RESUME

CONTACT Information Ashok Kumar Sharma

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EDUCATION

PhD in Computational Biology

Jan 2013 - May 2018

IISER Bhopal, Madhya Pradesh, India

- Advisor: Dr. Vineet Kumar Sharma
- Thesis: Development of Computational Models and Algorithms for Designing of Novel Microbiome-based Therapeutics.

Masters in Pharmacoinformatics

July 2010 - June 2012

NIPER Mohali, Punjab , India

- Advisor: Prof. Prasad V. Bharatam
- \bullet Thesis: Modelling and Designing of Glycogen Synthase Kinase 3 Inhibitors.

Bachelor of Pharmacy

Aug 2006 - June 2010

Dr. H. S. Gour University, Sagar, Madhya Pradesh, India

RESEARCH EXPERIENCE

Scientist II, Computational Biology

April 2023 - Present

Drug Safety Research Evaluation (DSRE)

Takeda

As a lead computational biologist at DSRE, I drive safety risk assessment through predictive modeling and large-scale transcriptomic data analysis, while spearheading pipeline development to enhance efficiency and consistency in drug discovery research.

- Developing AI/ML models for prediction of toxicological endpoints, specifically Hepatotoxicity (DILI) and Neurotoxicity.
- Supporting the Oligo working group by developing **off-target identification strategies** (e.g., developing a splicing analysis pipeline for the SSO program)
- Developed transcriptomic data analysis pipeline, ensuring its adaptability and efficiency for use across various projects, and utilized it to update **CellMap visualization dashboard**
- Contributing towards the development of **transcriptomic biomarkers** predictive of tumorigenic pathways as part of an ongoing industry consortium, with Takeda being one of the participating members.

Postdoctoral Scientist

Feb 2021 - Dec 2022

Casero Lab, Cedars-Sinai Medical Center

Advisor: <u>Dr. David Casero</u>, Director, Translational Multiomics, Inflammatory Bowel Immunobiology Research Institute

I was deeply involved in the analysis and integration of multi-omics datasets aimed at identifying pathogenic factors in Inflammatory Bowel Disease (IBD). Here is a summary of the specific projects I worked on during that time:

- Investigated **cell-specific gene expression** alterations in the host epithelium of Ulcerative Colitis (UC) patients post-IPAA surgery, and conducted **multi-omics datasets** to uncover microbial regulation of immune responses within creeping fat in Crohn's disease.
- Quantified **bacterial growth rates from metagenomic datasets** to identify activities associated with Crohn's disease progression.

Postdoctoral Associate

March 2018 - Feb 2021

Microbiomics Lab, University of Minnesota

Mentor: Dr. Andres Gomez

As the first computational team member in the department, I was responsible for managing and establishing computational pipelines on a high-performance computing facility at UMN. Furthermore, I played a crucial role in advancing various mutli-omics projects, detailed as follows:

- Analyzed whole genome sequencing data to comprehend the impact of diet and lifestyle on gut microbial taxonomic and functional compositions across traditional human and nonhuman primate populations.
- Identified gastrointestinal gene expression patterns in response to functional alterations in the gut microbiome, leveraging **meta-transcriptomic data analysis** techniques.
- Involved in developing innovative and advanced computational models for **meta-analysis** of diverse microbiome datasets, aimed at exploring microbiome-host interactions.

Senior Research Fellow (SRF) MetaBioSys Lab, IISER Bhopal Mentor: Dr. Vineet Kumar Sharma June 2015 - March 2018

- Developed machine learning-based computational methods for the analysis of complex genomic and metagenomic datasets. Also, conducted high-throughput predictions of gut microbial-mediated drug metabolism and toxicity using chemical structure-centric approaches.
- Identified taxonomic, functional, and metabolic markers associated with colorectal cancer patients in India.

