The Faculty of Medicine of Harvard University **Curriculum Vitae**

Date Prepared: July 29, 2022

Name: Richard Allen Guyer

Office Address: General Surgery Residency Program

Massachusetts General Hospital

55 Fruit St, GRB-425 Boston, MA 02114

Home Address:

Work Phone: (908) 447-3462

Work Email: rguyer@mgh.harvard.edu

Education:

2002-2006	BA	Economics and Mathematics	Davidson College
2006-2008	No degree	Special Sciences Program	University of Pennsylvania
2008-2012	No degree	Medicine and Microbiology (Laboratory of Ian Macara)	University of Virginia
2012-2016	MD, PhD	Medicine and Cell and Developmental Biology (Laboratory of Ian Macara)	Vanderbilt University

Postdoctoral Training:

06/16-	Resident physician	General Surgery	Massachusetts General Hospital
07/19-	Research fellow	Pediatric Surgical Research Laboratories (Laboratory of Allan Goldstein)	Massachusetts General Hospital

Appointments at Hospitals/Affiliated Institutions:

2016	Resident	Surgery	Massachusetts General
			Hospital

2016-	Clinical Fellow	Surgery	Harvard Medical School
2016-	Resident	Surgery	Newton-Wellesley Hospital
2017-	Resident	Surgery	North Shore Medical Center

Major Administrative Leadership Positions:

Local

Academic Development Administrative 2021-Massachusetts General Hospital Resident

(Surgery)

Professional Societies:

2015-American College of Surgeons

2016-Association for Academic Surgery

2016-Massachusetts Medical Society

2017-American Pediatric Surgical Association

> 2020-2022 Resident Member, Research Committee

Grant Review Activities:

2020-Research Committee American Pediatric Surgical Association

> 2020-2022 Permanent Member

Editorial Activities:

Ad hoc Reviewer

Biomolecules

International Journal of Molecular Sciences

Stem Cells Translational Medicine

Computational and Structural Biotechnology Journal

Neural Regeneration Research

Other Editorial Roles

Award

Honors and Prizes:			
2004	Richard R. Bernard Society for Mathematics	Davidson College Department of Mathematics	Institutional undergraduate mathematics honor society
2005	Omicron Delta Epsilon Economics Honor Society	Davidson College Department of Economics	National undergraduate economics honor society
2007	Ruth and William Silen, MD Award	Harvard Medical School DICP	Best oral presentation at the 2007 New England Science Symposium
2009	Trevor Evans Award	Mathematical Association of America	For an exceptional article accessible to undergraduates published in Math Horizons
2011	NIH T32 Cancer Research Training Grant	University of Virginia	Research of breast cancer metastasis
2016	Medical Student Research Prize	Association for Academic Surgery/Association for Academic Surgery Foundation	Research of breast cancer metastasis conducted in Ian Macara's laboratory
2019	Marshall K. Barlett Research Fellowship	Massachusetts General Hospital Department of Surgery	Support for research in Allan Goldstein's laboratory
2020	National Research Service Award (F32) Postdoctoral Fellowship	National Institutes of Health	Support for research of the transcriptional control of postnatal enteric neurogenesis
2022	Patricia K. Donahoe, MD, Surgeon- Scientist Research Program Resident Catalyst	Massachusetts General Hospital Department of Surgery	Support for laboratory research during senior residency training

Report of Funded and Unfunded Projects

Past

2020-2022 Transcription Factors Mediate Postnatal Neurogenesis from Enteric Glial

Cells

NIH/NIDDK; F32 DK121440

PI (\$141,936)

This project utilizes single-cell RNA sequencing and conditional gene

knockout models to identify transcriptional drivers of postnatal neurogenesis

in the enteric nervous system.

Current

2022-2024 A Developmentally-Informed Approach to Neuroblastoma Biology

Massachusetts General Hospital Department of Surgery; Research Grant

PI (\$100,000)

This project applies both cutting-edge sequencing tools and traditional

methods to uncover the developmental and epigenetic basis for

neuroblastoma tumorigenesis and response to therapy

Report of Local Teaching and Training

Clinical Supervisory and Training Responsibilities:

2016- Daily supervision of HMS students on

the Surgery rotation of the Principle

Clinical Experience and on general surgery sub-internships

Harvard Medical School Students

Massachusetts General Hospital

20 hours / week

2017- Daily supervision of third-year Tufts

medical students undertaking their

surgical clerkship at Newton-Wellesley

Hospital

Tufts University Medical Students

Newton-Wellesley Hospital

20 hours / week

Local Invited Presentations:

 \boxtimes No presentations below were sponsored by 3^{rd} parties/outside entities

