BRIAN PATRICK ENGLISH

EDUCATION

PhD Harvard University

- MA Harvard University
- BA Cornell University
- Abitur Simpert Kraemer Gymnasium

PROFESSIONAL EXPERIENCE



HHMI/Janelia 19700 Helix Drive Ashburn, VA 20147 englishb@janelia.hhmi.org http://www.brianpenglish.com

| 11/2007 | Single Molecule Studies of Enzymatic Dynamic Fluctuations. Advisor: Xiaoliang Sunnev Xie |
|---------|---|
| 11/2003 | Chemistry and Chemical Biology |
| 01/2001 | Bachelor of Arts with Distinction in all Fields |
| 06/1996 | Krumbach, Bavaria, Germany |

| Howard Hughes Medical Institute Janelia Research Campus Ashburn VA | Senior Scientist Research Scientist (01/2015 – 12/2015) Research Specialist (01/2013 – 12/2014) | 01/2013 – Present | |
|---|---|-------------------|--|
| Albert Einstein College of Medicine | Postdoctoral Fellow | 00/2010 12/2012 | |
| Bronx NY | Anatomy and Structural Biology | 09/2010 - 12/2012 | |
| Uppsala University | Postdoctoral Fellow | 00/2007 08/2010 | |
| Uppsala Sweden | Cell and Molecular Biology | 09/2007 - 00/2010 | |
| Harvard University | Graduate Research Fellow | 00/2001 00/2007 | |
| Cambridge MA | Chemistry and Chemical Biology | 09/2001 - 06/2007 | |
| Cornell University | Research Technician | 01/2001 00/2001 | |
| Ithaca NY | Laboratory of Harold A. Scheraga | 01/2001 - 06/2001 | |
| Cornell University | Undergraduate Research Fellow | 00/1007 10/2000 | |
| Ithaca NY | Chemistry and Chemical Biology | 09/1997 – 12/2000 | |

HONORS

| Janelia & EMBL Biolmaging Seminar Series (<u>co-organizer</u>) | 05/2022 - Present |
|--|-------------------|
| 2015 AAAS Newcomb Cleveland Prize (Lattice light-sheet microscopy) | 02/2016 |
| Postdoctoral Representative to the Einstein Senate Albert Einstein College of Medicine | 10/2010 - 12/2012 |
| Young Researcher Participant of the 59th Meeting of Nobel Laureates in Lindau | 06/2009 |
| Student-nominated Fieser Speaker Harvard Chemistry and Chemical Biology | 04/2007 |
| Eli Lilly Poster Presentation Award 19th Annual Symposium of the Protein Society | 08/2005 |
| George C. Caldwell Prize Cornell Chemistry and Chemical Biology | 10/2001 |
| Phi Beta Kappa Honors Society | 05/2001 |
| 2000 Undergraduate Award in Analytical Chemistry American Chemical Society | 10/2000 |

COMPLETED RESEARCH SUPPORT

| Estonian Science Foundation (ETF) | PUT37 (co-applicant, PI: Vasili Hauryliuk) <i>ppGpp-mediated activation of RSH proteins: from the mechanism of allosteric regulation to computational properties of the stringent response system</i> | 01/2013 – 12/2015 |
|---|--|----------------------|
| Human Frontier Science Program (HFSP) | Cross Disciplinary Fellow (LT00829/2008, PI: Brian English) <i>Transcription factor dynamics in living bacterial cells at the single cell level</i> | 06/2008 – 08/2011 |
| Swedish Research Council (VR) | International Postdoctoral Fellowship (623-2007-8116, PI: Brian English) The Dynamics of Gene Regulation – The Study of Individual Transcription Factor Molecules in Living Cells | 01/2008 – 05/2008 |
| National Institutes of Health (NIH) | Molecular, Cellular and Chemical Biology Training Grant (5 T32 GM07598) Graduate Research Training Fellowship – Harvard University | 01/2002 – 08/2005 |

PUBLICATIONS

37. Galbraith *et al.* Compartmentalized Cytoplasmic Flows Direct Protein Transport to the Cell's Leading Edge. *bioRxiv*. doi: <u>10.1101/2024.05.12.593794</u> (2024)