Publications

- 1. Sarode, G.V., Mazi, T.A., Neier, K., Shibata, N.M., Jospin, G., Harder, N.H., Caceres, A., Heffern, M.C., **Sharma, A.K.**, More, S.K. and Dave, M., **2023**. The role of intestine in metabolic dysregulation in murine Wilson disease. **Hepatology Comm**, 7(10).
- Guse, K., Sharma, A.K., Weyenberg, E., Davison, S., Ma, Y., Choi, Y., Johnson, A.J., Chen, C. and Gomez, A., 2023. Regular consumption of lacto-fermented vegetables has greater effects on the gut metabolome compared with the microbiome. Gut Microbiome , 4, p.e11.
- 3. Dina G. Moussa, **Sharma, A.K.**, Tamer Mansour, Bruce Witthuhn; Jorge Perdigao; Joel D. Rudney; Conrado Aparicio; Andres Gomez, **2022**. Functional Biomarkers of Ex-vivo Dental Caries Onset. **Journal of oral microbiology**, 14(1), p.2123624.
- Rosa, F., <u>Sharma, A.K.</u>, Gurung, M., Casero, D., Matazel, K., Bode, L., Simecka, C., Elolimy, A.A., Tripp, P., Randolph, C. and Hand, T.W., <u>2022</u>. Human Milk Oligosaccharides Impact Cellular and Inflammatory Gene Expression and Immune Response. <u>Frontiers in Immunology</u>. 13: 907529. doi: 10.3389/fimmu.
- Saxena, R., Prasoodanan PK, V., Gupta, S., Gupta, S., Waiker, P., Samaiya, A., <u>Sharma, A.K.</u> and Sharma, V.K., 2022. Assessing the effect of smokeless tobacco- consumption on oral microbiome in healthy and oral cancer patients. Frontiers in <u>Cellular and Infection Microbiology</u>, p.331.
- 6. Omontese, B.O., Sharma, A.K., Davison, S., Jacobson, E., DiConstanzo, A., Webb, M.J. and Gomez, A., 2022. Microbiome network traits in the rumen predict average daily gain in beef cattle under different backgrounding systems. Animal Microbiome, 4(1), pp.1-15.
- 7. Sharma, A.K., & Sam Davison; Barbora Pafco; Jonathan B. Clayton, Jessica M. Rothman, Matthew R. McLennan, Marie Cibot, Terence Fuh, Roman Vodicka, Carolyn

- Jost Robinson, Klara Petrzelkova, and Andres Gomez, **2022**. The primate gut mycobiome bacteriome interface is impacted by environmental and subsistence factors. **npj Biofilms Microbiomes** 8(1), pp.1-11..
- 8. Vishnu Prasoodanan P K, & <u>Ashok K Sharma</u>, Shruti Mahajan, Darshan B Dhakan, Abhijit Maji, Joy Scaria, Vineet K Sharma, **2021**. Western and non-western gut microbiomes reveal new roles of Prevotella in carbohydrate metabolism and mouth-gut axis. **npj Biofilms Microbiomes**, Oct 7;7(1):77.
- 9. Sharma, A.K., & Petrzelkova, K., Pafco, B., Robinson, C.A.J., Fuh, T., Wilson, B.A., Stumpf, R.M., Torralba, M.G., Blekhman, R., White, B. and Nelson, K.E., Leigh S.R., Gomez A, 2020. Traditional human and nonhuman primate populations show parallel gut microbiome adaptations to analogous dietary conditions. mSystems, 5(6).
- 10. Gomez, A., Sharma, A.K., Grev, A., Sheaffer, C. and Martinson, K., 2020. The horse gut microbiome responds in a highly individualized manner to forage lignification. Journal of Equine Veterinary Science, 96, p.103306.
- 11. Sharma, A.K., & Debusk, W.T., Stepanov, I., Gomez, A. and Khariwala, S.S., 2020. Oral microbiome profiling in smokers with and without head and neck cancer reveals variations between health and disease. Cancer Prevention Research, 13(5), pp.463-474.
- 12. Gupta, A., Dhakan, D.B., Maji, A., Saxena, R., PK, V.P., Mahajan, S., Pulikkan, J., Kurian, J., Gomez, A.M., Scaria, J. and Amato, K.R., **Sharma, A.K.** and Sharma V.K, **2019**, Association of Flavonifractor plautii, a flavonoid degrading bacterium, with the gut microbiome of colorectal cancer patients in India. **mSystems**, 4(6).
- 13. Gomez, A.*, Sharma, A.K.*, Mallott, E.K., Petrzelkova, K.J., Robinson, C.A.J., Yeoman, C.J., Carbonero, F., Pafco, B., Rothman, J.M., Ulanov, A. and Vlckova, K, 2019. Plasticity in the human gut microbiome defies evolutionary constraints. mSphere, 4(4), pp.e00271-19.
- 14. **Sharma, A.K.**, Pafčo, B., Vlčková, K., Červená, B., Kreisinger, J., Davison, S., Beeri, K., Fuh, T., Leigh, S.R., Burns, M.B. and Blekhman, R., Gomez, A, **2019**. Mapping gastrointestinal gene expression patterns in wild primates and humans via fecal RNA-seq. **BMC genomics**, 20(1), p.493.
- 15. Pafčo, B., Sharma, A.K., Petrželková, K.J., Vlčková, K., Todd, A., Yeoman, C.J., Wilson, B.A., Stumpf, R., White, B.A., Nelson, K.E. and Leigh, S., 2019. Gut microbiome composition of wild western lowland gorillas is associated with individual age and sex factors. American journal of physical anthropology, 169(3), pp.575-585.
- 16. Dhakan, D.B., Maji, A., <u>Sharma, A.K.</u>, A.K., Saxena, R., Pulikkan, J., Grace, T., Gomez, A., Scaria, J., Amato, K.R. and Sharma, V.K, **2019**. The unique composition of Indian gut microbiome, gene catalogue, and associated fecal metabolome deciphered using multi-omics approaches. **GigaScience**, 8(3), p.giz004.
- 17. Kumar, K., Dhoke, G.V., **Sharma, A.K.**, Jaiswal, S.K. and Sharma, V.K., **2019**. Mechanistic elucidation of amphetamine metabolism by tyramine oxidase from human gut microbiota using molecular dynamics simulations. **Journal of cellular biochemistry**, 120(7), pp.11206-11215.
- 18. Kumar, K., Jaiswal, S.K., Dhoke, G.V., Srivastava, G.N., Sharma, A.K. and Sharma, V.K., 2018. Mechanistic and structural insight into promiscuity based metabolism of cardiac drug digoxin by gut microbial enzyme. Journal of cellular biochemistry, 119(7), pp.5287-5296.

