### **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: Sharma, Gaurav

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

POSITION TITLE: Assistant Professor of Cardiovascular and Thoracic Surgery, Biomedical Engineering and Advanced Imaging Research

EDUCATION/TRAINING (Begin with baccalaureate or other initial professional education, such as nursing,

include postdoctoral training and residency training if applicable.)

INSTITUTION AND LOCATION	DEGREE	START	COMPLETION	FIELD OF STUDY
	(if	DATE	DATE	
	applicable)	MM/YYYY	MM/YYYY	
School of Life Sciences, Dr. B. R.	MS	09/2007	11/2009	Biotechnology
Ambedkar University, Agra, Uttar				
Pradesh				
All India Institute of Medical Sciences,	PhD	07/2010	04/2016	Biomedical Sciences
New Delhi				
Quantic School of Business &	MBA	07/2019	07/2020	Management,
Technology, Washington, DC				Leadership, etc.
UT Southwestern Medical Center,	Postdoctoral	06/2016	06/2022	Intermediary Metabolism
Dallas, TX	Fellow			and Metabolic Imaging

### A. Personal Statement

At the Sharma Lab at UT Southwestern, our research focuses on advancing metabolic imaging, stable isotope tracing, and multi-omics approaches to better understand cardiovascular disease and improve outcomes in heart failure and transplantation. A central area of our work is the development and application of hyperpolarized carbon-13 MRI (HP <sup>13</sup>C-MRI), which enables real-time, noninvasive assessment of myocardial metabolism. By integrating metabolic imaging with clinical insight and systems biology, we aim to develop tools that support surgical decision-making, identify viable myocardium, and personalize therapy for ischemic cardiomyopathy. The Sharma Lab leads multiple translational studies, including clinical trials that combine HP 13C-MRI with FDG-PET imaging. In parallel, we are working to improve donor heart and lung preservation through optimized perfusion strategies and metabolic profiling. Our research is supported by the American Heart Association, UK Research and Innovation, the Australian National Health and Medical Research Council, and the Canadian Institutes of Health Research.

Most recently, we have expanded our efforts into artificial intelligence for medicine, using machine learning to analyze complex imaging and metabolic datasets. These Al-driven approaches are designed to improve outcomes in both transplant medicine and cardiometabolic disease.

The Sharma Lab is dedicated to training the next generation of scientists in translational imaging, computational biology, and AI-enabled medicine. We also contribute to peer review and editorial leadership across journals and funding agencies worldwide. Our mission is to translate innovations in metabolism, imaging, and AI into practical tools for improving cardiovascular care and transplantation outcomes.

Selected publications closely related to the proposed studies:

- 1. **Sharma G**, McNeil S, Lin SH, Harrison C, Park JM, Peltz M, Malloy CR, Jessen ME. Hyperpolarized Carbon-13 Metabolic Imaging Detects Changes in Mitochondrial Metabolism in Patients Before and After Coronary Artery Bypass Graft Surgery. *Circulation.* 150 (2024) doi: 10.1161/circ.150.suppl 1.4115015
- 2. **Sharma G**, McNeil S, Lin SH, Harrison C, Park JM, Peltz M, Malloy CR, Jessen ME. Hyperpolarized C Metabolic Imaging in Human Subjects with Advanced Coronary 13 Artery Disease: Initial Experience and Thoughts. *Proc. Intl. Soc. Mag. Res. Med.*, 4757 (2024)
- 3. **Sharma G.**, Park JM., Peltz M., Malloy CR, Jessen M E. *Hyperpolarized 13C Pyruvate MRI and PET in Single Exam Using PET-MR for Ischemic Heart Disease*. *Patent Cooperation Treaty (PCT) Application* No. PCT/US2024/061542. Filing Date: December 20, 2024.

