

Madeleine S. Gastonguay

(860) 578-7177 | mgaston1@jh.edu

<https://madeleine-gastonguay.netlify.app/> | ORCID: [0000-0002-5700-8543](https://orcid.org/0000-0002-5700-8543)

Education

Johns Hopkins School of Medicine, Baltimore, MD

August 2022 - present

PhD in Biomedical Engineering

Advisor: Dr. Alison L. Hill

University of Connecticut, Storrs, CT

August 2016 - May 2020

Bachelor of Science, Applied Mathematics

Summa Cum Laude with Honors; GPA: 3.98/4.00

Minor: Bioinformatics

Thesis: A Quantitative Pipeline for The Identification of Combinations of Targets for Claudin-Low Triple Negative Breast Cancer Reversion

Advisor: Dr. Paola Vera-Licona

La Sorbonne University, Paris, France

Jan. 2018 - May 2018

Course de Civilisation Française

Research Experience

Johns Hopkins Institute for Computational Medicine

August 2022 – present

Biomedical Engineering PhD Candidate

Topic: Integrating mathematical models and statistical methods for within-host viral dynamics with quantitative systems pharmacology models of novel immunotherapies

- Building mathematical models and evaluating their dynamic properties
- Developing Bayesian methods for parameter estimation and uncertainty quantification
- Implementing nonlinear-mixed effects modeling to quantify interindividual variability and understand therapeutic mechanisms of action
- Designing stochastic simulations to predict long-term behavior of the system and evaluate therapeutic interventions
- Collaborating with experimentalists in the field of infectious diseases

The Jackson Laboratory (JAX), Bar Harbor, ME

June 2020 – July 2022

Research Data Analyst I

Topic: A Bayesian approach to mediation analysis of complex traits with measurement noise

- Contributed to the development and validation of an R package implementation of a Bayesian model selection approach to mediation analysis that is flexible in both data inputs and potential inferences, and extended it to moderated mediation
- Diagnosed the effect of measurement noise on the inference of mediation
- Applied these tools to better understand the mechanisms underlying the effects of sex and diet on protein and gene expression in the livers of genetically diverse mice

UConn Health Center for Quantitative Medicine, Farmington, CT

Sept. 2018 - May 2020

Undergraduate Research Assistant

Topic: A quantitative pipeline for cancer reversion analysis in triple negative breast cancer

- Constructed a static intracellular signaling network for a claudin-low triple negative breast cancer (CL TNBC) cell line with multi-omics data using bioinformatics techniques
- Applied a structure-based control method for nonlinear systems, implemented in python, to identify putative targets that steer the system to any desired attractor
- Conducted virtual screenings using a network-based approach and a signal propagation algorithm to estimate long term behaviors to identify concerted perturbations of control nodes resulting in reversion of the CL TNBC phenotype

Metrum Research Group, Simsbury, CT

June 2018 - Aug. 2018

Summer Intern

Topic: A maternal-fetal physiologically based pharmacokinetic model for drugs metabolized by cytochrome P450 isoenzymes

- Modeled maternal and fetal drug exposures at varying gestational ages by incorporating anatomical, biochemical, and physiological changes associated with pregnancy as a system of ordinary differential equations using R and *mrgsolve*
- Performed local sensitivity analysis, optimized model parameters, and validated the model by comparing model predictions to external published data

Topic: The effect of host genetic variability on Epstein Barr Virus (EBV)-associated cancer susceptibility

- Reviewed relevant literature to identify genes that may impact EBV-associated cancer susceptibility
- Used wet lab techniques such as Polymerase Chain Reactions, Gel Electrophoresis, and Sanger Sequencing to sequence the genes of interest in several EBV-associated cancer cell lines and a non-cancerous EBV+ control cell line
- Aligned the genetic sequence of target genes to identify common and distinct single nucleotide polymorphisms (SNPs) across EBV-associated cancer cell lines using the software Geneious and BLAST

Children's Hospital of Philadelphia, Philadelphia, PA, Metrum Research Group, Simsbury, CT

Sept. 2015 - Aug. 2016

High School Independent Study

Topic: A pharmacogenomic study of midazolam and morphine clearance in critically ill pediatric patients

- Analyzed variability of SNPs in the sample dataset for inclusion in the analysis
- Estimated the effect of pediatric risk of mortality score on drug clearance using population pharmacokinetic modeling
- Identified a SNP in UGT2B7 whose minor allele frequency is associated with increased midazolam clearance

Publications and Preprints

Gastonguay, M. S., Keele, G. R., & Churchill, G. A. (2023). The trouble with triples: Examining the impact of measurement error in mediation analysis. *Genetics*, 224(1), iyad045. <https://doi.org/10.1093/genetics/iyad045>

