## Curriculum Vitae **Kathleen Mulvaney**

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### **EDUCATION**

#### UNIVERSITY OF NORTH CAROLINA CHAPEL HILL PhD in Cell Biology

#### Dissertation: Proteomic dissection of KEAP1/NRF2 signaling identifies novel pathway interactors

- Courses in Cell Biology, Biochemistry, Molecular Biology, Cancer Biology and Immunology
- Workshops in Critical Thinking, Grant Writing and Presenting Scientific Research

#### UNIVERSITY OF ROCHESTER

#### **Bachelors of Science in Biology**

#### **Concentration in Molecular Genetics with Distinction in Research**

Advanced coursework in Molecular Biology, Genetics, Biochemistry, Tumor Immunology, Microbiology, Cell Biology, Cognitive Science and Neuroscience

### **RESEARCH EXPERIENCE**

#### THE BROAD INSTITUTE: DANA FARBER CANCER INST. Postdoctoral Fellow with Dr. William (Bill) Sellers

- Research focus on targeting protein-protein interactions for therapeutic development in cancer •
- Therapeutically targeting PRMT5 on a novel binding interface in pancreatic cancer
- Led a productive team of 2-3 research associates funded by NIH and Deerfield Management support •
- Collaborated with the Center for Development of Therapeutics to complete drug discovery project •
- Culminated in two research publications and one patent •

#### UNIVERSITY OF NORTH CAROLINA CHAPEL HILL Postdoctoral Fellow with Dr. Ben Major

Completed screen and manuscript identifying NRF2 interacting proteins and transcriptional regulators •

### UNIVERSITY OF NORTH CAROLINA CHAPEL HILL

#### Graduate Student with Dr. Ben Major

- Research focus on the KEAP1-NRF2 signaling pathway in cancer •
- Described a function for KEAP1 in cell cycle regulation through ubiquitylation of MCM3
- Proteomic screening identified novel NRF2 interacting proteins and transcriptional regulators
- Culminated in successful defense of PhD, a first-author paper, a first-author review and three additional manuscripts

#### DANA FARBER CANCER INSTITUTE, HARVARD UNIVERSITY **Research Assistant with Dr. Alan Engelman**

Performed ChIP-Sequencing project to determine lens epithelial derived growth factor, LEDGF, • binding sites and thereby identify preferential HIV-1 integration sites throughout the genome

#### UNIVERSITY OF ROCHESTER Lab Assistant with Dr. Shawn Murphy

ROCHESTER, NY February 2007 - May 2009

ROCHESTER, NY May 2010

CHAPEL HILL, NC

December 2016

CHAPEL HILL. NC December 2016-May 2017

CHAPEL HILL. NC

### Spring 2011-December 2017

BOSTON, MA

**Summer 2009** 

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# CAMBRIDGE. MA

June 2017-Present

- Research Focus on Diffuse Large B-Cell Lymphoma and expression of MHC Class II and CIITA with the underlying goal of identifying the association between expression levels and patients' immunological responses to DLBCL tumors
- Culminated in writing and successful defense of senior thesis and two co-authored manuscripts

