BIOGRAPHICAL SKETCH

Provide the following information for the Senior/key personnel and other significant contributors. Follow this format for each person. **DO NOT EXCEED FIVE PAGES.**

NAME: Austin Minton

eRA COMMONS USER NAME (credential, e.g., agency login): AUSTIN.MINTON

POSITION TITLE: Graduate Student

EDUCATION/TRAINING (Begin with baccalaureate or other initial professional education, such as nursing, include postdoctoral training and residency training if applicable. Add/delete rows as necessary.)

| INSTITUTION AND LOCATION | DEGREE (if applicable) | Start Date MM/YYYY | Completion Date MM/YYYY | FIELD OF STUDY |
|--|---------------------------|-----------------------|-------------------------|-------------------|
| Kentucky Wesleyan College, Owensboro, Kentucky | BS (Hons) | 08/2018 | 04/2022 | Chemistry |
| Kentucky Wesleyan College, Owensboro, Kentucky | BS (Hons) | 08/2018 | 04/2022 | Biology |
| University of Kentucky, Lexington, Kentucky | PHD | 08/2022 | 05/2027 | Physiology |

A. Personal Statement

The mystery surrounding my sister's neuromuscular disorder first sparked my interest in translational science. Her geneticist identified an abnormality in chromosome 6 but could not determine how it contributed to her condition. This uncertainty—how genetic changes manifest in disease—ignited my passion for biomedical research. During my career, I hope to be able to help transform genealogical discoveries into clinical actions.

I quickly realized that accessing research opportunities was not going to be easy. I grew up in rural western Kentucky without access to resources that many of my PhD classmates take for granted. I was very fortunate to be able to attend Kentucky Wesleyan College as an undergraduate, but it is a small liberal arts institution with limited infrastructure for research. However, I persisted and secured an opportunity in the lab of Rachel Pritchard, PhD, where I studied soil-derived antibiotics as tools to combat the growing crisis of drug-resistant pathogens. I used preliminary data from this project to build a successful application for the Wesleyan Fellowship, which funded 2 years of research. I disseminated first-author works at various local, national, and international conferences.

After admission into a PhD program at the University of Kentucky, I joined the lab of my primary sponsor, Ken Campbell, PhD where I was given the opportunity to shift my focus towards the genetics of human heart failure. Non-ischemic cardiomyopathies have a strong genetic component, yet nearly half of cases are idiopathic, much like my sister's condition. Despite this, genetic screening remains vastly underutilized in clinical settings. Our lab has spent 17 years building a cardiac biobank that now contains more than 20,000 specimens matched to clinical data. I spearheaded our lab's initiative to obtain sequencing data for 350 patients, presenting findings at numerous conferences and laying the foundation for a multi-omic atlas of heart failure patients in the greater Bluegrass region.

My PhD project centers on titin-truncating variants (TTNtvs), which cause premature protein translation stoppage and are amongst the most prevalent genetic contributors to non-ischemic cardiomyopathy. However, how TTNtvs lead to pathophysiology remains unclear. My project aims to address this gap by investigating if and how TTNtvs lead to overloaded cellular turnover pathways, accelerated aggregation of cytosolic residuals, and truncated titin filaments in sarcomeres. By integrating genomic, (immuno)histological, and biomechanical analyses, I aim to uncover novel therapeutic targets that will advance current treatments towards proactive, genetics-informed interventions.

Under the mentorship of Dr. Campbell, I will use my previous experiences in drug discovery and scientific communication as a conduit to provide insight into the pathophysiological underpinnings of heart failure. My training will be a first-hand perspective of the interconnectedness between research and medicine. I believe that this, including my previous research experiences, will provide a solid foothold for my long-term goal of leading a research team in cardiovascular genetics.

- a. **Minton AT**, Wellette-Hunsucker AG, Gulbulak U, Milburn GN, Yackzan AT, Campbell KS. Multi-Omic and Biochemical Profiling of Heart Failure Specimens at the University of Kentucky. University of Kentucky Center for Clinical and Translational Research Spring Conference. 2025 (Podium Talk)
- b. **Minton AT**, Wellette-Hunsucker AG, Gulbulak U, Milburn GN, Yackzan AT, Campbell KS. Genomic and Biochemical Profiling of Heart Failure at the University of Kentucky. Biophysical Society Annual Meeting. 2025 (Poster)
- c. **Minton AT**, Yackzan AT, Wellette-Hunsucker AG, Milburn GN, Gulbulak U, Campbell KS. Genomic Characterization of Patients with Advanced Heart Failure at the University of Kentucky. Madison Myofilament Meeting. 2024 (Poster)
- d. Cortazar AS, **Minton AT**, Gulbulak U, Campbell KS. Whole Exome Sequencing of a Myocardial Repository at the University of Kentucky. National Institutes of Health STEP-UP Program Annual Conference. 2024 (Podium Talk)