36. Holland *et al*. A series of spontaneously blinking dyes for super-resolution microscopy. *bioRxiv*. doi: 10.1101/2024.02.23.581625 (2024)

35. Jradi *et al.* Coumarin as a general switching auxiliary to prepare photochromic and spontaneously blinking fluorophores. *bioRxiv*. doi: <u>10.1101/2024.05.12.593749</u> (2024)

34. Feng *et al.* Inhibition of coronavirus HCoV-OC43 by targeting the eIF4F complex. *Frontiers in Pharmacology* 13:1029093. doi: 10.3389/fphar.2022.1029093 (2022) (*open access*)

33. V Gandin*, <u>BP English</u>* *et al.* Cap-dependent translation initiation monitored in living cells. *Nature Communications* 13, 6558 (2022) (*open access*)

32. L Xie et al. BRD2 Compartmentalizes the Accessible Genome. Nature Genetics. doi: 10.1038/ s41588-022-01044-9 (2022)

31. V Gandin*, <u>BP English</u>* *et al.* Cap-dependent translation initiation monitored in living cells. *bioRxiv*. doi: <u>10.1101/2021.05.21.445166</u> (2021)

30. A Ranjan *et al.* Live-cell single particle imaging reveals the role of RNA polymerase II in histone H2A.Z eviction. *eLife.* doi: <u>10.7554/eLife.55667</u> (2020)

29. L Xie, P Dong *et al.* 3D ATAC-PALM: Super-resolution Imaging of the Accessible Genome. *Nature Methods* 17, 430-436 (2020)

28. A Ranjan *et al.* Live-cell single particle imaging reveals the role of RNA polymerase II in histone H2A.Z eviction. *bioRxiv*. doi: <u>10.1101/2020.02.13.947119</u> (2020)

27. L Xie, P Dong *et al.* Super-resolution Imaging Reveals 3D Structure and Organizing Mechanism of Accessible Chromatin. *bioRxiv*. doi: <u>10.1101/678649</u> (2019)

26. JB Grimm, TA Brown, <u>BP English</u> *et al.* Synthesis of Janelia Fluor HaloTag and SNAP-tag ligands and their use in cellular imaging experiments. In: Erfle H. (eds) *Super-Resolution Microscopy*. *Methods in Molecular Biology*, Vol.1663, Humana Press, New York, NY; doi: <u>10.1007/978-1-4939-7265-4_15</u> (2017)

25. JB Grimm*, <u>BP English</u>* *et al.* Bright photoactivatable fluorophores for single-molecule imaging. *Nature Methods* 13(12), 985-988 (2016)

24. YJ Yoon, B Wu *et al.* Glutamate-induced RNA localization and translation in neurons. *PNAS* 113(44), E6877-86 (2016) (*open access*)

23. Z Zhang, <u>BP English</u> *et al.* Rapid Dynamics of General Transcription Factor TFIIB Binding During Preinitiation Complex Assembly Revealed by Single-Molecule Analysis. *Genes and Development* 30, 2106-2118 (2016) (*open access*)

22. LD Lavis*, JB Grimm, <u>BP English</u> *et al.* Bright photoactivatable fluorophores for single-molecule imaging. *bioRxiv*. doi:<u>10.1101/066779</u> (2016)

21. PW Tillberg, F Chen *et al.* Expansion Microscopy of Biological Specimens with Protein Retention. *Nature Biotechnology* 34, 987–992 (2016) (*cover art*)

20. T Morisaki *et al.* Real-time quantification of single RNA translation dynamics in living cells. *Science* 352(6292), 1425-1429 (2016) (*F1000Prime*)

19. WK Cho, N Jayanth, <u>BP English</u> *et al.* RNA Polymerase II cluster dynamics predict mRNA output in living cells. *eLife* 2016;10.7554/eLife.13617 (2016) (*open access*)