### PUBLICATIONS

- <u>Mulvaney, KM</u>; Blomquist, C; Acharya, N; Li, R; Ranaghan, MJ; O'Keefe, M; Rodriguez, DJ; Young, MJ; Kesar, D; Pal, D; Stokes, M; Nelson, AJ; Jain, SS; Yang, A; Mullin-Bernstein, Z; Columbus, J; Bozal, FK; Skepner, A; Raymond, D; ... Sellers, WR. Molecular basis for substrate recruitment to the PRMT5 methylosome. *Molecular Cell*. 2021, 81(17)17, P3481-3495.
- McKinney, DC; McMillan, BJ; Ranaghan, MJ; Moroco, JA; Brousseau, M; Mullin-Bernstein, Z; O'Keefe, M; McCarren, P; Mesleh, M; <u>Mulvaney, KM</u>; Robinson, F; Singh, R; Bajrami, B; Wagner, FF; Hilgraf, R; Drysdale, MJ; Campbell, AJ; Skepner, A; Timm, DE; Porter, D; Kaushik, VK; Sellers, WR; Ianari, A. Discovery of a First-in-Class Inhibitor of the PRMT5–Substrate Adaptor Interaction. *Journal of Medicinal Chemistry*. 2021, 64 (15), 11148-11168.
- Song, S; Nguyen, V; Schrank, T; <u>Mulvaney, KM</u>; Walter, V.; Wei, D.; Orvis, T.; Desai, N.; Zhang, J.; Hayes, DN; Zheng, Y.; Major, MB; Weissman, B. Loss of SWI/SNF chromatin remodeling alters NRF2 signaling in non-small cell lung carcinoma. *Mol Cancer Res.* 2020, Dec; 18(12):1777-1788.
- Bowman BM; Montgomery SA Schrank TP, Simon JM, Ptacek TS, Tamir TY, <u>Mulvaney KM</u>, Weir SJ, Nguyen TT, Murphy RM, Makowski L, Hayes DN, Chen XL, Randell SH, Weissman BE, Major MB. A conditional mouse expressing an activating mutation in NRF2 displays hyperplasia of the upper gastrointestinal tract and decreased white adipose tissue. *J Pathol.* 2020 Oct;252(2):125-137.
- Mulvaney, KM.; Matson, J.; Siesser, P.; Tamir, TY; Goldfarb, D.; Jacobs, T.; Cloer, EW; Cook, JG; Major, MB. Identification and Characterization of MCM3 as a KEAP1 Substrate. *Journal of Biological Chemistry*, 2016 Nov 4; 291(45): 23719–23733.
- 6. Tamir, TY\*; <u>Mulvaney, KM\*</u>; Major, MB. Dissecting the Keap1/Nrf2 pathway through proteomics. *Current Opinion in Toxicology*, (\*equal contribution) 2016, 1:118-24.
- Guntas, G; Lewis, S; <u>Mulvaney, KM</u>; Cloer, E; Tripathy, A; Lane, T; Major, MB; Kuhlman, B. Engineering a genetically encoded competitive inhibitor of the KEAP1-NRF2 interaction via structure- based design and phage display. *Protein Eng Des Sel.* 2016 Jan;29(1):1-9.
- Hast, BE; Goldfarb D\*; <u>Mulvaney KM\*</u>; Hast MA; Siesser PF; Yan F; Hayes DN; Major MB. Proteomic analysis of ubiquitin ligase KEAP1 reveals associated proteins that inhibit NRF2 ubiquitination. *Cancer Research*. 2013 1;73(7):2199-210. (\*equal contribution)
- Cycon, KA; <u>Mulvaney, KM</u>; Rimsza, LM; Persky, D.; Murphy, SP. Histone deacetylases (HDACs) contribute to the silencing of CIITA expression in DB diffuse large B cell lymphoma. *Immunology*. 2013 Oct; 140(2): 259–272.
- Bushway, M.; Cycon, KA; <u>Mulvaney, KM</u>; Murphy, SP. Coordinate loss of MHC class II expression in the diffuse large B cell lymphoma cell line OCI-Ly2 is due to a novel mutation in RFX-AP. *Immunogenetics*. 2010, 62(2):109-16.

### PUBLICATIONS IN PROGRESS

- 1. <u>Mulvaney, KM</u>; Bozal, FK; Sellers, WR. Monitoring and disrupting cellular PRMT5-substrate adaptor complex formation in in-tact and permeabilized cells using a split luciferase complementation assay. *STAR Protocols, Cell Press.* Manuscript in preparation.
- Pal, D; Rodriguez, D; Chang, L; Robinson, F; Potter, D; Kesar, D; Li, R; Van Tienen, L; <u>Mulvaney</u>, <u>KM</u>; Jung, N; Jain, S; Park, J; Letai, A; Sellers, WR. WSB2 is a novel regulator of the cellular

apoptotic pathway and a therapeutic target in melanoma. Submitted to Cancer Cell.

### **PROJECTS IN PROGRESS**

- 1. Identifying mechanisms of resistance and sensitizers to PRMT5 chemical inhibitors using single and digenic CRISPr KO screens.
- 2. In collaboration with Dr. Eric Fischer: Structurally resolving a full-length methylosome-substratesubstrate adaptor complex by Cryo-EM.

### PATENTS

1. Arthur Campbell, Martin Drysdale, Robert Hilgraf, Alessandra Ianari, Patrick McCarren, David McKinney, Brian McMillan, Kathleen Mulvaney, Dale Porter, William Sellers, Ritu Singh, Florence Wagner. Co-inventor, PCT Application Serial No. PCT/US2021/045016, filed on August 6, 2021 and entitled, "Substrate Adaptor Inhibitors of PRMT5 and Uses Thereof". (inventors listed alphabetically)

### **AWARDS**

•	NIH F32 Ruth L. Kirschstein Postdoctoral Individual National Research Service Award	(2018-2021)
•	NIH LRP Award	(2019-2021)
•	Sigma Xi Research Society Graduate Student Travel Award	(2015)
•	National Science Foundation Graduate Research Fellowship Honorable Mention	(2011, 2012)
•	Degree with Distinction in Research: an honor awarded by the University of Rochester	(2009)
•	National Science Foundation David T. Kearns Scholar	(2008-2010)
•	Take Five Scholar: Awarded tuition-free fifth year by the University of Rochester to study "Influences	
	on Cognitive and Personality Development"	(2009-2010)
•	Academic Competitiveness Grant awarded by New York State	(2007-2008)
•	International Baccalaureate Scholar awarded as a four-year merit scholarship	(2005-2009)

