# Henry E. Miller

PhD Candidate, F31 NIA/NIH Fellow

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#### **Personal statement**

My long-term research focus is to understand the factors which drive epigenetic aging and rejuvenation to develop novel therapies for aging and age-related diseases. During the remainder of my PhD training, I will continue to solidify my skills as a bioinformatician. After graduating (expected Dec. 2022), I hope to leverage these skills to address fundamental questions in epigenetic aging and rejuvenation. Success in this research direction may yield deeper insights into the aging epigenome, novel aging clocks, and drug targets for clinical translation.

#### Education

| Aug 2018 – Current  | UT Health San Antonio<br>Doctor of Philosophy<br>Biology of Aging<br>GPA: 4.0<br>Qualifying exam: Passed with distinction |
|---------------------|---|
| Aug 2016 – May 2018 | SUNY University at Buffalo<br>Master of Science<br>Biomedical Engineering<br>GPA: 3.8                                     |
| Aug 2012 – May 2016 | Christopher Newport University<br>Bachelor of Science<br>Double major, Neuroscience and Philosophy<br>GPA: 3.1            |

#### **Research Experience**

| 2018 - Current | <b>Bishop Laboratory</b><br>Supervisor: Alexander Bishop, DPhil<br><i>UT Health San Antonio</i>   |
|----------------|---|
|                | As a PhD candidate, I use bioinformatics to understand the role of R-loops in cancer, development, and aging.   |
| 2018 - 2018    | <b>Chen Laboratory</b><br>Supervisor: Yidong Chen, PhD<br><i>UT Health San Antonio</i>  |
|                | As a rotation student, I benchmarked methodologies for RNA-Seq splicing analysis.   |
| 2018 - 2018    | <b>Frost Laboratory</b><br>Supervisor: Bess Frost, PhD<br><i>UT Health San Antonio</i>  |
|                | As a rotation student, I analyzed calcium-dependent transcriptional signatures and transposable-<br>element expression in human, mouse, and fly RNA-Seq data. |

| 2017 - 2017 | Lovell Nanomedicine Laboratory<br>Supervisor: Jonathan Lovell, PhD<br>SUNY University at Buffalo  |
|-------------|---|
|             | I generated three peanut allergy mouse models and formulated a novel drug based on phos-<br>phatidylserine liposomes designed to attenuate peanut allergies.  |
| 2017 - 2017 | <b>Engineered Theranostics Laboratory</b><br>Supervisor: Yun Wu, PhD<br><i>SUNY University at Buffalo</i>   |
|             | I completed a short project which involved designing and fabricating microfluidic chips to enhance<br>the interaction of liposome nanoparticles and lung-cancer cell-derived exosomal miRNAs.                                     |
| 2015 - 2016 | <b>Eye-blink Conditioning Laboratory</b><br>Supervisor: Matthew Campolattaro, PhD<br><i>Christopher Newport University</i>  |
|             | As an undergraduate research student, I assisted the Campolattaro laboratory with data collection, tissue histology, and perfusion. Additionally, I performed rodent stereotaxis surgery and learned post-operative care methods. |
| 2015 - 2016 | <b>Velkey Fish Behavior Laboratory</b><br>Supervisor: Andrew Velkey, PhD<br><i>Christopher Newport University</i>   |
|             | As an undergraduate research student, I assisted the Velkey laboratory with running behavioral trials and collecting data.  |

# Publications

# In press

| 2022 | <ul> <li>Miller HE, Ilieva M, Bishop AJR, Uchida S. Current Status of Epitranscriptomic Marks Affecting lncRNA Structures and Functions. Non-coding RNA. 2022, Mar 28;8(2):23. doi: https://doi.org/10.3390/ncrna8020023.</li> <li>Ilieva M, Miller HE, Agarwal A, Paulus GK, Madsen GH, Bishop AJR, Kauppinen S, Uchida S. Expression analysis of protein-coding and long non-coding RNA genes in fibrosis. Non-coding RNA. 2022, Jan 28;8(1):13. doi: https://doi.org/10.3390/ncrna8010013.</li> </ul>  |
|------|---|
| 2021 | • Miller HE, Bishop AJR. Correlation AnalyzeR: functional predictions from gene co-expression correlations.<br>BMC Bioinformatics. 2021 Apr 20;22(1):206. doi: 10.1186/s12859-021-04130-7. PMID: 33879054; PMCID: PMC8056587.   |
| 2020 | <ul> <li>Miller HE, Gorthi A, Bassani N, Lawrence LA, Iskra BS, Bishop AJR. Reconstruction of Ewing Sarcoma Developmental Context from Mass-Scale Transcriptomics Reveals Characteristics of EWSR1-FL1 Permissibility. Cancers. 2020 Apr 11;12(4):948. doi: 10.3390/cancers12040948. PMID: 32290418; PMCID: PMC7226175.</li> <li>Pan H, Jin M, Ghadiyaram A, Kaur P, Miller HE, Ta HM, Liu M, Fan Y, Mahn C, Gorthi A, You C, Piehler J, Riehn R, Bishop AJR, Tao YJ, Wang H. Cohesin SA1 and SA2 are RNA binding proteins that localize to RNA containing regions on DNA. Nucleic Acids Res. 2020 Jun 4;48(10):5639-5655. doi: 10.1093/nar/gkaa284.</li> <li>Mahoney R, Ochoa Thomas E, Ramirez P, Miller HE, Beckmann A, Zuniga G, Dobrowolski R, Frost B. Pathogenic Tau Causes a Toxic Depletion of Nuclear Calcium. Cell Rep. 2020 Jul 14;32(2):107900. doi: 10.1016/j.celrep.2020.107900. PMID: 32668249; PMCID: PMC7428851.</li> </ul> |