18. ZB Katz*, <u>BP English*</u> *et al.* Mapping translation 'hot-spots' in live cells by tracking single molecules of mRNA and ribosomes. *eLife* 2016;5:e10415 (2016) (*open access*, *F1000Prime*)

17. <u>BP English</u>, RH Singer. Tracking multiple single molecules in living cells. **SPIE Newsroom**.

doi: 10.1117/2.1201509.006125 (November 2, 2015) (pdf)

16. <u>BP English</u>*, RH Singer. A three-camera imaging microscope for high-speed single-molecule tracking and superresolution imaging in living cells. *Proc. SPIE 9550, Biosensing and Nanomedicine VIII*, 955008 (*invited paper*); doi:10.1117/12.2190246 (2015) (*PMC article*)

15. N Monnier *et al.* Inferring transient particle transport dynamics in live cells. *Nature Methods* 12(9), 838-840 (2015) (*PMC article*)

14. S Viswanathan *et al.* High-performance probes for light and electron microscopy. *Nature Methods* 12(6), 568–576 (2015) (*PMC article*, *F1000Prime*)

13. JB Grimm, <u>BP English</u> *et al.* A general method to improve fluorophores for live-cell and single-molecule microscopy. *Nature Methods* 12(3), 244 - 250 (2015) (*PMC article, featured in: 1, cover art*)

12. H Jiang*, <u>BP English</u>* *et al.* Tracking surface glycans on live cancer cells with single molecule sensitivity. *Angewandte Chemie International Edition* 54(6), 1765-1769 (2015) (*PMC article*, *C&EN News*)

11. BC Chen, WR Legant, K Wang *et al.* Lattice Light Sheet Microscopy: Imaging Molecules to Embryos at High Spatiotemporal Resolution. *Science* 346(6208), 1257998 (2014) (*PMC article*, *F1000Prime*, *cover art*)

10. KD Piatkevich, <u>BP English</u> et al. Photoswitchable Red Fluorescent Protein with a Large Stokes Shift. **Chemistry & Biology** 21, 1402–1414 (2014) (<u>open access</u>)

9. V Shyp *et al.* Positive allosteric feedback regulation of the stringent response enzyme ReIA by its product. *EMBO Reports* 13, 835-839 (2012) (*open access, featured in: 1*)

8. A Kuzemenko*, S Tankov*, <u>BP English</u>* *et al.* Single molecule tracking fluorescence microscopy inmitochondria reveals highly dynamic but confined movement of Tom40. *Scientific Reports* 1, 195; doi:10.1038/srep00195 (2011) (*open access, SGD curated paper*)

7. <u>BP English</u> *et al.* Single Molecule Investigations of the Stringent Response Machinery in Living Bacterial Cells. **PNAS** 108(31), E365-373 (2011) (*open access*, *E1000Prime*, *featured in*: <u>1</u>, <u>2</u>, <u>3</u>, <u>4</u>)

6. <u>BP English</u>*, A Sanamrad* *et al.* Tracking of individual freely diffusing fluorescent protein molecules in the bacterial cytoplasm. *arXiv* <u>1003.2110v1</u> [q-bio.QM] (2010)

5. <u>BP English</u> et al. Ever-fluctuating single enzyme molecules: Michaelis-Menten equation revisited. **Nature Chemical Biology** 2, 87-94 (2006) (<u>F1000Prime</u>, featured in: <u>1</u>, <u>2</u>, <u>3</u>, <u>cover art</u>)

4. W Min *et al.*. When Does the Michaelis-Menten Equation Hold for Fluctuating Enzymes? **Journal of Physical Chemistry B** 110, 20093-20097 (2006)

3. SC Kou *et al.* Single-Molecule Michelis-Menten Equations. *Journal of Physical Chemistry B* 109, 19068-19081 (2005) (*cover art*)

2. W Min, <u>BP English</u> et al.. Fluctuating Enzymes: Lessons from Single-Molecule Studies. Accounts of Chemical **Research** 38, 923-931 (2005)

1. <u>BP English</u> *et al.* Development of a Novel Method To Populate Native Disulfide-Bonded Intermediates for Structural Characterization of Proteins: Implications for the Mechanism of Oxidative Folding of RNase A. *Journal of the American Chemical Society* 124, 4995-4999 (2002)

