Alex J Guseman, PhD

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Education

2014-2018 University of North Carolina at Chapel Hill

Ph.D., Chemistry,

Advisor: Dr. Gary J. Pielak

2010 - 2014 University of Maryland College Park

B.S. Biochemistry

Research Advisor(s): Drs. David Fushman and David Hawthorne

Research Experience

2018-present University of Pittsburgh School of Medicine – Department of Structural Biology

Postdoctoral Scholar, Advisor: Dr. Angela Gronenborn

Merck Fellow of the Life Science Research Foundation

Burroughs Wellcome Fund Postdoctoral Enrichment Program Fellow

Projects: Mechanism of cataract formation in lens-like environments

• Performed biophysical and structural characterization of deamidation variants of human vD crystallin

Expanding the toolbox of in cell Nuclear Magnetic Resonance

• Developed methods for ¹⁹F Nuclear Magnetic Resonance (NMR) in mammalian cells

2014-2018 University of North Carolina at Chapel Hill– Department of Chemistry

Ph.D. Student, Ruth L. Kirschstein Predoctoral Fellow, Advisor: Dr. Gary Pielak Dissertation: Protein Dimerization in Physiologically Relevant Environments

- Developed GB1 homodimers into model system for studying dimerization in living cells
- Determined influence of macromolecular crowding on protein dimerization using ¹⁹F Nuclear Magnetic Resonance spectrometry.
- Demonstrated importance of crowding-induced chemical interactions to protein dimerization
- Adapted and tested Scaled Particle Theory to predict influence of macromolecule crowding on two GB1 homodimers
- Developed methods to quantify test protein concentration in *Escherichia coli* using combination of LC-MS and flow cytometry.
- Developed GB1 variant to probe cytoplasmic pH of living E. coli using in-cell NMR
- Determined structural perturbations to heterochromatin protein-1 upon mutation of trimethyl lysine binding pocket using NMR

2012 - 2014 University of Maryland- Department of Chemistry and Biochemistry

HHMI Undergraduate Research Fellow, Advisor: Dr. David Fushman

Project: Chemical methods to ubiquitinate histones H2A and H2B.

- Purification, chemical ligation, and structural characterization of ubiquitin histone conjugates.
- Developed chemical methods to generate nonnative histone-ubiquitin conjugates

• Performed NMR studies on the folding and binding of histones H2A and H2B

2010 - 2012 University of Maryland- Department of Entomology

Undergraduate Researcher, Advisor: Dr. David Hawthorne

Project: Multi-Drug Resistance Transporters and a Mechanism-Based Strategy for Assessing risks of Pesticide Combinations on Honey Bees

- Maintained populations of *A. mellifera and D. melanogaster* and *A. mellifera* in fields using standard beekeeping methods
- Developed mortality-based bioassays to screen pesticide combinations that result in detrimental synergisms in *A. mellifera and D. melanogaster*.
- Developed behavior-based bioassays to determine neurological impacts of sub-lethal doses of pesticides to *D. melanogaster*.

2009 - 2010 Igene Biotechnologies

Research Intern, Advisor: Dr. Anu Sunderarajan

Project: Optimization of P. Rhodozyma for fermentation of the Carotenoid Astaxathan

- Performed random mutagenesis and screens of *P. Rhodozyma* to discover hyperproducers of Astaxanthin
- Generated auxotrophic strains of *P. Rhodozyma* for downstream use in protoplast fusions

Grants & Fellowships (Total value \$400,308)

2020 - present NIH Loan Repayment Program Grant (\$19,400)

National Eye Institute Health Disparities LRP

University of Pittsburgh, Department of Structural Biology

2019 - present Merck Fellow of The Life Science Research Foundation (\$186,000)

Life Science Research Foundation Postdoctoral Fellow

Sponsor: Merck

University of Pittsburgh, Department of Structural Biology

2019 - present Burroughs Wellcome PDEP Awardee (\$60,000)