# In preparation, preprint, or under review/revision

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- Miller HE & Mahmud S, Mukhopadhyay P, Movasati A, Shafiha R, Vines A, Bishop A & Venkitaraman A. BRCA2 and PAF1 coordinate in the regulation of transcriptional pausing. (In preparation).
- Borbeck, J, Miller HE, Shekhovtsov, S, Montemayor, D, Bernstein, M, Bishop A. autoPLIER: a pathwayconstrained autoencoder for biologically interpretable low-dimensional embedding. (In preparation).
- Romero JC, Tonapi S, Loranc E, Bassani N, Lawrence LA, **Miller HE**, Robledo DG, Cao L, Nie J, Kanda K, Gorthi A, Lane AN, Fan TWN, Zha S, Musi N, Bishop AJR. ATM protein kinase regulates CD98HC dependent antiport activities to control cell metabolism. (**Under revision, Molecular Cell**).
- Miller HE, Montemayor D, Li J, Levy SA, Pawar R, Hartono S, Sharma K, Frost B, Chedin D, Bishop AJR. Exploration and analysis of R-loop mapping data with RLBase. **bioRxiv**; doi: https://doi.org/10.1101/2021.11.01.466854. (preprint).
- Miller HE, Montemayor D, Abdul J, Vines A, Levy SA, Hartono S, Sharma K, Frost B, Chedin D, Bishop AJR. Quality-controlled R-loop meta-analysis reveals the characteristics of R-Loop consensus regions. bioRxiv; doi: https://doi.org/10.1101/2021.11.01.466823. (preprint, Under Revision, Nucleic Acids Res.).
- Iskra B, Davis L, Miller HE, Chiu YC, Bishop AJR, Chen Y, Aune GJ.Assessing the heterogeneity of cardiac non-myocytes and the effect of cell culture with integrative single cell analysis. bioRxiv; doi: https://doi.org/10.1101/2020.03.04.975177. (preprint, Under Review, AJP Heart).

# Conference presentations and speaking engagements

| 2020 | Sarcoma Working Group Meeting  |
|------|--|
|      | Greehey Children's Cancer Research Institute   |
|      | Presented talk entitled "Reconstruction of Ewing Sarcoma Developmental Context from Mass-Scale Transcriptomics |
|      | Reveals Characteristics of EWSR1-FLI1 Permissibility"  |
| 2020 | Mikiten Research Symposium   |
|      | UT Health San Antonio  |
|      | Presented talk, "Bioinformatics for Biologists."   |
|      |  |

2017 Northeast Bioengineering Conference
 New Jersey Institute of Technology
 Presented poster, "Slit-based linear-array photoacoustic tomography system for vascular imaging in human."

# Funding

| Sep 2021 - Current  | NIA/NIH F31 predoctoral fellowship (F31AG072902). |
|---------------------|---|
| Sep 2020 - Sep 2021 | Greehey Foundation predoctoral fellowship         |

# Work Experience

| Mar 2017 - May 2018 | <b>Technology Research Intern</b><br>University of Buffalo School of Engineering   |
|---------------------|--|
|                     | The University of Buffalo School of Engineering hired me as an Entrepreneurship Intern. In this experience, I developed a business plan and marketing materials for a new internet technology startup. |