RESEARCH INTERESTS

My aim at Janelia is to develop quantitative single cell and multi-color single molecule tracking assays with high spatial and temporal resolution to study when and where molecules are interacting inside living cells and where enzymes are active. At Harvard I developed turnover assays to study activity fluctuations of individual enzyme molecules *in vitro*. The microscope at Uppsala facilitated the *in vivo* tracking of even fast freely diffusing protein molecules. At Einstein we extended this approach to mapping translation by simultaneous tracking thousands of mRNA and ribosome molecules.

AD HOC REVIEWER

| Biomicrofluidics | Biophysical Journal | J of Nanobiotechnology | Philosophical Transactions B | Protein Expression and Purification | Scientific Reports |
|--|--|---|---|---|---------------------|
| TEACHING FELLO | DW AT HARVAI | RD CHEMISTRY | | | |
| Honors Intro | ductory Chem | istry Principle | es of Chemistry | Frontiers in Mo | olecular Biophysics |
| INVITED SEMINA | RS | | | | |
| UT Southwestern <i>Computational</i> & Systems Biology Seminar Series | | Multiplexed Single-mo Nature of Complex Bi | olecule Live-cell Im iological Reactions | aging Reveals the Dyr | namic 11/2017 |
| Massachusetts Ir Technology Biopl | n <mark>stitute of</mark> hysics Semina | Insights into translation ar ribosomes and mRNA | on by simultaneous As | single particle tracking | g of 11/2013 |
| Umeå University International Semi | inar Series | Insights into mRNA tr and mRNAs, and by i | anslation by simult maging of cytoskel | aneous tracking of ribo letal structures in live o | somes 10/2013 |
| Duke University and Chemistry Se | Joint Biology minar | Mechanistic insights f enzymes, ribosomes | from single molecul and mRNAs in bac | le tracking of individua teria and mammalian | l 04/2013 cells |
| University of Tartu Biomedical Technology Seminar | | Simultaneous single r r ribosome | Simultaneous single molecule tracking of β -actin mRNA and the ribosome | | |
| University of Mu Gene Center Sem | n ich (LMU) Inar | Single molecule inves compartmentalization | stigations into β-act | tin mRNA localization a | and 10/2012 |
| University of Mui Invited SFB 594 S | n ich (LMU) Seminar | Live-cell imaging and mammalian cells with microscopy | single molecule tra laser feedback int | acking in bacteria and erference and fluoresc | ence 11/2011 |
| Delft University Applied Physics S | eminar | Stringent Response – | - From the Test-Tub | be to Living Cell | 04/2009 |
| University of Tart Biomedical Techno | t u ology Seminal | A Single Molecule Ap r Living Bacterial Cells | proach to Enzymol | ogy – From the Test-T | ube to 12/2008 |
| Harvard Universi Student-nominated Lecture | ty d Fieser | Fluctuating Single Mo | olecules – Zooming | in on Enzyme Kinetic | s 04/2007 |
| Georgia Institute Molecular Biophys | of Technolog | Fluctuating Single En | zyme Molecules | | 04/2007 |
| Dickinson Colleg | je blloquium | Biophysics of Single I | Molecules – Zoomi | ng in on Enzyme Kinei | tics 10/2006 |

SELECTED CONFERENCE LECTURES

| 4D-Nucleome Annual Meeting North Bethesda | Imaging of Multiple Single-Molecules Reveals the Dynamic Nature of Complex Biological Reactions | 09/2017 |
|--|--|---------|
| 10th Berlin Summer Meeting Berlin | Simultaneous Live-Cell Imaging of Multiple Single-Molecules Reveals the Dynamic Nature of Complex Biological Reactions | 06/2017 |
| 60th Annual Meeting of the Biophysical Society Los Angeles | Simultaneous High-Speed Tracking of Multiple Single-Molecules Reveals Functional Interactions in Living Cells (<u>abstract</u>) | 02/2016 |
| SPIE Optics + Photonics San Diego | A three-camera imaging microscope for high-speed single-molecule tracking and super-resolution imaging in living cells (<i>invited talk</i>) | 08/2015 |