Burroughs Wellcome Fund Postdoctoral Enrichment Program

University of Pittsburgh, Department of Structural Biology

2017 - 2018 National Institutes of Health – F31 GM126763 (\$66,808)

Ruth L. Kirschstein NRSA Predoctoral Fellowship to Promote Diversity University of North Carolina at Chapel Hill, Department of Chemistry

2016 - 2017 National Science Foundation- Supplement to MCB1051819 (\$30,000)

Diversity Supplement

University of North Carolina at Chapel Hill, Department of Chemistry

2015 - 2016 National Institutes of Health - T32 GM008570-20 (\$24,000)

Molecular and Cellular Biophysics Training Program

University of North Carolina at Chapel Hill, Department of Chemistry

2014 - 2015 National Institutes of Health- R25 GM055336 (\$25,000)

Diversity Excellence Fellowship, Initiative for Maximizing Student Diversity

University of North Carolina at Chapel Hill, Biological and Biomedical Sciences

Program

2013 - 2014 Howard Hughes Medical Institute Undergraduate Research Fellow (\$6,000)

HHMI, University of Maryland Howard Hughes Medical Institutional Grant

University of Maryland, Department of Chemistry and Biochemistry

2013 Maryland Summer Scholars Fellowship (\$2,500)

Office of Undergraduate Research University of Maryland University of Maryland, Department of Chemistry and Biochemistry

<u>Publications (*denotes equal contribution co-first authorship)</u>

- 14.) Speer, S.L.; Zheng, W.; Jiang, X.; Chu, I.; **Guseman, A.J.;** Liu, M.; Pielak, G.J; Li, C.; "The intracellular environment tunes protein-protein interactions" (Submitted)
- 13.) **Guseman, A.J.;** Whitley, M.J.; Gonzalez, J.J.; Rathi, N.; Ambarian, M; Gronenborn, A.M.; "Assessing the Structures and Interactions of γ D-Crystallin Deamidation Variants" Structure (In Press)
- 12.) Krone, K.W.*; Albanese K.I.*; Leighton, G.O.; He, C.Q.; Lee, G.Y.; Garcia-Borras, M.; **Guseman, A.J.**; Williams D.C.J; Houk, K.N.; Brustad, E.M.; Waters, M.L.; (2020) "Thermodynamic Consequences of Tyr to Trp Mutations in the Cation-π-Mediated Binding of Trimethyllysine by the HP1 Chromodomain" Chemical Science 11 (13) 3495-3500
- 11.) **Guseman, A.J.**; Pielak, G.J.; (2020) Chapter 12: Protein Stability and weak intracellular interactions: In-cell NMR Spectroscopy: From Molecular Sciences to Cell Biology Shirakawa, M. Döstch, V. and Ito, Yutaka. (The Royal Society of Chemistry) pp 188-206
- 10.) Free, M.E.; Stember, K.G.; Hess, H.J.; McInnis, E.A.; Lardinois, O.; Hogan, S.L.; Hu, Y.; Mendoza, C.; Le, A.K.; Guseman A.J.; Pilkinton, M.A.; Bortone, D.S.; Cowens, K.; Sidney, F.; Karosiene, E.; Peters, B.; James, E.; Kwok, W.W.; Vincent, B.G.; Mallal, S.A.; Jennette, C.J.; Ciavatta, D.J.; Falk, R.J. (2019) "Restricted Myeloperoxidase Epitopes Drive the Adaptive Immune Response in ANCA Vasculitis" Journal of Autoimmunity 106:102306
- 9.) Speer, S.L.; **Guseman, A.J;** Patteson, J.B.; Ehrmann B.M.; Pielak, G.J.; (2019) "Controlling and quantifying protein concentration in *Escherichia coli* cells" Protein Science 28:1307-1311
- 8.) **Guseman, A.J.;** Gronenborn A.M.; (2019) "Isomerization, an Achilles Heel to Long-Lived Proteins" Journal of Biological Chemistry 294:7556-7557
- 7.) Piszkiewicz, S.P.; Gunn, K.H.; Warmuth, O.; Propst, A.; Mehta, A.; Nguyen. K.H.; Kuhlman, E.; Guseman, A.J.; Stadmiller. S.S.; Boothby T.C.; Neher, S.B.; Pielak, G.J.; (2019) "Protecting Activity of Desiccated Enzymes" Protein Science 28:5 941-951
- 6.) **Guseman, A.J.**; Perez Goncalves, G.M.; Speer, S.L.; Young, G.B.; Pielak G.J.; (2018) "Protein Shape Modulates Crowding Effects" Proceedings of the National Academies of Sciences 115 (43):10965-10970
- 5.) **Guseman, A.J.*;** Speer, S.L.*; Perez Goncalves, G.M.; Pielak G.J.; (2018) "Surface-Charge Modulates Protein-Protein Interactions in Physiologically Relevant Environments" Biochemistry 57:1681-1684.
- 4.) Stadmiller, S.S.; Gorensek-Benitez, A.H; **Guseman, A.J.;** Pielak, G.J.; (2017) "Osmotic-Shock Induced Protein Destabilization and its Reversal by Glycine Betaine" Journal of Molecular Biology 429 (8), 1155-1161
- 3.) **Guseman, A.J.**; Pielak, G.J.; (2017) "Cosolute and Crowding Effects on a Side-By-Side Protein Dimer" Biochemistry 56 (7):971-976
- 2.) **Guseman, A.J.**; Miller, K.; Kunkle, G.; Dively, G.J.; Pettis, J.S.; Evans, J.D.; vanEngelsdorp, D.; Hawthorne, D.J.; (2016) "Multi-Drug Resistance Transporters and a Mechanism-Based Strategy for Assessing Risks of Pesticide Combinations on Honey Bees" PLoS ONE 11(2): e0148242.