2018	Blunt Abdominal Trauma / Lecture to Tufts University medical students
	Department of Surgery, Newton-Wellesley Hospital
2018	Compartment Syndromes / Lecture to Tufts University medical students
	Department of Surgery, Newton-Wellesley Hospital
2019	Reanalysis of published single cell RNA sequencing of enteric nervous system cells / monthly departmental research seminar Department of Surgery, Massachusetts General Hospital
2020	Single cell transcriptomics of postnatal enteric glial cells reveals similarity with embryonic enteric neuronal progenitors / Weekly research conference
	Pediatric Surgical Research Laboratories, Massachusetts General Hospital
2020	Using Single-Cell Transcriptomics to Understand and Treat Hirschsprung Disease / Pediatric Surgery/Pediatric Anesthesia Grand Rounds
	Departments of Pediatric Surgery and Anesthesia, MassGeneral Hospital for Children
2021	A Developmentally-Informed Approach to Neuroblastoma Biology / Weekly research conference
	Pediatric Surgical Research Laboratories, Massachusetts General Hospital
2021	Single cell analysis of enteric glial cells identifies a molecular basis for postnatal enteric neurogenesis / Surgical Investigators' Club Department of Surgery, Massachusetts General Hospital
2022	Chromatin Remodeling and Cell Fate in the Neural Crest / Pediatric Surgery/Pediatric Anesthesia Grand Rounds Departments of Pediatric Surgery and Anesthesia, MassGeneral Hospital for Children
2022	Chromatin Remodeling and Cell Fate in the Neural Crest / Surgical Investigators' Club Department of Surgery, Massachusetts General Hospital

2022	Heterogeneity and plasticity in neuroblastoma cell lines / Weekly research conference
	Pediatric Surgical Research Laboratories, Massachusetts General Hospital
	-
2022	Management of Blunt Abdominal Trauma / Lecture to Tufts University medical students
	Department of Surgery, Newton-Wellesley Hospital
2022	Overcoming the Review – Getting a F32 Funded / Academic Development Career Seminar Series
	Department of Surgery, Massachusetts General Hospital
2022	Postoperative management / Lecture to incoming house officers Newton-Wellesley Hospital
2022	Presenting Patients on Surgical Rounds / Lecture to Tufts University medical students
	Department of Surgery, Newton-Wellesley Hospital

Report of Scholarship

ORCID: 0000-0001-5075-1767

Peer-Reviewed Scholarship in print or other media:

Research Investigations

- 1. Yu J, Ishii M, Law M, Woodburn JM, Emami K, Kadlecek S, Vahdat V, **Guyer RA**, Rizi RR, Optimization of scan parameters in pulmonary partial pressure oxygen measurement by hyperpolarized 3He MRI. (2008) Magn Reson Med, 59(1):124-31.
- 2. Emami K, Cadman RV, Fischer MC, Zhu J, Woodburn JM, Kadlecek SJ, **Guyer RA**, Law M, Vahdat V, Friscia ME, Ishii M, Yu J, Shrager JB, Rizi RR, Early changes of lung function and structure in an elastase model of emphysema--a hyperpolarized 3He MRI study. (2008). J of Appl Physiol, 104(3):773-86.
- 3. **Guyer RA**, Hellman M, Emami K, Kadlecek SJ, Cadman RV, Law M, Ishii M, Vadhat V, Yu J, Rizi RR, A robust method for measuring regional pulmonary parameters in the presence of noise. (2007) Acad Radiol, 15(6):740-52.
- 4. LaPar DJ, Stukenborg GJ, **Guyer RA**, Stone ML, Bhamidipati CM, Lau CL, Kron IL, Ailawadi GA. Primary Payer Status is Associated with Mortality and Resource

