

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 γ D crystallin
 - Developed methods to study crystallin biophysics in intact lens tissues.
- Expanding the toolbox of in cell Nuclear Magnetic Resonance*
- Developed methods for ^{19}F Nuclear Magnetic Resonance (NMR) in mammalian cells
- Development of lectins as inhibitors to SARS-CoV2 spike protein.*
- Performed biological, biophysical, and structural characterization of lectins and the SARS-CoV2 Spike protein to determine mechanism of inhibition
- 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 ^{19}F 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
- 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.
- 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
- Developed mortality-based bioassays to screen pesticide combinations that result in detrimental synergisms in *A. mellifera* and *D. melanogaster*.

Grants & Fellowships (Total value \$405,308)

- 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 (\$191,000)**
 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

Publications (*denotes equal contribution co-first authorship)

- 17.) **Guseman, A.J.**; Nambulli, S.K.; Murphy, L.; Bhinderwhala, F.; Vergara, S.; Duprex, W.P.; Gronenborn A.M.; “Targeting Spike Glycans as an Inhibitory Mechanism of SARS-CoV2 Viral Entry” (in preparation)
- 16.) Zhu, W.; **Guseman A.J.**; Bhinderwala, F.; Lu, M.; Su, X.; Gronenborn, A.M. “Visualizing Proteins in Mammalian Cells by ¹⁹F NMR” (in preparation).
- 15.) Zhao, H.; Wu, H.; **Guseman, A.J.**; Abeykoon, D.; Camara, C.; Dalal, Y.; Fushman, D.; Papoian, G.; “The Principles of Histone Folding in Archaea and Eukarya” (in preparation)

- 14.) Speer, S.L.; Zheng, W.; Jiang, X.; Chu, I.; **Guseman, A.J.**; Liu, M.; Pielak, G.J.; Li, C.; (2021) “The intracellular environment tunes protein-protein interactions” *Proceedings of the National Academies of Sciences* 118:e2019918118.
- 13.) **Guseman, A.J.**; Whitley, M.J.; Gonzalez, J.J.; Rathi, N.; Ambarian, M; Gronenborn, A.M.; (2021) “Assessing the Structures and Interactions of γ D-Crystallin Deamidation Variants” *Structure* 29:3:284-291
- 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, KG.; 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.) Piszkiwicz, 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.; Gorenssek-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

- 2021 35th annual Meeting of the Protein Society**
Assessing the Structures and Interactions of γ D-Crystallin Deamidation Variants- *Invited Talk*
- 2021 Great Lakes Regional Meeting of the American Chemical Society- *Invited Talk***
Assessing the Structures and Interactions of γ D-Crystallin Deamidation Variants. - *Invited Talk*
- 2021 Future Faculty Symposium, University of Chicago Department of Chemistry**
Development of non-RBD targeting inhibitors of SARS-Cov2 Spike protein. - *Invited Talk*
- 2020 International Council for Magnetic Resonance in Biological Systems Early Career Series**
*Deamidation of γ D-crystallin – Effects on Structure and Interactions Properties - *Invited Talk**
- 2020 64th Annual meeting of the Biophysical Society**
*Deamidation of γ D-crystallin – Effects on Structure and Interactions Properties -*Invited Talk**
- 2019 Rising Stars in Biomedical URM, Massachusetts Institute of Technology**
*Expanding the tool box of NMR in living mammalian cells - *Invited Talk**
- 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– *Invited Talk**
- 2018 Gordon Research Symposium on Protein Folding Dynamics**
*Protein dimerization in physiologically relevant conditions – *Invited Talk and Poster**
- 2018 Graduate Student Research and Policy Expo**
*Protein Dimerization in living cells – *Invited Talk**
- 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**
- 2014 Howard Hughes Medical Institute Undergraduate Research Symposium**
*Generating site specific ubiquitin histone conjugates for study by NMR— *Poster**
- 2010 Howard County Public Schools High School Internship Expo**
*Optimization of P. Rhodozyma for production of Carotenoids— *Invited Talk**

Honors and Awards

- 2021** Diversity Award, The Protein Society
- 2021** Dr. Eddie Méndez Award, Fred Hutch Cancer Research Center
- 2021** Future Faculty Conference, University of Chicago Department of Chemistry
- 2021** Intersections Fellows Symposium Associate
- 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
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
2010 Karl Wolfe Scholarship

Teaching & Mentoring Experience

- 2015 - 2016** **BIOC 649 Mathematics and Macromolecules-** Co-Instructor
2016 - 2019 **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
 Raixie Melendez-Pacheco (Back to University of Puerto Rico Rio Piedras)
2019-Present Jeremy Gonzalez-Roman, (Back to University of Puerto Rico (soon to be back at Pitt!))
2011-2014 **Undergraduate Chemistry Lab, UMD** – Teaching Assistant for leading lab activities

Memberships and Scientific Service

- 2022** Gordon Research Symposium Protein Folding Dynamics Co-Chair
2020-2021 Biopolymers *In Vivo* Postdoctoral Member
2020 Discussion leader Protein Folding Dynamics Gordon Research Symposium
2020 – present Member of the Biophysical Society
2019 Gibbs Bio thermodynamics 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, International Journal of Macromolecules, Journal of Molecular Biology and others.
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
2018 Diversity in STEM, Co-Organizer and Fundraising Chair, UNC-CH
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

Pending funding

K99 MOSAIC NIGMS/NIAID “Developing Lectins as Inhibitors of Coronavirus Spike Proteins”
 Burroughs Wellcome Fund Careers At Scientific Interfaces “Peering into the Crystallin Ball of the Eye Lens To See the Mechanism of Cataract Formation”