4. **Sharma G**, Wen W, Meptue NR Hever T, Malloy CR, Sherry AD, Khemtong C. Co-Polarized [1-13C]Pyruvate and [1,3-13C2]Acetoacetate Provide a Simultaneous View of Cytosolic and Mitochondrial Redox in a Single Experiment. **ACS Sensors. 2021**. PMID: 34761912 \*Featured on Journal Cover and Favorite Papers from the First 10 Years of ACS Sensors (DOI: 10.1021/acssensors.4c03746)

### **B.** Positions and Honors

### **Positions and Employment**

- 2025- Present Member, Harold C. Simmons Comprehensive Cancer Center, UT Southwestern Medical Center, Dallas, Texas, United States
- 2024 present Assistant Professor, Department of Biomedical Engineering (Secondary), UT Southwestern Medical Center, Dallas, TX, USA
- 2022 present Assistant Professor, Department of Cardiovascular and Thoracic Surgery (**Primary**), UT Southwestern Medical Center, Dallas, TX, USA
- 2022 present Assistant Professor, Advanced Imaging Research Center (Secondary), UT Southwestern Medical Center, Dallas, TX, USA
- 2016 2022 Postdoctoral Researcher, Advanced Imaging Research Center, UT Southwestern Medical Center, Dallas, TX, USA
- 2015 2016 Research Associate, Jawaharlal Nehru University, New Delhi, India
- 2012 2015 Senior Research Fellow, All India Institute of Medical Sciences, New Delhi, India
- 2010 2012 Junior Research Fellow, All India Institute of Medical Sciences, New Delhi, India

### Other Experience and Professional Memberships

- 2024-present Grant Reviewer, American Lung Association
- 2024-present Ad hoc Grant Reviewer, National Science Foundation (NSF)
- 2024-present ECR Grant Reviewer, NIH-Surgery, Anesthesiology and Trauma Study Section [SAT]
- 2024-present Ad hoc Grant Reviewer, UK Research and Innovation (UKRI)
- 2024-present Member, American College of Cardiology (ACC)
- 2024-present Member, Radiological Society of North America
- 2024-present Editorial Board Member, NPJ-Imaging (Springer Nature)
- 2023-present Member, College of Reviewers, Canadian Institutes of Health Research (CIHR)
- 2023-present Grant Reviewer, National Health and Medical Research Council, Australia
- 2023-present Member, American Physiological Society (APS)
- 2023-present Associate Editor, The Journal of Heart and Lung Transplantation-Open
- 2023-present Member, Institutional Review Board (IRB), UT Southwestern Medical Center
- 2023-present Ad hoc early career grant reviewer, National Institute of Health
- 2023-present Grant Reviewer, Innovative Projects Award, American Heart Association
- 2022-present Grant Reviewer, Career Development Award, American Heart Association
- 2022-present Grant Reviewer, Cardiac Biology & Regulation, American Heart Association
- 2022-present Member, The International Society for Heart and Lung Transplantation (ISHLT)
- 2020-present Associate Editor, Medicine (LWW journals, Wolters Kluwer Health, Inc.)
- 2020-present Associate Editor, BMC Research Notes (Springer Nature)
- 2020-present Associate Editor, BMC Cardiovascular Disorders (Springer Nature)
- 2020-present Member of Reviewer Board, International Journal of Molecular Sciences (MDPI), Journal of
  - Imaging (MDPI), Metabolites (MDPI) and Biomolecules (MDPI)
- 2020-present Review Editor, Frontiers in Cardiovascular Medicine (Frontiers Media S.A.)
- 2020-present Academic Editor, PeerJ (O'Reilly and SAGE)
- 2020-present Associate Editor, BMC Cancer (Springer Nature)
- 2020-present Associate Editor, Journal of Translational Medicine (Springer Nature)
- 2020-present Topic Editor, Bioengineering (MDPI)
- 2020-present Member of Editorial Advisory Board, Heliyon (Cell Press)
- 2019-present Ad Hoc Reviewer, manuscripts submitted in peer-reviewed indexed journals (>250)
- 2019-present Review Editor, Frontiers in Molecular Biosciences (Frontiers Media S.A.)
- 2016-present Life Member, ISMRM Indian Chapter
- 2016-2017 Member, American Association for the Advancement of Science (AAAS)