- Utilization for Coronary Artery Bypass Grafting Operations. (2012). Circulation, 126(11 Suppl 1):S132-9.
- 5. **Guyer RA**, Macara IG, Loss of the Polarity Protein PAR3 Activates STAT3 Signaling via an Atypical Protein Kinase C (aPKC)/NF-kB/Interleukin-6 (IL-6) Axis in Mouse Mammary Cells. (2015) The Journal of Biological Chemistry, 290(13):8457-68.
- 6. Dennis BM, Gondek SP, **Guyer RA**, Hamblin SE, Gunter OL, Guillamondegui OD, Use of an evidence-based algorithm for patients with traumatic hemothorax reduces need for additional interventions. (2017) J Trauma Acute Care Surg, 82(4):728-732.
- 7. Wandell G, Miller C, Rathor A, Thee Wei T, **Guyer RA**, Turner J, Hwang P, Davis G, Humphreys I, A Multi-Institutional Review of Outcomes in Biopsy-Proven Acute Invasive Fungal Sinusitis. (2018) Int Forum of Allergy Rhinol, 8(12):1459-1468.
- 8. Nagy N, **Guyer RA**, Hotta R, Zhang D, Newgreen DF, Halasy V, Kovacs T, Goldstein AM, RET overactivation leads to concurrent Hirschsprung disease and intestinal ganglioneuromas. (2020) Development, 147(21):dev190900.
- 9. Bhave S, Arciero E, Baker C, Ho WL, **Guyer RA**, Goldstein AM, Pan-enteric neuropathy and dysmotility are present in a mouse model of short-segment Hirschsprung disease and may contribute to post-pullthrough morbidity. (2021) J Pediatr Surg, 56(2):250-256.
- 10. Stavely R, Bhave S, Ho WLN, Ahmed M, Pan W, Rahman AA, Ulloa J, Bousquet N, Omer M, **Guyer RA**, Nagy N, Goldstein AM, Hotta R, Enteric mesenchymal cells support the growth of postnatal enteric neural stem cells. (2021) Stem Cells, 39(9):1236-1252.
- 11. Westfal ML, Okiemy O, Chung PHY, Feng J, Lu C, Miyano G, Tam PKH, Tang W, Wong KKY, Yamataka A, **Guyer RA**, Doody DP, Goldstein AM, Optimal timing for soave primary pull-through in short-segment Hirschsprung disease: A meta-analysis. (2021) J Pediatr Surg, 57(4):719-725.
- 12. Bhave S, Ho WLN, Cheng K, Omer M, Bousquet N, **Guyer RA**, Hotta R, Goldstein AM, Tamoxifen administration alters gastrointestinal motility in mice. (2022) Neurogastroenterol Motil 34(5):e14357.
- 13. Stavely RS, Hotta R, Picard N, Rahman AA, Pan W, Bhave S, Omer M, Ho WL, **Guyer RA**, Goldstein AM. Schwann cells in the subcutaneous adipose tissue have neurogenic potential and can be used for regenerative therapies. (2022) Sci Transl Med 14:eabl8753.

- 14. Cramm SL, Lipskar AM, Graham DA, Kunisaki SM, Griggs CL, Allukian M, Russell RT, Chandler NM, Santore MT, Aronowitz DI, Blakely ML, Campbell B, Collins DT, Commander SJ, Cowles RA, DeFazio JR, Echols JC, Esparaz JR, Feng C, Guyer RA, Hanna DN, He K, Kahan AM, Keane OA, Lamoshi A, Lopez CM, McLean SE, Pace E, Regan MD, Scholz S, Tracy ET, Williams SA, Zhang L, Rangel SJ, for the Eastern Pediatric Surgery Network. Association of Gangrenous, Suppurative, and Exudative Findings With Outcomes and Resource Utilization in Children With Nonperforated Appendicitis. (2022) JAMA Surgery doi:10.1001/jamasurg.2022.1928.
- 15. *Bhave SB, *Guyer RA, Picard N, Omer M, Hotta R, Goldstein AM. Ednrb/ mice with hirschsprung disease are missing Gad2-expressing enteric neurons in the ganglionated small intestine. (2022) Front Cell Dev Biol doi:10.3389/fcell.2022.917243

Non-peer reviewed scholarship in print or other media:

Reviews, chapters, and editorials

- 1. Macara IG, **Guyer RA**, Richardson G, Huo Y, Ahmed M, Epithelial Homeostasis. (2014) Current Biology, 24:R815-25.
- 2. **Guyer RA**, Mueller JL, Goldstein AM, Applications of Single-Cell Sequencing Technology to the Enteric Nervous System. (2022) Biomolecules 12(3):452.

Case reports

- 1. **Guyer RA**, Turner JH, Delayed Presentation of Traumatic Cerebrospinal Fluid Rhinorrhea: Case Report and Literature Review. (2015). Allergy and Rhinology, 6:188-190.
- 2. Mueller JL, **Guyer RA**, Adler JT, Mullen JT, Metastatic renal cell carcinoma to the small bowel: three cases of GI bleeding and a literature review. (2018) CEN Case Reports, 7:39-43.

Other non-peer reviewed scholarship

- 1. **Guyer RA**, Radiology Paging a Good Mathematician: Why Math Can Contribute More to Medicine Than You Might Think. (2008) Math Horizons.
- 2. **Guyer RA**, Schwarze ML, Gosain A, Maggard-Gibbons M, Keswani SG, Goldstein AM, Top ten strategies to enhance grant-writing success. (2021) Surgery S0039-6060(21)00656-5.

- 3. **Guyer RA**, Stavely RS, Robetson K, Bhave S, Hotta R, Kaltschmidt JA, Goldstein AM, Single-Cell Multiome Sequencing Clarifies Enteric Glial Cell Diversity and Identifies an Intraganglionic Population Poised for Neurogenesis. (2022) BioRxiv 10.1101/2021.08.24.457368.
- 4. *Guyer RA, *Picard N, Mueller JL, Murphy AJ, Cornejo KM, Hotta R, Goldstein AM. Differentiated neuroblastoma cells remain epigenetically poised for de-differentiation to an immature state. (2022) BioRxiv 10.1101/2022.07.10.499470.

