asahu@broadinstitute.org

RESEARCH SUMMARY

My two research programs interface cancer immunology with Artificial Intelligence and probabilistic approaches. **First**, my cancer-immunology program identifies drugs with dual anticancer potential through novel AI/statistical methods. It has identified a new class of drugs called bipotent drugs, for which I was awarded the prestigious Michelson Prize by Human Vaccines Project and was also selected among the "40 most promising young Cancer Immunology scientists" by Irving Foundation. **Second**, my single-cell research program, funded through K99 pathway to independence award, develops probabilistic approaches for single-cell and spatial genomics data. My lab will extend these programs to precision medicine and antibody design.

ACADEMIC POSITION

University of Colorado Anschutz Medical Campus

2022-

Cell: (1) 2403918125

Asst. Clinical Professor, Department of Obstetrics and Gynecology

Broad Institute 2018-2022

Human Vaccine Project Fellow, Data Science Adviser: Prof. X. Shirley Liu, Prof. Rafael Irizarry, Prof. Keith T Flaherty

ACADEMIC AWARDS

NCI K99 Pathway to Independence Career Award

• National Cancer Institute K99 Career Award, 2020.

Arthur L. Irving Family Foundation Cancer Immunology

• 40 promising young scientists in Cancer Immunology, 2020.

Michelson Medical Research Foundation and Human Vaccines Project

• Michelson Prize, 2019.

Quantitative Challenges in Cancer Immunology and Immunotherapy

• Stellar Research Award, 2019.

National Institute of Minority Health and Health Disparities fellowship

• Health Disparity Research Fellowship, 2016-2017.

Invention of the year, UMD, 2016

• Harnessing Synthetic Rescues to Assess and Counteract Resistance to Treatment in Cancer, University of Maryland, 2016 Invention of the year.

National Institute of Health GPP research fellowship

• Graduate Partnership Program Fellowship, 2012.

Max-Planck-Institut für Informatik fellowship

• Summer research German Govt. fellowship, 2012.

Dean fellowship, University of Maryland

• Dean Fellowship, 2011.

EDIC fellowship

• EDIC Swiss Doctoral Fellowship, 2011.

MSc Research Scholar

• MSc Research Scholar Fellowship, EPFL 2010.

Indian Statistical Institute Calcutta Entrance Exam

• All India Rank 2, Indian Statistical Institute Entrance exam, 2003.

EXTERNAL RESEARCH SUPPORT

K99/R00 Career Development Award

2020 - 2025

Identifying drug synergistic with cancer immunotherapy. NIH/NCI~K99CA248953 PI (\$971,726)

Michelson Med Res Foundation, Young Investigator Award

2019 - 2022

AI-predictor to identify immunomodulatory drugs. MP19-02-190-Sahu-A

PENDING RESEARCH SUPPORT

Novartis Drug Discovery and Translational Research Program Pending

Developing AI-based precision oncology by targeting inter-lesion and intra-tumor heterogeneity in highly resistant patients.

Selected for final round

PI: Sahu & Juric

EDUCATION

University of Maryland, College Park,

PhD., Computer Science (2016),

Adviser: Prof. Eytan Ruppin, Prof. Sridhar Hannenhalli

Thesis: Data driven approaches to identify determinants of heart disease and cancer

resistance.

École Polytechnic Fédérale de Lausanne, Switzerland (and University of Washington, Seattle)

M.S, Genome Science, Computer Science (2011)

Adviser: Prof. Bernard Moret, Prof. William Stafford Noble

Master Thesis: ENCODE: Unsupervised Segmentation of ENCODE data

National Institute of Technology, Trichy, India,

B.Tech.(2007), Electronics and Communication Engineering NIT

Bachelor Thesis: Hiding audio in video with optimized source and channel coding

Adviser: Dr. P. Palaniswamy

Advises/Co-

ADVISEES Mentoring role, Resarch area, Accomplishments, Institution

Postdoctoral trainees:

2015-2017

• Nishanth U. Nair, Ph.D.