| 8th Berlin Summer Meeting Berlin | Insights into translation by co-movement analysis of ribosomes and mRNAs | 06/2015 |
|--|---|---------|
| EMBO EMBL Symposium: Seeing is Believing 2013 Heidelberg | Insights into mRNA compartmentalization and translation by simultaneous single particle tracking of ribosomes and mRNAs, and by super-resolution imaging of cytoskeletal structures in live cells (<u>meeting</u> <u>booklet</u>) | 10/2013 |
| Focus on Microscopy 2011 Konstanz | Live-cell imaging of invadopodia formation with simultaneous phase- shifted laser feedback interference and fluorescence microscopy (<u>abstract</u>) | 04/2011 |
| 9th HFSP Meeting and 20th Anniversary Celebration,Tokyo | Single Molecule Approach to Stringent Response in Individual Living Bacterial Cells | 06/2009 |
| 232nd American Chemical Society Meeting San Francisco | Ever-fluctuating single enzyme molecules: Michaelis-Menten equation revisited (<u>abstract</u>) | 09/2006 |
| 40th IUPAC World Chemistry Congress Beijing | From Single Molecule Enzymology to Imaging Gene Expression in Live Cells, One Molecule at a Time | 08/2005 |
| SELECTED CONFERENCE PRES | ENTATIONS | |
| EMBO EMBL Symposium: Seeing is Believing 2019 Heidelberg | Initiation of cap-dependent translation monitored by fluorescence auto- and cross-correlation spectroscopy and single particle tracking in living cells | 10/2019 |
| Focus on Microscopy 2015 Göttingen | A three-camera imaging setup and novel cell-permeable dyes for multiplexed single-molecule live cell experiments (<u>abstract</u>) | 04/2015 |
| Focus on Microscopy 2011 Konstanz | Single Molecule Investigations of the Stringent Response Machinery in Living Bacterial Cells (<u>abstract</u>) | 04/2011 |
| 4th Mechanobiology Workshop - Biophysical | Imaging Adhesions with Phase-Shifted Laser Feedback Interference Microscopy | 11/2010 |
| 54th Biophysical Society Meeting San Francisco | Single Molecule Tracking Inside Individual Living Bacterial Cells (<u>abstract</u>) | 02/2010 |
| 9th International Conference on Systems Biology | A Single Molecule Approach to Stringent Response in Individual Living Bacterial Cells | 08/2008 |
| 19th Symposium of the Protein Society Boston | A Michaelis-Menten Study of Individual Beta-Galactosidases | 07/2005 |
| 228th American Chemical Society Meeting Philadelphia | Enzymatic dynamics of individual Beta-Galactosidases (abstract) | 08/2004 |

PROCEDURAL EXPERTISE

Development of biophysical assays, data analysis routines, and simulation algorithms. Live cell multi-color superresolution imaging and single-molecule tracking.

- microscopy design and construction of custom microscopes for single-particle tracking, PALM and STORM super resolution imaging, and light-sheet illumination with code written in Labview (hardware timing) and micro-manager. Construction of a diSPIM microscope. Operation and alignment of the multifocus microscope (MFM). PALM-imaging in combination with expansion microscopy. Phase-shifted laser feedback interference microscopy. Hosting of two commercial STED and STED-FCS microscopes for extended demonstrations for which I was responsible for the independent operation of the instruments, as well as for preparing suitable sample preparations for the entire Janelia community.
- data analysis development of co-movement algorithms, and custom-analysis scripts for the Janelia transcription imaging consortium. Bayesian trajectory analysis. Igor Pro, Matlab, Micro-Manager and LabView programming.

PROFESSIONAL MEMBERSHIPS

| Biophysical Society | American Chemical Society |
|--|---|
| The International Society for Optics and Photonics | The 4D nucleome project: Imaging Tools Initiative Consortium Member |

NATIONALITY

Dual German/American citizen, fluent in both German and English.