B. Positions, Scientific Appointments and Honors

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|---------------|---|
| | Scientific Appointments |
| 2024 – | Member, American Heart Association |
| 2024 – | Member, Biophysical Society |
| 2023 – | Graduate Research Assistant, Dr. Kenneth Campbell's Laboratory, Dept of Physiology, |
| | University of Kentucky College of Medicine |
| 2020 - 2022 | Laboratory Assistant, Div of Nat Sciences & Mathematics, Kentucky Wesleyan College |
| 2020 - 2022 | Peer Tutor & Instructor, Student Success Center, Kentucky Wesleyan College |
| 2020 - 2022 | Member, American Society for Microbiology |
| 2019 - 2022 | Directed Researcher, Div of Nat Sciences & Mathematics, Kentucky Wesleyan College |
| <u>Honors</u> | |
| 2023 | Featured in Fall 2023 Issue of <i>Pillars</i> as GOLD Alumnus, institution's alumni magazine, |
| 2020 | Kentucky Wesleyan College |
| 2022 | Invited Guest Speaker, STEM Bridge Program, Kentucky Wesleyan College |
| 2018 – 2022 | Presidential Scholarship, partial tuition scholarship, Kentucky Wesleyan College |
| 2020 – 2022 | Ellie Magnuson Memorial Endowed Fellowship Scholarship, awarded to selected |
| 2020 – 2022 | |
| 2020 2022 | researchers majoring in chemistry, Kentucky Wesleyan College |
| 2020 – 2022 | Wesleyan Fellowship, awarded to selected researchers, Kentucky Wesleyan College |
| 2020 – 2022 | Dean's List , ≥3.5 semester grade point average, Kentucky Wesleyan College |
| 2022 | President's Award, awarded to a selected Greek Life member, Kentucky Wesleyan College |
| 2022 | Program of the Year Award, awarded to a selected organization leader who hosted a |
| | successful community-wide program, Kentucky Wesleyan College |
| 2022 | Fraternity and Sorority Life Hall of Fame, awarded to selected Greek Life members, |
| | Kentucky Wesleyan College |
| 2022 | Student Government Association Senator of the Year, Kentucky Wesleyan College |
| 2022 | Order of Oak & Ivy Nominee, institution's highest honor, Kentucky Wesleyan College |
| 2022 | Chemistry Alumni Award, awarded to a selected student majoring in chemistry, Kentucky |
| | Wesleyan College |
| 2022 | Henry Milton Pyles Biology Award, awarded to a selected student majoring in biology, |
| | Kentucky Wesleyan College |
| 2021 | Interviewed on Bench Talk: The Week in Science, selected based on conference |
| | presentation award, Kentucky Academy of Science |
| 2021 | Philip R. Edwards Microbiology Award, awarded to a selected researcher in microbiology, |
| | Kentucky Wesleyan College |
| 2021 | Oral Presentation Award Winner, Kentucky Academy of Science, Eastern Kentucky |
| | University |
| 2021 | Fraternity Man of the Year, awarded to a selected fraternity member, Kentucky Wesleyan |
| | College |
| 2021 | Dr. Ernest W. Abernathy Scholarship , awarded to selected students majoring in chemistry |
| ZUZ I | or biology, Kentucky Wesleyan College |
| 2020 | Oral Presentation Award Winner, American Society for Microbiology, Vanderbilt University |
| 2020 | Ciai resentation Award Willier, American Society for Microbiology, Value bit Offiversity |

Presidential Scholarship, partial tuition scholarship, Kentucky Wesleyan College

C. Contributions to Science

2018

1. Genetic Variants in Heart Failure

Approximately 50% of patients with heart failure receive an idiopathic diagnosis. Moreover, animal models of many types of heart failure are nonrepresentative due to comorbidities such as hypertension, diabetes, and chronic lung diseases. There is a poor understanding of the link between heart failure and genetics, primarily due to the lack of genetic testing in this patient population. With Dr. Kenneth Campbell, I selected an experimental kit necessary to extract and purify nucleic acid eluants from specimens within the lab's myocardial repository. I used the extraction kit to derive a high-throughput protocol of extracting DNA and RNA from cryopreserved cardiac samples, which was utilized to extract DNA and RNA from 394 specimens. I coordinated with numerous genomics companies to determine the best sequencing platform, depth, coverage, and enrichment system to identify causal variants. 350 samples were sent for library preparation and whole exome/transcriptome sequencing. The collected data provided a genetic atlas of specimens within the myocardial repository, representative of heart transplant and ventricular assist device recipients in the greater Bluegrass region. Further analyses revealed trends in sequencing results and matched clinical data. Moreover, this dataset fostered research collaborations nationally and internationally.