- 19. Sharma, A.K., Jaiswal, S.K., Chaudhary, N. and Sharma, V.K., 2017. A novel approach for the prediction of species-specific biotransformation of xenobiotic/drug molecules by the human gut microbiota. Scientific reports, 7(1), pp.1-13.
- 20. Sharma, A.K., Srivastava, G.N., Roy, A. and Sharma, V.K., 2017. ToxiM: A toxicity prediction tool for small molecules developed using machine learning and chemoinformatics approaches. Frontiers in pharmacology, 8, p.880.
- 21. Gupta, S., Sharma, A.K.*, Shastri, V., Madhu, M.K. and Sharma, V.K., 2017. Prediction of anti-inflammatory proteins/peptides: an insilico approach. Journal of translational medicine, 15(1), pp.1-11.
- 22. Gupta, A., Kumar, S., Prasoodanan, V.P., Harish, K., **Sharma**, **A.K.** and Sharma, V.K., **2016**. Reconstruction of bacterial and viral genomes from multiple metagenomes. **Frontiers in microbiology**, 7, p.469.
- 23. Gupta, S., Sharma, A.K.*, Jaiswal, S.K. and Sharma, V.K., 2016. Prediction of biofilm inhibiting peptides: an in silico approach. Frontiers in microbiology, 7, p.949.
- 24. Chaudhary, N., Sharma, A.K.*, Agarwal, P., Gupta, A. and Sharma, V.K., 2015. 16S classifier: a tool for fast and accurate taxonomic classification of 16S rRNA hypervariable regions in metagenomic datasets. PLOS One, 10(2), p.e0116106.

CONFERENCE PUBLICATIONS

- Sharma, A.K., Martin, A., Moskowitz, J.E., Bora, S., Legree, K., Dorrestein, P., Underhill, D., Knight, R., Chen, P. and Devkota, S., 2022. 1166: In-patient antibiotic exposure promotes sars-cov-2 persistence in the gi tract in covid-19 admitted patients. Gastroenterology, 162(7), pp.S-279].
- Lahcene, N.L., Moskowitz, J.E., Sharma, A.K., Martin, A., Merchant, A., Fleshner, P. and Devkota, S., 2022. 1059: Spatial characterization of immune cells and bacterial co-localization in crohn's disease's creeping fat. Gastroenterology, 162(7), pp.S-241.