Thesis:

1. Tumor Suppressor Mechanisms of the Polarity Protein Par-3, Doctoral thesis submitted to the faculty of Vanderbilt University

Abstracts, Poster Presentations, and Exhibits Presented at Professional Meetings:

- 1. Bousquet N, **Guyer RA**, Hotta R, Goldstein AM, CD49b Expression In Neuroblastoma Distinguishes Chemosensitive, Proliferative Cells From Chemoresistant, Quiescent Cells. Virtual podium presentation at American College of Surgeons Clinical Congress, virtual conference, 10/2021.
- 2. **Guyer RA**, Bhave S, Stavely S, Hotta R, Goldstein AM, Single cell RNA sequencing reveals enteric glial subpopulations with therapeutic potential in Hirschsprung disease. Virtual podium presentation at American College of Surgeons Clinical Congress, virtual conference, 10/2020.
- 3. **Guyer RA**, Bhave S, Stavely R, Hotta R, Li Z, Ngan E, Goldstein AM, Single-cell Analysis Shows Postnatal Enteric Glial Cells Closely Resemble Embryonic Neural Stem Cells. Virtual oral quickshot presentation at Academic Surgical Congress, virtual conference, 2/2021.
- 4. **Guyer RA**, Mueller J, Bhave S, Stavely R, Hotta R, Goldstein AM, Reduced Expansion In Culture of Enteric Glial Cells From Hirschsprung Disease Mice. Oral podium presentation at Academic Surgical Congress, Orlando, FL. 2/2022.
- 5. **Guyer RA**, Bhave S, Stavely RS, Hotta R, Goldstein AM, Joint Single-Nucleus ATAC and RNA Sequencing Reveal Chromatin Closing at Distal Regulatory Elements During Glial-to-Neuron Differentiation. Oral podium presentation at the 6th International Symposium on Development of the Enteric Nervous System, Ferrara, Italy, 4/2022.
- 6. **Guyer RA**, Picard N, Mueller JS, Hotta R, Goldstein AM, Neuroblastoma Cells Include Heterogeneous Populations of Highly Plastic Quiescent and Proliferative Cells Marked

by CD49b Expression. Oral podium presentation at the American Pediatric Surgical Association Annual Meeting, San Diego, CA, 5/2022.

* Co-first author

Narrative Report

I am a general surgery resident with formal training as a cell and developmental biologist, and with aspirations to become an academic pediatric surgeon. I was a dual-degree MD/PhD student at the University of Virginia and Vanderbilt University, and I earned a PhD in Ian Macara's laboratory. Following medical school, I came to Harvard Medical School and Massachusetts General Hospital for clinical training in surgery and a postdoctoral fellowship in the MGH Pediatric Surgical Research Laboratories. I am scheduled to graduate from residency in June 2024, after which I hope to undertake a clinical fellowship in pediatric surgery.

My research expertise is focused on understanding how cells achieve and maintain a differentiated phenotype, with particular emphasis on the epigenetic mechanisms that control gene expression. I seek to understand how chemical changes to DNA and DNA-associated proteins direct neural crest to commit to differentiated fates, and how perturbations to this process contribute to neoplasia. My work has been supported by numerous grant awards, both during graduate school and during my postdoctoral training. These grants include appointment to the University of Virginia Cancer Research T32 training grant, the Marshall K. Barlett Research Fellowship from the MGH Department of Surgery, an NIH F32 postdoctoral fellowship, and, most recently, the Patricia K. Donahoe Resident Research Catalyst Award from the MGH Department of Surgery. I have published 15 peer-reviewed research papers, and I have several more currently in preparation. I have also presented at numerous national and international meetings, including the American Society for Cell Biology, International Symposium on the Development of the Enteric Nervous System, American College of Surgeons, Academic Surgical Congress, and American Pediatric Surgical Association. I am currently working to understand how chromatin structure directs specification of enteric neurons, and how disruption of epigenetic mechanisms lends inappropriate plasticity to neuroblastoma cells. These projects leverage traditional methods in cell and developmental biology as well as utilizing cutting-edge tools for single-cell transcriptomic and epigenetic analysis.

In my role as a clinical general surgery resident, I care for patients with surgical disease at Massachusetts General Hospital, Newton-Wellesley Hospital, and North Shore Medical Center. Since coming to Harvard Medical School as a trainee, I have endeavored to provide outstanding surgical care to patients, contribute to the education of medical students and junior residents through teaching, and to advance medical science through basic discovery. Through teaching, lectures, written work, basic biological research, and involvement with professional societies, I am seeking to improve the care of babies and children suffering from congenital malformations and cancer.