1.) Cohen, R.D.; **Guseman, A.J.**; Pielak, G.J.; (2015) "Intracellular pH Modulates Quinary Structure" Protein Science 24 (11):1748-1755

Presentations

- **2020** International Council for Magnetic Resonance in Biological Systems Early Career Series Deamidation of VD-crystallin Effects on Structure and Interactions Properties <u>Invited Talk</u>
- **2020 64**th **Annual meeting of the Biophysical Society** *Deamidation of γD-crystallin Effects on Structure and Interactions Properties -<u>Invited Talk</u>*
- 2019 Rising Stars in Biomedical URM, Massachusetts Institute of Technology Expanding the tool box of NMR in living mammalian cells <u>Invited Talk</u>
- **2019** Frontiers of Biophysics, International School of Biological Magnetic Resonance Expanding the toolbox of NMR in living mammalian cells Invited Talk
- **2018 Duke University BioCoRE symposium**Developing zebrafish oocytes as a model system for in-cell NMR– <u>Invited Talk</u>
- **2018 Gordon Research Symposium on Protein Folding Dynamics**Protein dimerization in physiologically relevant conditions <u>Invited Talk and Poster</u>
- **2018** Graduate Student Research and Policy Expo Protein Dimerization in living cells – <u>Invited Talk</u>
- **2017** University of Virginia Invited Candidate Symposium

 Protein Dimerization in physiologically relevant environments and in living cells—Invited Talk
- **2017 Diversity in STEM**Crowding and protein dimerization—Poster
- 2017 31st Annual Meeting of the Protein Society
 - Dimer Shape determines effect of macromolecular crowding-Poster
- 2016 30th Annual Meeting of the Protein Society Crowding and Protein Dimerization-Poster
- 2016 60th Annual Meeting of the Biophysical Society Crowding and Protein Dimerization—*Poster*
- **2015 UNC Initiative for Maximizing Student Diversity Symposium** Crowding and protein dimerization *Poster*
- **Mid-Atlantic Prep and IMSD Research Symposium (MAPRS)**Using ¹⁹F NMR to probe the solvation of GB1 in living cells. Poster
- **2014** Howard Hughes Medical Institute Undergraduate Research Symposium