2013-present Member, World Molecular Imaging Society (WMIS)
 2017-present Member, American Heart Association (AHA)
 2013-present Member, International Society for Magnetic Resonance in Medicine (ISMRM)

# **Honors**

2025	Fellow, CV Section, American Physiology Society, USA
2024	CVSA Early Career Investigator Abstract Award, American Heart Association, Dallas, USA
2024	Fellow, American College of Cardiology, Washington, D.C, USA
2024	Honored Listee, Marquis Who's Who in America, NY, USA
2023	Second Century Early Faculty Independence Award, American Heart Association, Dallas, TX
2021	EBM of the month, BMC Cardiovascular Disorders (Springer Nature)
2019	1 <sup>st</sup> place in the Scientific Poster Award, National Center for In Vivo Metabolism, Dallas, USA
2018 - 2020	AHA Postdoctoral Award, American Heart Association, Dallas, TX, USA
2017, 2019	Educational Stipend, International Society of Magnetic Resonance in Medicine
	Travel award' to attend the Annual scientific meeting of WMIC, Seoul, Republic of Korea, Indian
2014	Council of Medical Research (ICMR), New Delhi, India
	'Travel award' to attend the Annual scientific meeting of ISMRM, USA, Indian Council of Medical
2013	Research (ICMR), New Delhi, India
	ICMR Junior/Senior Research Fellowship (highly coveted, ~1% success rate), Indian Council of
2010 - 2016	Medical Research (ICMR), New Delhi, India
	Graduate Aptitude Test for Engineering (GATE) in Life Sciences (1) and Biotechnology (2),
2009-2011	Jointly by the Indian Institute of Science (IISc) and the Indian Institute of Technology (IITs)

## C. Contribution to Science

- 1. Novel hyperpolarized <sup>13</sup>C methods for the non-invasive assessment of metabolism: I have been engaged in developing and deploying new technologies to access high-impact metabolic diseases. Hyperpolarized <sup>13</sup>C MR spectroscopy (MRS) using dynamic nuclear polarization (DNP) is an emerging technique to noninvasively assess mitochondrial metabolism in real-time. I have studied altered in-vivo and ex-vivo real-time metabolism in disease models using hyperpolarized <sup>13</sup>C MRS. Studies are underway to explore the utility of using hyperpolarized [<sup>13</sup>C] metabolites to analyze metabolism in-vivo and in perfused isolated organs from rodent models of diseases. My current research is focused on translating these agents and technologies into in-vivo imaging applications. Key advancements from ongoing work are reported in:
  - a. **Sharma G**, Maptue N, Rahim M, Trigo Mijes ML, Wen W, Hever T, Funk AM, Malloy CR, Young JD, Khemtong C. Oxidation of hyperpolarized [1-<sup>13</sup>C]pyruvate in isolated rat kidneys. *NMR Biomed.* 2023. PubMed PMID: 36285844\*Featured on Journal Cover
  - b. **Sharma G**, Wen W, Meptue NR Hever T, Malloy CR, Sherry AD, Khemtong C. Co-Polarized [1- 

