biology (RECOMB).	2019		
Book chapters			
K. Johnston*, C. Fannjiang*, B. Wittmann*, <b>B. Hie</b> *, K. Yang*, and Z. Wu*.  "Machine learning for protein engineering."  Machine Learning in Molecular Sciences, Challenges and Advances in Computational Chemistry and Physics. Springer Nature.	2023		
Preprints			
G. Brixi*, M. Durrant*, J. Ku*, M. Poli*, G. Brockman, D. Chang, G. Gonzalez, S. King, D. L. A. Merchant, M. Naghipourfar, E. Nguyen, C. Ricci-Tam, D. Romero, G. Sun, A.	i,		

Taghibakshi, A. Vorontsov, B. Yang, et al., D. Burke, H. Goodarzi, P. Hsu<sup>†</sup>, and **B. Hie**<sup>†</sup>.

"Genome modeling and design across all domains of life with Evo 2."	
bioRxiv, DOI: 10.1101/2025.02.18.638918.	2025
J. Ku, E. Nguyen, D. Romero, G. Brixi, B. Yang, A. Vorontsov, A. Taghibakhshi, A. Lu, D. Burke, G. Brockman, S. Massaroli, C. Ré, P. Hsu, <b>B. Hie</b> , S. Ermon, and M. Poli. "Systems and algorithms for convolutional multi-hybrid language models at scale." arXiv, DOI: 10.48550/arXiv.2503.01868.	2025
A. Merchant, S. King, E. Nguyen, and <b>B. Hie</b> . "Semantic mining of functional de novo genes from a genomic language model." bioRxiv, DOI: 10.1101/2024.12.17.628962.	2024
<b>B. Hie*</b> , S. Candido*, Z. Lin, O. Kabeli, R. Rao, N. Smetanin, T. Sercu, and A. Rives. "A high-level programming language for generative protein design." bioRxiv, DOI: 10.1101/2022.12.21.521526.	2022
C. Itoh, C. Gunnarson, G. Babunovic, A. Nibasumba, Ngomu A., M. Wadsworth III, T. Hughes II, S. Solomon, <b>B. Hie</b> , B. Berger, A. Shalek, S. Fortune, and B. Bryson. "GM-CSF differentiation of human monocytes stabilizes macrophage state via oxidative signaling."	
bioRxiv, DOI: 10.1101/2020.09.29.318352.	2020

# **SOFTWARE**

Evo 2, Primary Developer

https://github.com/arcinstitute/evo2

Evo, Primary Developer

https://github.com/evo-design/evo

Scanorama, Primary Developer

https://github.com/brianhie/scanorama

Geosketch, Primary Developer

https://github.com/brianhie/geosketch

Evolocity, Primary Developer

https://github.com/brianhie/evolocity

esm, Contributor

https://github.com/facebookresearch/esm

scverse, Contributor

https://github.com/scverse

# **TEACHING**

Stanford University, Palo Alto, CA

Instructor, Data Science for Computational Molecular Biology (DATASCI 194B/294B)

2025

Machine learning and its applications to biology. Topics include protein language modeling, protein structure prediction, molecular representation learning, Bayesian optimization, single-cell genomics, and genomic language modeling.

# Instructor, Data Science and Machine Learning Approaches in Chemical and Materials Engineering (CHEMENG 177/277, MATSCI 166/176)

2025

Applications of data science and machine learning approaches to modern problems in chemical, materials, and biological engineering.

# Massachusetts Institute of Technology, Cambridge, MA

#### Teaching Assistant, Algorithms for Inference (6.438)

2019

Graduate-level course on statistical inference with probabilistic graphical models. Responsible for preparing exams/assignments, leading discussion sections, and holding office hours.

#### ACADEMIC SERVICE

#### Mentorship

•	Stanford University, Doctoral thesis research	2021-Present
•	Stanford University, Undergraduate thesis research	2024-Present
•	Stanford University, ADVANCE Undergraduate Institute	2025-Present
•	University of Toronto, iGEM competition team	2021
•	Massachusetts Institute of Technology, Masters of Engineering thesis research	2020-2021

#### **Public Engagement and Science Communication**

Press interviews for Freethink, Ideas Roadshow, IEEE Spectrum, Inverse, Al Jazeera, MIT Technology Review, NSF The Discovery Files, Nature News, Nautilus, Nikkei, Quanta Magazine, Science News, Swiss Radio, Tagesspiegel, Wall Street Journal, WebMD/Medscape, and Wired

#### **Policy and Ethics**

- National Academies of Sciences, Engineering, and Medicine. The Age of Al in the Life
   Sciences: Benefits and Biosecurity Considerations, consultant.
- National Institutes of Health. Toward an Ethical Framework for Artificial Intelligence in Biomedical and Behavioral Research, workshop participant

#### **Peer Review**

Contributed reviews to Bioinformatics, BMC Bioinformatics, Cell, Cell Systems, Frontiers in Genetics, GigaScience, Intelligent Systems for Molecular Biology (ISMB), Journal of Molecular Biology, Nature, Nature Biotechnology, Nature Communications, Nature Methods, NeurIPS Workshop on Machine Learning in Structural Biology (MLSB), Nucleic Acids Research, PLOS Computational Biology, PLOS ONE, PNAS, PNAS Nexus, Research in Computational Molecular Biology (RECOMB), Science, and Science Translational Medicine

#### PATENTS AND PATENT APPLICATIONS

B. Hie. V. Shanker, and P.S. Kim.

"Antibody compositions and optimization methods."

US Patent App. PCT/US23/17977.

2023

A. Thugabere Jagadeesh and B. Hie.

"Hyperspectral scanning to determine skin health."

US Patent No. 11,532,400.	2022		
	2022		
<b>B. Hie</b> , B. Berger, and H. Cho.  "Realizing private and practical pharmacological collaboration using a neural network			
architecture configured for reduced computation overhead."			
US Patent No. 11,450,439.	2022		
	2022		
<b>B. Hie</b> , B. Bryson, and B. Berger.  "Escape profiling for therapeutic and vaccine development."			
US Patent No. 11,011,253.	2021		
03 Faterit No. 11,011,255.	2021		
AWARDS AND FELLOWSHIPS			
Innovation Investigator, Arc Institute	2024-Present		
STAT Wunderkinds	2022		
Stanford Science Fellows Program	2021-2023		
Department of Defense (DoD) National Defense Science and Engineering Graduate			
(NDSEG) Fellowship	2019-2021		
RECOMB/National Science Foundation Travel Fellowship Award	2019		
Hoefer Prize for Writing in the Major, Nominated, Stanford University	2016		
Tau Beta Pi Engineering Honors Society	2015-Present		
Lunsford Award for Oral Presentation, Nominated, Stanford University	2014		
Boothe Prize for Excellence in Writing, Honorable Mention, Stanford University	2013		
President's Award for Academic Achievement, Stanford University	2013		

2012

National Merit Scholarship Finalist