Crouse, W. L., Keele, G. R., Gastonguay, M. S., Churchill, G. A., & Valdar, W. (2022). A Bayesian model selection approach to mediation analysis. *PLOS Genetics*, 18(5), e1010184. <https://doi.org/10.1371/journal.pgen.1010184>

Utsey, K., Gastonguay, M. S., Russell, S., Freling, R., Riggs, M. M., & Elmokadem, A. (2020). Quantification of the Impact of Partition Coefficient Prediction Methods on Physiologically Based Pharmacokinetic Model Output Using a Standardized Tissue Composition. *Drug Metabolism and Disposition*, 48(10), 903 LP – 916. <https://doi.org/10.1124/dmd.120.090498>

Zuppa, A. F., Benitez, G. R., Zane, N. R., Curley, M. A. Q., Bradfield, J., Hakonarson, H., Gastonguay, M. S., Moorthy, G., Prodell, J., & Gastonguay, M. R. (2019). Morphine Dose Optimization in Critically Ill Pediatric Patients With Acute Respiratory Failure. *Critical Care Medicine*, 47(6), e485–e494. <https://doi.org/10.1097/CCM.00000000000003741>

Zuppa, A. F., Conrado, D. J., Zane, N. R., Curley, M. A. Q., Bradfield, J., Hakonarson, H., Gastonguay, M. S., Moorthy, G., Prodell, J., & Gastonguay, M. R. (2019). Midazolam Dose Optimization in Critically Ill Pediatric Patients With Acute Respiratory Failure. *Critical Care Medicine*, 47(4), e301–e309. <https://doi.org/10.1097/CCM.00000000000003638>

Oral Presentations

Gastonguay MS, Gerold J, Juillard F, Szymula A, Hill AL, Kaye K, *Quantifying the dynamics of Kaposi's sarcoma-associated herpesvirus persistence*, Epidemics, Bologna, Italy, November 27th, 2023

Gastonguay MS, Russell S, Freling R, Utsey K, and Elmokadem A, *Prediction of maternal-fetal exposures of CYP450-metabolized drugs using physiologic pharmacokinetic modeling implemented in R and mrgsolve.*, R/Pharma Conference, Cambridge, MA, August 23rd, 2019

Gastonguay MS, Marazzi L, Vera-Licona P, *Identification of Combinations of Targets for Claudin-Low Triple Negative Breast Cancer Reversion*, UConn Center for Quantitative Medicine, July 30th, 2019

Gastonguay MS, Marazzi L, Vera-Licona P, *Identification of Combinations of Targets for Claudin-Low Triple Negative Breast Cancer Reversion*, UConn Center for Cell Analysis and Modeling Summer Seminar, July 26th, 2019

Gastonguay MS, Russell S, Freling R, Utsey K, and Elmokadem A, *Development of an Open and General Physiologically Based Pharmacokinetic Model to Predict Maternal-Fetal Exposures for Drugs Metabolized by CYP Isoenzymes*, R/Medicine Conference, New Haven, CT, September 8th, 2018

Gastonguay MS, *The Effect of Host Genetic Variability on Epstein Barr Virus-associated cancer susceptibility*, UConn Holster Scholar Symposium, October 2017

Poster Presentations

Gastonguay MS, Gerold J, Juillard F, Szymula A, Hill AL, Kaye K, *Quantifying the dynamics of Kaposi's sarcoma-associated herpesvirus persistence*, MIDAS Network Annual Meeting, Bethesda, MD, November 18th, 2024

Gastonguay MS, Gerold J, Juillard F, Szymula A, Hill AL, Kaye K, *Quantifying the dynamics of Kaposi's sarcoma-associated herpesvirus persistence*, International Workshop on Kaposi's Sarcoma Herpesvirus (KSHV) & Related Agents, Boston, MA, July 7th, 2024

Gastonguay MS, Marazzi L, Vera-Licona P, *Identification of Combinations of Pharmacologic Targets for Claudin-Low Triple Negative Breast Cancer Reversion*, International Society of Pharmacometrics Quantitative Systems Pharmacology Student Symposium, Virtual, April 28th, 2021

Gastonguay MS, Marazzi L, Vera-Licona P, *Identification of Combinations of Targets for Claudin-Low Triple Negative Breast Cancer Reversion*, Joint Meeting in Mathematics, Denver, CO, January 15th – 18th, 2020

