# Madeleine S. Gastonguay

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#### **Education**

## University of Connecticut, Storrs, CT

May 2020

Bachelor of Science, Applied Mathematics Summa Cum Laude with Honors in the Major

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

January 2018-May 2018

Course de Civilisation Française

## **Research Experience**

## The Jackson Laboratory (JAX), Bar Harbor, ME

June 2020 - present

Research Data Analyst I

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

- Contributing to the development and validation of an R package implementing a Bayesian model selection approach to
  mediation analysis that is flexible in both data inputs and possible inferences, and extending it to allow for inference of
  moderated mediation
- Diagnosing the effect of measurement noise in the inference of mediation
- Applying developed tools to identify the effect of sex and diet on mechanisms of protein expression in Diversity Outbred Mouse Liver

## UConn Health Center for Quantitative Medicine, Farmington, CT

September 2018 - May 2020

Undergraduate Research Assistant

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

- Awarded a Summer Undergraduate Research Fund through UConn to fund my work
- Constructed a static intracellular signaling network for a claudin-low triple negative breast cancer (CL TNBC) cell line with multi-omics data using Cytoscape and GeneXplain
- Applied a structure-based control method for nonlinear systems implemented in python to identify putative targets that can steer the system to any desired attractor
- Conducted virtual screenings using the topology of the network and a signal propagation algorithm to identify concerted perturbations of control targets resulting in reversion of the CL TNBC phenotype

## Metrum Research Group, Simsbury, CT

June - August 2018

Summer Intern

Topic: Developing an open and general maternal-fetal physiologically based pharmacokinetic model for drugs metabolized by cytochromes P450 isoenzymes

- Modeled maternal and fetal drug exposures at different gestational ages by incorporating anatomical, biochemical, and physiological changes associated with pregnancy as a system of 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

University of Connecticut, Department of Molecular and Cellular Biology, Storrs, CT

January - September 2017

Undergraduate Research Assistant/Holster Scholar

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

- Reviewed relevant literature to identify genes that may impact EBV-derived cancer susceptibility
- Utilized wet lab techniques such as Polymerase Chain Reactions, Gel Electrophoresis, Cloning, and Sanger Sequencing to sequence the genes of interest in several EBV-derived cancer cell lines
- Aligned the genetic sequence of target genes to identify common and distinct single nucleotide polymorphisms across EBVderived cancers using the software Geneious and BLAST

### **Publications and Pre-Prints**

Gastonguay, M. S., Keele, G. R., & Churchill, G. A., (2021). The impact of measurement noise on mediation analysis of complex traits. *Manuscript in preparation for submission* 

Crouse, W. L., Keele, G. R., Gastonguay, M. S., Churchill, G. A., & Valdar, W. (2021). A Bayesian model selection approach to mediation analysis. *BioRxiv*, 2021.07.19.452969. https://doi.org/10.1101/2021.07.19.452969, *under review at PLOS genetics* 

Utsey K, Gastonguay MS, Russell S, Freling R, Riggs MM, Elmokadem A. Quantification of the Impact of Partition Coefficient Prediction Methods on Physiologically Based Pharmacokinetic Model Output Using a Standardized Tissue Composition. *Drug Metab Dispos*. 2020;48(10):903 LP-916. doi:10.1124/dmd.120.090498

Zuppa AF, Benitez GR, Zane NR, Curley MAQ, Bradfield J, Hakonarson H, Gastonguay MS, Moorthy G, Prodell J, Gastonguay MR. Morphine Dose Optimization in Critically III Pediatric Patients With Acute Respiratory Failure: A Population Pharmacokinetic-Pharmacogenomic Study. Crit Care Med. 2019 Jun;47(6):e485-e494. doi: 10.1097/CCM.000000000003741. PMID: 30920410.

Zuppa AF, Conrado DJ, Zane NR, Curley MAQ, Bradfield J, Hakonarson H, Gastonguay MS, Moorthy G, Prodell J, Gastonguay MR. Midazolam Dose Optimization in Critically Ill Pediatric Patients With Acute Respiratory Failure: A Population Pharmacokinetic-Pharmacogenomic Study. Crit Care Med. 2019 Apr;47(4):e301-e309. doi: 10.1097/CCM.000000000003638. PMID: 30672747; PMCID: PMC6432942.

#### **Oral Presentations**

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 23<sup>rd</sup>, 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 30<sup>th</sup>, 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 26<sup>th</sup>, 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 8<sup>th</sup>, 2018

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

#### **Poster Presentations**

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, April 28<sup>th</sup>, 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 15<sup>th</sup> – 18<sup>th</sup>, 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 16<sup>th</sup>, 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<sup>2</sup> 2019

#### **Fellowships and Grants**

Summer Undergraduate Research Fund (SURF) Trimble Family Award
University of Connecticut Office of Undergraduate Research (\$4,000)

May 2019

Holster Scholar

May 2017

University of Connecticut Honors Program (\$4,000)

#### **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; basic Python, Matlab, SQL, Julia, and bash shell; Git; LaTeX; OpenRefine; High Performance Computing with SLURM; *mrgsolve*; *shiny*; *tidyverse*; 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

## **Professional Development and Continuing Education**

Attendee	
Advanced Shiny, Phil Bowsher with RStudio	Sept. 2021
Introduction to Bayesian Data Analysis, Juliacon Carpentries Instructor Training, The Carpentries	July 2021 Mar. 2021
Quantitative Trait Mapping in the Diversity Outbred, University of Wisconsin-Madison	Dec. 2020
Containerization with Singularity, JAX	Oct. 2020
Introduction to HPC, JAX	Sep. 2020
Human and Mammalian Genetics and Genomics: The 61st McKusick Short Course, JAX	July 2020
Shiny Reproducibility, Joe Cheng at R/pharma 2019	Aug. 2019
Machine Learning, Max Kuhn at R/pharma 2019	Aug. 2019
Instructor or Teaching Assistant	
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 Associations**

Society for Industrial and Applied Mathematics (SIAM) International Society of Pharmacometrics (ISoP)

American Statistics Association (ASA)

## Volunteer Work and Extra-Curricular Activities

Rubyfruit A Cappella, The University of Connecticut

Sept. 2016 - May 2020

- 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

• Tutored high school freshmen from underprivileged schools in Hartford, CT

iGEM Genetic Engineering Team, The University of Connecticut

Sept. 2016 - Dec. 2017

• Developed and presented a project proposal for the iGEM jamboree with a team of students

Husky for a Day Volunteer, The University of Connecticut

Sept. 2016 - May 2017

• Guided prospective students around campus and answered questions about UConn