    13C]Pyruvate and [1,3-13C2]Acetoacetate Provide a Simultaneous View of Cytosolic and Mitochondrial Redox in a Single Experiment. *ACS Sensors. 2021*. PMID: 34761912 \*Featured on Journal Cover
  - c. **Sharma G**, Wu CY, Wynn RM, Gui W, Malloy CR, Sherry AD, Chuang DT, Khemtong C. Real-time hyperpolarized <sup>13</sup>C magnetic resonance detects increased pyruvate oxidation in pyruvate dehydrogenase kinase 2/4-double knockout mouse livers. *Sci Rep. 2019* Nov 11;9(1):16480. PubMed PMID: 31712597; PubMed Central PMCID: PMC6848094.
  - d. Chen W, Sharma G, Jiang W, Maptue NR, Malloy CR, Sherry AD, Khemtong C. Metabolism of hyperpolarized <sup>13</sup> C-acetoacetate to β-hydroxybutyrate detects real-time mitochondrial redox state and dysfunction in heart tissue. *NMR Biomed.* 2019 Jun;32(6):e4091. PubMed PMID: 30968985; PubMed Central PMCID: PMC6525062.
- 2. Understanding the intermediary metabolism in cardiovascular pathophysiology: The vast majority of cardiovascular diseases are associated with disruption of metabolic pathways. Although research in cell biology has produced key information about how metabolic pathways and dependencies are regulated but provides limited information about altered oxidative metabolism in diseases. Tracing intermediary metabolism with isotope infusions/perfusion is safe and highly informative in rodents and humans. I have been currently involved in the understanding role of altered intermediary metabolism in rodent disease models using <sup>13</sup>C NMR spectroscopy, and mathematical modeling. Ongoing work demonstrates that <sup>13</sup>C

isotopomer analysis can be used to study the substrate competition and metabolic state in gene knockouts. Key findings from ongoing work are:

- a. Li Q, Li C, Elnwasany A, **Sharma G**, An YA, Zhang G, Elhelaly WM, Lin J, Gong Y, Chen G, Wang M, Zhao S, Dai C, Smart CD, Liu J, Luo X, Deng Y, Tan L, Lv SJ, Davidson SM, Locasale JW, Lorenzi PL, Malloy CR, Gillette TG, Vander Heiden MG, Scherer PE, Szweda LI, Fu G, Wang ZV. PKM1 Exerts Critical Roles in Cardiac Remodeling Under Pressure Overload in the Heart. *Circulation. 2021.* Aug 31;144(9):712-727. PMID: 34102853; PMCID: PMC8405569.
- b. Dai C, Li Q, May HI, Li C, Zhang G, **Sharma G**, Sherry AD, Malloy CR, Khemtong C, Zhang Y, Deng Y, Gillette TG, Xu J, Scadden DT, Wang ZV. Lactate Dehydrogenase A Governs Cardiac Hypertrophic Growth in Response to Hemodynamic Stress. *Cell Rep.* 2020 Sep 1;32(9):108087. PubMed PMID: 32877669.
- c. Cardoso AC, Lam NT, Savla JJ, Nakada Y, Pereira AHM, Elnwasany A, Menendez-Montes I, Ensley EL, Petric UB, Sharma G, Sherry AD, Malloy CR, Khemtong C, Kinter MT, Tan WLW, Anene-Nzelu CG, Foo RS, Nguyen NUN, Li S, Ahmed MS, Elhelaly WM, Abdisalaam S, Asaithamby A, Xing C, Kanchwala M, Vale G, Eckert KM, Mitsche MA, McDonald JG, Hill JA, Huang L, Shaul PW, Szweda LI, Sadek HA. Mitochondrial Substrate Utilization Regulates Cardiomyocyte Cell Cycle Progression. *Nat Metab.* 2020 Feb;2(2):167-178. PubMed PMID: 32617517; PubMed Central PMCID: PMC7331943.
- d. Makarewich CA, Baskin KK, Munir AZ, Bezprozvannaya S, **Sharma G**, Khemtong C, Shah AM, McAnally JR, Malloy CR, Szweda LI, Bassel-Duby R, Olson EN. MOXI Is a Mitochondrial Micropeptide That Enhances Fatty Acid β-Oxidation. *Cell Rep. 2018* Jun 26;23(13):3701-3709. PubMed PMID: 29949755; PMCID: PMC6066340.