| Mar 2017 - May 2018 | Lead Venture Coach<br>Blackstone Launchpad at The University of Buffalo  |
|---------------------|--|
|                     | Blackstone Launchpad at The University of Buffalo provides training and guidance for student<br>and faculty entrepreneurs. As the Lead Venture Coach, I recruited, trained, and led a team of<br>eight student entrepreneurs to coach more than 100 entrepreneurs.   |
| Apr 2017 - May 2018 | Chief Communications Officer<br>POP Biotechnologies Inc  |
|                     | POP Biotechnologies Inc is a Buffalo startup developing nanomedicine therapeutics for treating cancer and infectious diseases. As Chief Communications Officer, I handled business development activities, such as pitching investors, issuing press releases, and executive recruitment.  |
| Aug 2016 - Dec 2017 | <b>Director of Operations</b><br>Sonioptix LLC   |
|                     | Sonioptix LLC is a Buffalo startup creating cutting-edge cancer screening devices. As director of operations, I handled all business development activities, such as pitching investors. During my tenure, Sonioptix LLC recieved funding to begin clinical studies in Western New York for their breast cancer screening device.                |
| May 2016 - Aug 2016 | Environmental Health Technician<br>Fairfax County Health Department  |
|                     | I was a summer employee for the Environmental Health division of Fairfax County Health<br>Department. In this position, I assisted with surveillance, trapping, and analysis of mosquitoes<br>and ticks in local parks. I also provided technical assistance in repairing broken traps.  |
| Jan 2016 - May 2016 | <b>Biochemistry Course Tutor</b><br>Professor Lasseter, Christopher Newport University   |
|                     | I was a tutor for the Biochemistry for Pre-Medical Students course at Christopher Newport<br>University. In this capacity, I met with students for one-on-one tutoring sessions in which I assisted<br>them in mastering challenging course topics.  |
| Aug 2014 - May 2016 | <b>Philosophy Department Tutor</b><br>Christopher Newport University   |
|                     | I was the department tutor for the Philosophy and Religious Studies department at Christopher<br>Newport University. In this capacity, I met with students for one-on-one tutoring sessions in<br>which I assisted them in mastering challenging topics from a variety of department courses. I also<br>substitute lectured for several classes. |
| May 2014 - Aug 2014 | Environmental Health Technician<br>Fairfax County Health Department  |
|                     | I was a summer employee for the Environmental Health division of Fairfax County Health<br>Department. In this position, I performed health and safety inspections of local public pools and<br>water parks.  |

#### Leadership and Volunteer Experience

| Oct 2020 - Current  | Founder and President<br>Bioinformatics Research Network   |
|---------------------|--|
|                     | The Bioinformatics Research Network (BRN) is a 501(c)(3) non-profit dedicated to making high-<br>quality bioinformatics training and professional opportunities accessible to all people. It provides<br>free online bioinformatics training and project opportunities. I founded BRN in 2020 and have<br>guided it as president of the board since September 2021. Under my leadership, BRN has reached<br>a membership of 400 people, completed five research project, published two pre-prints and<br>one peer-reviewed article, and helped eight members achieve professional opportunities or PhD<br>positions in bioinformatics. |
| May 2020 - Nov 2021 | <b>Co-Founder and Training Chair</b><br>BIG Bioinformatics   |
|                     | BIG Bioinformatics is a UT Health student club that provides free online bioinformatics workshops. I co-founded BIG Bioinformatics in 2020 and have led the development of five full-length workshops. During my tenure as training chair, I created five workshops which were attended by more than 300 participants in total. The video recordings from these workshops have approximately 500 subscribers, 15k views, and 2k hours of watch time.   |

#### **Professional Organizations**

| 2020 - Current | Member, The International Society for Computational Biology (ISCB)                          |
|----------------|---|
| 2020 - 2021    | Graduate student representative, Bioinformatics Curriculum Committee, UT Health San Antonio |
| 2017 - 2018    | Member, Project Management Institute  |
| 2017 - 2018    | Member, Biomedical Engineering Society  |
| 2017 - 2018    | Member, Institute of Electrical and Electronics Engineers                                   |

#### Academic and Professional Honors

- 2022 VitaDAO Longevity Fellowship Award, Decentralized / Online
- 2020 Passed qualifying exam with distinction, UT Health San Antonio
- 2016 | Induction in Omicron Delta Kappa, Christopher Newport University
- 2013 Dean's List, Christopher Newport University

#### Open-source software, analyses, and web applications

\*Software on which I am the primary developer/maintainer

- **autoPLIER** (under development): An python package for using the autoPLIER pathway-constrained autoencoder. Source code
- FibroDB\*: An R-Shiny app for exploring results from fibroblast RNA-Seq studies. 
  Web App 
  Source code
- RLBase\*: An R-Shiny app for exploring and analyzing R-loop mapping datasets. 

  Web App 
  Source code
- **RLSeq**\*: An R/Bioconductor package for analyzing R-loop mapping datasets. ( Bioconductor page Source code
- RLHub\*: An R/Bioconductor ExperimentHub package containing useful data resources for R-loop data analysis.
   Bioconductor page 
   Source code

- RLPipes\*: An CLI/Bioconda package for upstream processing of R-loop data. 
  Bioconda page 
  Source code
- **onto-lib**: A python library for loading, querying, and processing biomedical ontologies from OBO files. **O** Source code
- Correlation Analyzer\*: An R-Shiny app for exploration of gene co-expression correlation data. ( Web app Source code
- correlationAnalyzeR\*: An R package for exploration and analysis of gene co-expression correlation data.
   Wignette 
   Source code
- Miller et al., bioRxiv 2021\*: Reproducible analysis from preprint, *Miller et al., bioRxiv 2021.* () Study link () Source code

#### Research skills, tools, techniques

\*Bold indicates comprehensive skill and knowledge of a tool, data type, or technique.