- a. **Minton AT**, Wellette-Hunsucker AG, Gulbulak U, Milburn GN, Yackzan AT, Campbell KS. Multi-Omic and Biochemical Profiling of Heart Failure Specimens at the University of Kentucky. University of Kentucky Center for Clinical and Translational Research Spring Conference. 2025 (Podium Talk)
- b. **Minton AT**, Wellette-Hunsucker AG, Gulbulak U, Milburn GN, Yackzan AT, Campbell KS. Genomic and Biochemical Profiling of Heart Failure at the University of Kentucky. Biophysical Society Annual Meeting. 2025 (Poster)
- c. **Minton AT**, Yackzan AT, Wellette-Hunsucker AG, Milburn GN, Gulbulak U, Campbell KS. Genomic Characterization of Patients with Advanced Heart Failure at the University of Kentucky. Madison Myofilament Meeting. 2024 (Poster)
- d. Cortazar AS, **Minton AT**, Gulbulak U, Campbell KS. Whole Exome Sequencing of a Myocardial Repository at the University of Kentucky. National Institutes of Health STEP-UP Program Annual Meeting. 2024 (Podium Talk)

2. Contribution of Variants in the Titin Gene to the Pathology of Dilated Cardiomyopathy

The Campbell Lab maintains and utilizes tissue from one of the world's largest human cardiac biobanks to perform cardiovascular research. Experimentation ranges from the single-myofibril to whole-organ level, providing insight applicable at the bench and the clinic. I collated clinical and whole exome sequencing data to identify patients who met dilated cardiomyopathy criteria and contained variants in the *TTN* gene. Using samples from these patients, I assisted in evaluating phosphorylation of proteins involved in myofilament calcium sensitivity (regulatory light chain, troponin I, and myosin-binding protein C) and relative abundances of contributors to intra/extracellular passive tension (collagen, alpha-tubulin, and titin). Findings differed from those previously collected by our lab, which included patients with truncating *TTN* variants (irrespective of dilated cardiomyopathy diagnosis). This hinted towards possible associations with the location of a genomic variant. To enable such comparisons, I mapped the exonic location of *TTN* variants based on the corresponding region of the sarcomere. Collected data has supported several conference presentations and serves as the basis of my dissertation research.

- a. **Minton AT**, Campbell KS. Effects of SGLT2i Treatment in Patients with Cardiac Titin Variants. University of Kentucky College of Medicine Department of Physiology Seminar Series: Trainee Talk. 2024 (Podium Talk)
- b. Wilkerson E, **Minton AT**, Wellette-Hunsucker AG, Gulbulak U, Campbell KS. Evaluating TTN Variants in Dilated Cardiomyopathy at the University of Kentucky. Kentucky Chapter of the American Physiological Society Annual Meeting. 2024 (Poster)

3. Production of Novel Antibiotics from Soil Bacteria

Bacteria are becoming increasingly more resistant to commercially available antibiotics, leading to difficulty treating infections that were once subjective to such medications. Since antibiotics are commonly produced in bacteria inhabiting soil, this serves as a natural reservoir to identify and isolate novel antimicrobials. In coordination with Dr. Rachel Pritchard, I served as the lead investigator on a project that explored the ability to discover novel antibiotics from soil samples of various demographics. I revealed antibiotic production from thirteen bacterial isolates of four soil samples and assisted in optimizing an experimental technique to extract the antimicrobial compounds. Novelty of the bacteria was confirmed with 16S rRNA gene PCR, Sanger sequencing, and advanced biochemical testing. I cultivated stocks that were sent to the Tiny Earth Chemistry

Hub, a public database that preserves samples and records all experimental conditions, for use in further experimentation and possible application.

- a. **Minton AT**, Pritchard R. Analysis of Purified Extracts from Antibiotic-Producing Bacterial Isolates. Kentucky Academy of Science Annual Meeting. 2021 (Podium Talk Award Winner)
- b. **Minton AT**, Pritchard R. Analysis of Bacterial Isolates Found in the Soil: Executing the Tiny Earth Project. Kentucky-Tennessee American Society for Microbiology Meeting. 2020 (Podium Talk Award Winner)
- c. **Minton AT**, Pritchard R. Analysis of Antibiotic-Producing Bacterial Isolates: Executing the Tiny Earth Project. Kentucky Wesleyan College Scholar's Day. 2021 (Poster)
- d. **Minton AT**, Pritchard R. Analysis of Bacterial Isolates Found in the Soil: Executing the Tiny Earth Project. Tiny Earth Winter Symposium. 2020 (Podium Talk)