 Generating site specific ubiquitin histone conjugates for study by NMR—Poster
- **Vanderbilt VU-EDGE symposium**Generating site specific ubiquitin histone conjugates for study by NMR—<u>Invited Talk</u>
- **2012** University of Maryland Baltimore County Undergraduate Research Symposium Multipesticide exposure and Honeybee mortality—Poster (award)
- **2010 Howard County Public Schools High School Internship Expo** *Optimization of* P. Rhodozyma *for production of Carotenoids Invited Talk*

Honors and Awards

- 2020 NIH Loan Repayment Program Grant, National Eye Institute
- 2019 Rising Star in Biomedical URM, Massachusetts Institute of Technology
- 2019 Life Science Research Foundation Postdoctoral Fellowship, Sponsored by Merck
- 2019 Burroughs Wellcome Fund Postdoctoral Enrichment Program Fellowship
- 2018 Sigma Xi Research Honor Society
- **2018** Carl Storm Travel Fellowship to attend the Protein Folding Dynamics GRC

2017 Ruth L. Kirschstein NRSA Predoctoral Fellowship to Promote Dive	rsity
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- 2017 Ledoux travel award to attend the Protein Society
- 2016 National Science Foundation Diversity Supplement
- 2015 NIH T32 NRSA Predoctoral
- 2014 NIH R25 Diversity Excellence Fellowship
- 2014 UNC Biological and Biomedical Sciences Program Directors Award
- 2013 Howard Hughes Medical Institute Undergraduate Research Fellowship
- 2013 Maryland Summer Scholars Fellowship
- 2012 University of Maryland Department of Entomology Cory Scholarship
- 2012 University of Maryland College Park Scholars Life Sciences Citation
- 2012 UMBC Undergraduate Research Symposium Poster Award
- 2010 Karl Wolfe Scholarship

Teaching & Mentoring Experience

2015 - 2016	BIOC 649 Mathematics and Macromolecules- Co-Instructor
2016 - 2019	Graduate Students Mentored

Graduate Students Mentored

Shannon Speer, Rotation 2017 (UNC) Joseph Thole, Rotation 2018 (UNC) Jacob Wolfe, Rotation (Pitt)

2014-2018 Undergraduates mentored and where they are now

Gerardo Perez Goncalves, Graduate Student Massachusetts Institute of Technology

Stephen Lanier, Post Baccalaureate Research UNC

Jeremy Gonzalez-Roman. (Back to University of Puerto Rico) 2019-Present

Shivam Khairnar, Pitt

2011-2014 **Undergraduate Chemistry Lab, UMD** – Teaching Assistant for leading lab activities

Memberships and Scientific Service

Gordon Research Symposium Protein Folding Dynamics Co-Chair 2022

Biopolymers In Vivo Postdoctoral Member 2020-2021

2020 Discussion leader Protein Folding Dynamics Gordon Research Symposium

2020 – present Member of the Biophysical Society

Gibbs Biothermodynamics Session Moderator

2019 - present SACNAS abstract and travel award reviewer

2019 - present Society for the Advancement of Chicano/Hispanic and Native American

Scientist (SACNAS)

2018 - present Peer Review ACS Biochemistry, Cell Press Structure, ACS OMEGA, and ACS Journal of Physical Chemistry B

2018 - present Sigma Xi Research Honor Society

2014 - present University of North Carolina Initiative for Maximizing Student Diversity

2011 - present Alpha Chi Sigma Professional Chemistry Fraternity

Conferences Organized

- 2022 Gordon Research Symposium Protein Folding Dynamics
- Diversity in STEM, Co-Organizer and Fundraising Chair, UNC-CH 2018
- 2017 Diversity in STEM, Co-Organizer, UNC-CH
- 2016 Initiative for Maximizing Student Diversity Symposium, Co-Organizer, UNC-CH
- 2015 Initiative for Maximizing Student Diversity Symposium, Co-organizer, UNC-CH