### PRESENTATIONS

NATIONAL MEETINGS			
FASEB Biological Methylation Conference	June 2022, Accepted Abstract-Talk		
AACR Annual Meeting April 2022, Accepted Abstract-Poster			
AACR Annual Meeting	April 2016		
<ul> <li>Presented "KEAP1 ubiquitinates MCM3 to regulate cell cycle in cancer" poster</li> </ul>			
FASEB Ubiquitin ConferenceJune 2014			
• Presented "KEAP1 ubiquitinates MCM3" poster to the ubiquitin-proteasome signaling community			
DANA FARBER HARVARD CANCER CENTER			
Mass. General Hospital Postdoc Association Invited Seminar	February 2022		
• Presented "Drugging Enzymes in Cancer" to a general audie	nce, invited seminar		
Ludwig Meeting Invited Seminar	July 2021		
• Presented "Resolving substrate recruitment to the PRMT5 methylosome" invited seminar			
Molecular and Cellular Oncology Retreat	June 2019		
• Awarded Best Poster Presentation: "PRMT5-substrate adaptor interface as a novel therapeutic target			
in MTAP null tumors"			
BROAD INSTITUTE			
Lunch and Learn July 2019			

#### Co-led department-wide invited chalk talk: "Opportunities and challenges in following up cancer • dependencies: lessons learned from PRMT5 and NXT1"

UNIVERSITY OF NORTH CAROLINA CHAPEL HILL

### **University Research Day**

Awarded Best Oral Presentation

### **IMSD** Symposium

Awarded Best Oral Presentation

### UNIVERSITY OF ROCHESTER

### **Office of Admissions**

Invited to present regionally to prospective students on undergraduate research opportunities •

### **TEACHING EXPERIENCE**

#### HARVARD UNIVERSITY, HARVARD EXTENSION SCHOOL, CAMBRIDGE, MA **BIOS E-30 Epigenetics & Gene Regulation Teaching Assistant and Guest Lecturer** Fall 2020, Fall 2021

- Led weekly independent recitation and study sections, guided and graded the writing assignments
- Teaching 2 lectures per seminar on cancer epigenetics (2021)

#### **BIOS E-16 Cell Biology Teaching Assistant**

- Led weekly independent recitation and study sections
- Guided and graded the writing assignments, problem sets, and exams

### BROAD INSTITUTE, CAMBRIDGE, MA

### **Graduate Student Mentoring**

• Mentoring Harvard BBS students rotating in the lab in critical thinking and scientific research

### **Research Associate Mentoring**

• Mentoring and promoting the career development of three post-undergraduate researchers in the lab.

### UNIVERSITY OF NORTH CAROLINA, CHAPEL HILL, NC

### **Science-at-Hand**

Give monthly scientific demonstrations and discuss careers in science with high school students. •

### **Undergraduate Mentoring**

• Mentoring two undergraduate researchers on research projects for credit in the Major Lab.

### **First Year Group**

Peer mentor to first year students, advise on lab rotations and giving scientific presentations.

### North Carolina DNA Day

• Taught high school students about pharmacogenomics and encouraged exploring careers in science.

### UNIVERSITY OF ROCHESTER, ROCHESTER, NY

### **Upward Bound Tutoring**

National initiative to assist high school students from disadvantaged backgrounds enter college

### **Refugee Tutoring**

### **Fall 2009-Spring 2010**

**Fall 2008-Spring 2011** 

# September 2017-Present

Spring 2019; Spring 2021

### June 2017-Present

March 2015

November 2014

December 2008

### **October 2014-May 2017**

### January 2014- May 2017

### Fall 2014

### April 2011; April 2013



• Program to assist high school, middle school refugees from Turkey and Somalia relocated to the US

#### Lab Teaching Assistant, Department of Chemistry

Fall 2006

• Prepared laboratory set-up for general chemistry students and reviewed and graded lab reports.

#### LABORATORY TECHNIQUES

**PROTEOMICS:** Experience with purification of protein complexes for identification by mass spectrometry (affinity, immunoprecipitation and BioID), isolation and detection of whole cell and targeted post-translational modifications (phosphorylation, ubiquitylation, methylation) by mass spectrometry, and analysis and visualization of mass spectrometry data (MaxQuant, Proteome Discoverer, Skyline, Spotlite, Cytoscape).

**MOLECULAR BIOLOGY & BIOCHEMISTRY:** Experience with mammalian cell culture (primary and immortalized), RNA and DNA isolation, qPCR, RNA-seq, genetic manipulation including CRISPr, cloning and site-directed mutagenesis, flow cytometry, chromatin immunoprecipitation, luciferase reporter assays, drug treatments and viability and proliferation assays, protein expression and purification, Western blotting, fixed and live-cell light and fluorescence microscopy, proximity ligation assays, mouse handling, MEF line derivation, subcutaneous xenografts, recombinant protein purification, size exclusion chromatography, and cellular and *in vitro* binding and ubiquitylation assays.

**GENOMICS, SCREENING AND NEXT-GEN SEQUENCING:** Experience with single and digenic CRISPr knockout screens, small molecule screening, library generation and Illumina sequencing.

### PROFESSIONAL MEMBERSHIPS

American Association for Cancer Research (AACR) (2016-present) American Association for the Advancement of Science (AAAS) (2015-present)

### JOURNAL REVIEWER

Molecular Cancer Research (2019-present)