BOOK CHAPTER

1. <u>Sharma, A.K.</u> and Dubey, V.S., **2021**. Metagenome Assembly for Gut Microbial Functional Diversity Associated with Xenobiotic Degradation. In Metagenomics and Microbial Ecology (pp. 79-87). **CRC Press**.

ORAL/POSTER PRESENTATIONS

- Selected speaker, 2022 'In-Patient Antibiotic Exposure Promotes SARS-CoV-2 Persistence in the GI Tract in COVID-19 Admitted Patients' at *Digestive Disease Week* (DDW) 2022, at San Diego, CA.
- **Delivered a lecture, 2020** 'Emergence of microbiome in therapeutics Ongoing efforts, challenges, and future opportunities' at *Department of Pharmaceutical Sciences*, Dr. Hari Singh Gaur University Sagar, India.
- Selected as one of the finalist's, 2019 in the Novartis Academia Hackathon event held on event to be held August 12th 23rd, 2019 on the Novartis Campus in Cambridge, MA, USA.
- Selected speaker, 2018 'Mapping gastrointestinal gene expression patterns from fecal RNA-seq' at EpiQ (Quantitative Epidemiology) seminar series, UMN Seminar.
- Selected speaker 2015 'Fast and Accurate Taxonomic Classification of 16S rRNA Hypervariable Regions in Metagenomic Datasets using 16S Classifier' in The Human Microbiome conference, at EMBL, Heidelberg, Germany.

EDITOR/REVIEWER FOR JOURNALS

• Associate Editor for Frontiers in Microbiology - Systems Microbiology; Guest Associate Editor for Frontiers in Microbiology - Microbiome in Health and Disease; Review Editor for Frontiers in Microbiology - Gastrointestinal Microbes

Achievements

- International Travel Grant, 2019 Received \$1000 grant to present my work at Keystone Symposium on "Microbiome: Therapeutic Implications (T1)" in October 2019 at Ireland.
- PBC Postdoctoral Fellowship, 2018 Received a Israel government fellowship for three years to pursue postdoctoral research at Bar-Ilan University, Israel.
- DST Travel Award, 2017 Received travel award from Department of Science and Technology, India to present my doctoral research in Symposium: NextGen Immunology at Rehovot, Israel.
- EMBL Grant, 2015 Received €1000 grant to present my work in The Human Microbiome Conference" at EMBL Germany.
- GATE, 2012 Secured 376 All India Rank in *Graduate Aptitude Test in Engineering* for Life Sciences conducted by IIT Delhi

TEACHING EXPERIENCE

- BIOL 1961, Foundations of Biology Lab I for Biological Sciences Majors (BIOL): Teaching basics of microbiome to computational microbiology students (16 contact hours/week, from July 2018).
- BIOL 3004, Foundations of Biology for Biological Sciences Majors, Part II Laboratory: Leading various groups of computational microbiology students for bioinformatics analysis of 16S rRNA data from published microbiome studies (16 contact hours/week, from July 2018).

SELECTED SKILLS

- Programming and Scripting: Proficient in R and Python; shell scripting in Unix/Linux environments
- Multi-omics data processing: Proficient in Genomics, Metagenomics, Metabolomics, and Transcriptomics (Bulk, and Single-cell), as well as Imaging Mass Cytometry (IMC). My foundational skills equip me to quickly adapt and effectively analyze any kind of sequencing dataset.
- Statistical analysis: Data mining, multivariate statistics, machine learning/AI, mixed effects modeling, and target/biomarker identification
- Tools and databases: Installation, maintenance, development of bioinformatics tools and pipelines within local and HPC environments, including workflow management with Nextflow
- Bioinformatics Libraries: Bioconductor, Seurat, mixOmics, DESeq2, caret, limma, ggplot2, MetaboAnalystR, Pandas, NumPy, SciPy, Matplotlib, TensorFlow, PyTorch, scikit-learn, Keras. Capable of adapting to incorporate additional libraries as required for comprehensive data analysis and interpretation.
- Visualization Dashboards: RShiny, Tableau
- Cloud Computing: high-performance computing (HPC); AWS services, including AWS S3 bucket