Gastonguay MS, Russell S, Freling R, Utsey K, and Elmokadem A, *Development of an Open-source Physiologically-Based Pharmacokinetic Model to Predict Maternal-Fetal Exposures of CYP450-Metabolized Drugs*, International Society of Pharmacometrics Regional Quantitative Systems Pharmacology Day, Princeton, NJ, July 16th, 2019

Gastonguay MS, Russell S, Freling R, Utsey K, and Elmokadem A, *Development of an Open-source Physiologically-Based Pharmacokinetic Model to Predict Maternal-Fetal Exposures of CYP450-Metabolized Drugs*, University of Connecticut Frontiers in Undergraduate Research, April 12th, 2019

Fellowships and Grants

Summer Undergraduate Research Fund , University of Connecticut Office of Undergraduate Research	May 2019
<ul style="list-style-type: none"> Awarded funding for a 9-week summer research project 	
Holster Scholar , University of Connecticut Honors Program	May 2017
<ul style="list-style-type: none"> Offered enrollment in a semester-long course to learn how to develop and write a project proposal Selected as one of 8 students awarded funding for a 10-week summer research project 	

Honors and Awards

Blue Ribbon Poster Award , ISoP Quantitative Systems Pharmacology Student Symposium	April 2021
Dean's List , The University of Connecticut	Sept. 2016 - May 2020
Academic Excellence Scholarship , The University of Connecticut	Sept. 2016 - May 2020
Babbidge Scholar , The University of Connecticut	Dec. 2017, 2019
New England Scholar , The University of Connecticut	Dec. 2018
Global Citizenship Scholarship , The University of Connecticut Education Abroad	Jan. 2017

Skills & Certifications

Technical: R; Python; Matlab, SQL, Julia, and bash shell; Git; LaTeX; OpenRefine; High Performance Computing with SLURM; Nonlinear mixed-effects modeling with Monolix; *mrgsolve*; *shiny*; *tidyverse*; *plotly*; JAGS; Bayesian Data Analysis
Wet Lab: Polymerase Chain Reactions, Gel Electrophoresis, Gel Extraction, Sanger sequencing
Language: Proficient in French conversation, reading, and writing; Certified in French level B1.2 by La Sorbonne in Paris
Teaching: Certified Instructor with The Carpentries

Teaching Experience

Dynamic modeling of infectious diseases in patients and populations , Johns Hopkins University (TA)	Oct. - Dec. 2024
Systems pharmacology and personalized medicine , Johns Hopkins University (TA)	Feb. - May 2024
Data carpentry ecology with R workshop , Bioinformatics Training Program at JAX (Instructor)	Dec. 2021
Data carpentry genomics workshop , Bioinformatics Training Program at JAX (Instructor)	Nov. 2021
Introductory statistics with R , Bioinformatics Training Program at JAX (TA)	Sept. 2021
Introduction to R and RStudio , Bioinformatics Training Program at JAX (TA)	June 2021

Professional Development and Continuing Education

Summer Institute in Statistics and Modeling in Infectious Diseases , Emory University	July 2024
Building Tidy R Packages , R/pharma 2021	Oct. 2021
Julia Language for R Programmers , R/pharma 2021	Oct. 2021
Advanced Shiny , RStudio	Sept. 2021
Introduction to Bayesian Data Analysis , Juliacon	July 2021
Carpentries Instructor Training , The Carpentries	Mar. 2021
Shiny, RMarkdown, and RStudio Connect , RStudio	Mar. 2021
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Quantitative Trait Mapping in the Diversity Outbred , University of Wisconsin-Madison	Dec. 2020
Containerization with Singularity , JAX	Oct. 2020
Introduction to HPC , JAX	Sept. 2020
Human and Mammalian Genetics and Genomics: The 61st McKusick Short Course , JAX	July 2020
Shiny Reproducibility , R/pharma 2019	Aug. 2019
Machine Learning , R/pharma 2019	Aug. 2019

Volunteer Work and Extra-Curricular Activities

Rubyfruit A Cappella , The University of Connecticut	Sept. 2016 - May 2020
<ul style="list-style-type: none"> Elected as President (2019-2020), Assistant Music Director (2018 - 2019), and Treasurer (2017- 2018) Coordinated funding, schedules, and rehearsals to record an album and distribute it on Spotify and Apple Music Communicated with other executive board members to run productive fundraisers, rehearsals, and gigs 	
Math Motivators , The University of Connecticut	Oct. 2017 - May 2019
<ul style="list-style-type: none"> Tutored high school freshmen from underprivileged schools in Hartford, CT 	
iGEM Genetic Engineering Team , The University of Connecticut	Sept. 2016 - Dec. 2017
<ul style="list-style-type: none"> Developed and presented a project proposal for the iGEM jamboree with a team of students 	