D. Scholastic Performance

| YEAR | COURSE TITLE | GRADE |
|------|--|-------|
| | KENTUCKY WESLEYAN COLLEGE | |
| 2016 | Fundamentals of General Chemistry | Р |
| 2016 | Fundamentals of General Chemistry Lab | Р |
| 2017 | Fundamentals of Organic Chemistry | Р |
| 2017 | Fundamentals of Organic Chemistry Lab | Р |
| 2017 | Medical Terminology from Greek & Latin | Р |
| 2017 | Writing I | Р |
| 2017 | Music Appreciation | Р |
| 2018 | Calculus AB | Р |
| 2018 | English II | Р |
| 2018 | General Biology I Lab | Α |
| 2018 | General Biology I | C+ |
| 2018 | General Chemistry Laboratory I | Α |
| 2018 | General Chemistry I | B+ |
| 2018 | Freshman Seminar | Α |
| 2018 | Introduction to Religion | Α |
| 2019 | General Biology II Lab | Α |
| 2019 | General Biology II | C+ |
| 2019 | General Chemistry Laboratory II | A- |
| 2019 | General Chemistry II | B+ |
| 2019 | Introduction to Psychology | Α |
| 2019 | Survey of Christian Traditions | Α |
| 2019 | Microbiology I | В |
| 2019 | Organic Chemistry Laboratory I | Α |
| 2019 | Organic Chemistry I | A- |
| 2019 | Analytical Chemistry | B- |
| 2019 | American Literature Survey | Α |
| 2020 | Genetics | A- |
| 2020 | Directed Student Research | Α |
| 2020 | Organic Chemistry Laboratory II | B+ |
| 2020 | Organic Chemistry | A- |
| 2020 | Fitness and Wellness | Α |
| 2020 | Directed Student Research | Α |
| 2020 | Statistics in the Behavioral Sciences | Α |
| 2020 | Natural Science Junior Seminar | Α |
| 2020 | Principles of Sociology | Α |
| 2020 | General Physics | Р |
| 2020 | College Physics Laboratory | Р |

| 2021 | Cellular/Molecular Biology | B+ |
|------|--|----|
| 2021 | Directed Student Research | Α |
| 2021 | Immunology | A- |
| 2021 | Inorganic Chemistry | Α |
| 2021 | Biochemistry | Α |
| 2021 | Introductory General Physics II | Α |
| 2021 | Introductory General Physics II Laboratory | Α |
| 2021 | Biology of the Mind | В |
| 2021 | Directed Student Research | Α |
| 2021 | Senior Seminar | Α |
| 2021 | Advanced Integrated Lab I | Α |
| 2021 | Computer Literacy | Р |
| 2021 | Introduction to Human Geography | Α |
| 2021 | Survey of American History I | Α |
| 2021 | Evolution | Α |
| 2022 | Physiological Psychology | Α |
| 2022 | Directed Student Research | Α |
| 2022 | Investigations in Molecular Cell Biology | Α |
| 2022 | Ecology | A- |
| 2022 | Instrumental Techniques of Biochemistry | В |
| 2022 | Advanced Integrated Lab II | A |
| | UNIVERSITY OF KENTUCKY | |
| 2022 | Biomolecules and Metabolism | В |
| 2022 | Molecular Biology and Genetics | В |
| 2022 | Seminar in Integrated Biomedical Sciences | S |
| 2022 | Research in Integrated Biomedical Sciences | Α |
| 2022 | Critical Scientific Readings | Α |
| 2022 | Practical Statistics | Α |
| 2023 | Ethics in Scientific Research | Α |
| 2023 | Cell Biology and Signaling | В |
| 2023 | Physiological Communication | Α |
| 2023 | Seminar in Integrated Biomedical Sciences | S |
| 2023 | Research in Integrated Biomedical Sciences | Α |
| 2023 | Genomics & Bioinformatics Tools | Α |
| | Joined the Campbell Muscle Lab | |
| 2023 | Systems, Cellular & Molecular Physiology | Α |
| 2023 | Graduate Seminar in Physiology | Α |
| 2023 | Readings in Systems, Cellular and Molecular Physiology | Α |
| 2024 | Fellowship Grant Writing Workshop | Α |
| 2024 | Advanced Topics in Physiology | Α |
| 2024 | Research in Physiology | Α |
| 2024 | Graduate Seminar in Physiology | A |
| 2024 | Qualifying Exam Residency Credit | Р |
| | | |

^{*}Kentucky Wesleyan College Grading System: Pass (≥70%), Fail (<70%); A (100-93%), A- (93-90%), B+ (89-87%), B (86-83%), B- (82-80%), C+ (79-77%), C (76-73%), C- (72-70%), D+ (69-67%), D (66-63%), D- (62-60%), F (<60%)

^{**&}lt;u>University of Kentucky Grading System</u>: Satisfactory (S; ≥70%), Non-Satisfactory (NS: <70%); A (100-90%), B (89-80%), C (79-70%), D (69-60%), F (<60%)