# Analysis techniques

**Transcriptomics:** Differential gene expression, differential transcript usage, differential splicing, *de novo* isoform prediction, pathway enrichment, GSEA, pausing index prediction, fusion-gene detection, transposon analysis, transcriptomic alignment, splice-aware genomic alignment, RNA-Seq data mining. <u>scRNA-Seq analysis</u>: upstream processing, scRNA-Seq data mining, **quality control**, integration, cell type prediction, **dimensionality reduction**, imputation, gene module scoring, trajectory analysis, differential expression analysis.

**Epigenomics:** Genomic alignment, **peak calling**, coverage analysis, **differential binding analysis**, differential accessibility analysis, motif analysis, **peak location meta-analysis and enrichment testing**, data mining, signal track correlation and clustering analysis, building custom UCSC browser sessions and track hubs. <u>scATAC-Seq analysis</u>: quality control, integration with scRNA-Seq, dimensionality reduction, differential accessibility.

Other bioinformatics techniques: Integrative multiomics data analyses, GWAS analysis, mutation/CNV calling, mass-spectrometry analysis, nanostring analysis, survival analysis.

Other data sci techniques: Dimensionality reduction, clustering, imputation, data viz, interactive data reporting, classification models, regression models, regular expressions.

# Data types and technologies

Languages: R, python, bash, markdown, YAML, HTML, CSS (minor), JS (minor), LaTeX (minor).

Data types: mRNA-Seq, lncRNA-Seq (ribo-depleted), sRNA-Seq, scRNA-Seq (10X, fluidigm c1), ATAC-Seq, scATAC-Seq, HiC-Seq, ChIP-Seq (and variants), WGS, RRBS, WGBS, DRIP-Seq (and variants), R-ChIP (and variants), microarray, bulk mass spectrometry, CyTOF, nanostring.

Bioinformatics technologies/tools: <u>R/Bioconductor packages</u>: DESeq2, EdgeR, limma, seurat, single-cell experiment, SGSeq, DiffBind, phateR, Rmagic, ChIPpeakAnno, ChIPseeker, GenomicRanges/GenomicFeatures, valr, clusterProfiler, fgsea, msigdbr, annotationDBI (and related packages), BiomaRt, ExperimentHub/AnnotationHub (usage and development of a hub), rtracklayer, tximport, SummarizedExperiment, SingleR, monocle, SRAdb, Biostrings, enrichR, flowCore, groHMM, nanostringr, RUVSeq, PLIER, regioneR, recount, Rsubread, Bioconductor (general usage & package development). <u>Command Line</u>: SRA Toolkit, bedtools, bioconda (general usage & package development), samtools, bwa, STAR, salmon, HISAT2, StringTie, MACS2/3, EPIC2, deepTools, Homer, MEME-Suite, TEtoolkit, fastp. <u>GUI/Web</u>: UCSC Table/Genome Browser, IGV, GEO, SRA, DAVID, enrichr, StringDB, cbioportal, GEPIA2.

Data sci technologies/tools: R Markdown, Rhtml, Jupyter Notebook, conda, R-tidyverse (dplyr, ggplot2, etc), python-Saucie, python-pandas, python-numpy, python-sklearn, python-keras, snakemake, R-caret, R-pheatmap, R-ComplexHeatmap, plotly, R-DBI, R-DT, R-kableExtra, R-rstatix, R-Rtsne, R-uwot, R-SuperExactTest, R-xgboost.

Other computational and web technologies/tools: git, GitHub, CI/CD (GitHub actions), Linux server admin, NGINX,

Wordpress, AWS (EC2, S3, RDS, DynamoDB, IAM), **R-shiny**, python-dash, python-flask, python-sqlalchemy, python-pytest, python-coverage python-click, R-devtools, R-parallel, R-future, R-httr, R-jsonlite, R-lintr, R-callr, R-renv.

# **Bench techniques**

Molecular biology: Cell culture, Western blot, plasmid prep, protein extraction, RNA extraction, Bradford assay, ELISA, exosome isolation, transfection, nucleofection, nanodrop analysis.

**Biomedical engineering**: Liposome formulation, liposome extrusion, nanosight analysis, UV Vis, Autodesk CAD/Inventor, stereolithography/extrusion <sub>3</sub>D printing, PDMS-based microfluidics.

Animal techniques: Neural cannulation, stereotaxis surgery, animal post-op care, perfusion, oral gavage, tissue histology.