- 3. Assessment of therapeutic interventions on metabolism in obesity or diabetes: The impaired regulation of carbohydrate and lipid metabolism is intricately linked with obesity. I investigated in vivo intermediary metabolism of glycerol in the liver, substrate competition, and therapeutic efficacy of drugs targeting mitochondrial pyruvate dehydrogenase kinases 1–4 (PDKs1–4) for obesity and type 2 diabetes using <sup>13</sup>C NMR isotopomer analysis and hyperpolarized <sup>13</sup>C-labeled substrates in diet-induced obese rodent models.
  - a. Jin ES, Malloy CR, **Sharma G**, Finn E, Fuller KNZ, Reyes YG, Lovell MA, Derderian SC, Schoen JA, Inge TH, Cree MG. Glycerol as a precursor for hepatic de novo glutathione synthesis in human liver. *Redox Biology.* 2023:102749. PubMed PMID: 37224695.
  - b. Wu CY, Satapati S, Gui W, Wynn RM, Sharma G, Lou M, Qi X, Burgess SC, Malloy C, Khemtong C, Sherry AD, Chuang DT, Merritt ME. A novel inhibitor of pyruvate dehydrogenase kinase stimulates myocardial carbohydrate oxidation in diet-induced obesity. *J Biol Chem.* 2018 Jun 22;293(25):9604-9613. PubMed PMID: 29739849; PubMed Central PMCID: PMC6016455.
  - c. Wu CY, Tso SC, Chuang JL, Gui WJ, Lou M, **Sharma G**, Khemtong C, Qi X, Wynn RM, Chuang DT. Targeting hepatic pyruvate dehydrogenase kinases restores insulin signaling and mitigates ChREBP-mediated lipogenesis in diet-induced obese mice. *Mol Metab. 2018* Jun;12:12-24. PubMed PMID: 29656110; PMCID: PMC6001905.
  - d. Thapa B, Suh EH, Parrott D, Khalighinejad P, Sharma G, Chirayil S, Sherry AD. Imaging β-Cell Function Using a Zinc-Responsive MRI Contrast Agent May Identify First Responder Islets. *Front Endocrinol* (*Lausanne*). 2022 Jan 31;12:809867. doi: 10.3389/fendo.2021.809867. PMID: 35173681; PMCID: PMC8842654.
- 4. Improving Donor Heart Preservation through Advanced Metabolic and Molecular strategies: My research has focused on enhancing the preservation of donor human hearts intended for transplantation by utilizing novel preservation systems and analyzing their metabolic profiles. I have investigated the effects of advanced hypothermic perfusion systems on donor heart metabolism and transcriptomics, contributing to better preservation strategies to translate for transplantation outcomes. Key findings include:
  - a. **Sharma G,** Vela R, Powell L, Malloy CR, Jessen M, Peltz M. Transcriptomic Signatures of Human Donor Hearts Preserved Using a Hypothermic Perfusion System. *The Journal of Heart and Lung Transplantation*. 2024;43(4):S248-S9. doi: 10.1016/j.healun.2024.02.1146.

- b. **Sharma G**, Vela R, Powell L, Malloy CR, Jessen ME, Peltz M. Metabolic Analysis of Human Hearts Preserved with a Novel Hypothermic Perfusion System. *The Journal of Heart and Lung Transplantation*. 2023;42(4):S185. doi: 10.1016/j.healun.2023.02.1696.
- c. **Sharma G**, Vela R, Powell L, Malloy CR, Jessen M, Peltz M. Transcriptomic Response of Human Donor Hearts to Temperature-Controlled Storage. *The Journal of Heart and Lung Transplantation*. 2024;43(4):S159. doi: 10.1016/j.healun.2024.02.321.
- d. **Sharma G**, Vela R, Powell L, Mizerska M, Deja S, Burgess S, Malloy CR, Jessen ME, Peltz M. Metabolic Indicators in Donor Hearts Following Conventional and Temperature Controlled Storage. **The Journal of Heart and Lung Transplantation.** 2023;42(4, Supplement):S378. doi: https://doi.org/10.1016/j.healun.2023.02.883.

# **Complete List of Published Work in My Bibliography:**

**ORCID ID:** 0000-0002-1754-7163

Bibliography: My Bibliography - NCBI (nih.gov)