Research co-advisor Big data and Oncology First-authored scholarship of mentored research University of Maryland • Songpeng Zu, Ph.D. 2020-2021 Research co-advisor Probablistic modelling First-authored scholarship under preparation Harvard University Ph.D. and MS students: • Xiaoman Wang 2018-2020 Research co-advisor Statistical methods Co-first authored publication of mentored research and PhD thesis Peking Union Medical College; Visiting student at DFCI 2020-• Phillip Nicol Ph.D. co-advisor Probablistic modelling First authored publication in submission Harvard School of Public Health • Gege Qian 2020-2021 Master thesis supervisor Single cell analysis Completed MS thesis on Exploit Immunological Information through Single-cell Sequence Analysis and Image Assay Harvard Medical School • Peter Shen 2018 MS supervisor Completed MS thesis on AI-predictor of immunotherapy response using transfer learning Harvard Medical School • Ya Han 2018 Research co-advisor Tonji University • Sailing Shi 2018 Research co-advisor Tonji University • Tong Han 2018 Research co-advisor Tonji University • Binbin Wang 2019

Research co-advisor Epigenetics Tonji University

 $\bullet\,$ Ben Kompa

Summer rotation supervisor

ΑI

Harvard Medical School

• UG students:

Chongming Ni
 Research supervisor
 AI and Immuno-oncology

2018

• Sanna Madan Research supervisor 2017

Rig data

Big data

University of Maryland

 Danielle Paulson 2019 (Summer student) High school RSI Research supervisor Single cell analysis Havard University

Software

• **BTAS**: AI predictor of immunotherapy response. https://rconnect.dfci.harvard.edu/BTAS/

• bipotentR: Method to identify gene targets that kill tumor by two mechanisms concurrently.

http://bipotentr.dfci.harvard.edu/

• MSSC: Bayesian model for robustly identifying differentially expressed genes in single cell data.

https://github.com/JunLiuLab/mssc

- CellPhen: Probablistic model to map immune populations to biological phenotypes such as patient survival or immunotherapy response through bulk tumor transcriptomes. https://github.com/vinash85/CellPhen
- scDiffPop: Single cell probablistic inference of cell populations (e.g. immune cells) perturbed by biological conditions such as disease or perturbation experiments. https://github.com/vinash85/scdiffpop/tree/phil
- **DeepImmune**: AI-predictor of immunomodulatory drugs through perturbation model of tumor immune microenvironment.

 https://github.com/vinash85/deepImmune
- **TRIM**: Statistical approach to identify Transcription Regulator of Immune-Metabolism (TRIM).

https://github.com/vinash85/TRIM

• INCISOR: Identification of synthetic rescue interactions. The genetic interactions determine resistance to cancer targeted and immunotherapy. (Patented) https://github.com/vinash85/INCISOR Sahu-CV5

• GOAL: Bayesian tool to find causal expression regulatory polymorphisms by integration of genetic and epigenetic data. https://www.cbcb.umd.edu/software/goal

- ISLE: Identification of clinically relevant synthetic lethal interactions. https://github.com/jooslee/ISLE
- Hirdaya: SVM predictor of driver genes in idiopathic dilated Image result for cardiomyopathy.
- CellToPhenotype: Machine learning predictor of migration and proliferation potential https://github.com/nishanth83/CellToPhenotype
- ChIPnorm: A statistical method for identifying interesting regions in histone modification libraries.

http://lcbb.epfl.ch/software.html

	Course	Role	Institution & level	Responsibilities
	Big data statistics in genomic and genetic research, 2020	Teaching Assistant	Harvard Grad	Create/presented a lecture.
	Introduction to Computational Biology and Bioinformatics, 2019	Teaching Assistant	Harvard UG	Create/presented a lecture & grade a excercise.
TEACHING EXPERIENCE	Big data and systems biology, 2017	Assistant Instruc- tor	UMD Grad	Grade exercises and class projects. Create/present 4 lectures: TCGA data analysis, Flux balance analysis, and Identify synthetic lethal and synthetic rescue interactions.
	Introduction to Artificial Intelligence, 2011	Teaching Assistant	UMD UG	Homework lectures, Teaching hour assistance, grade weekly excercises.
	Systems biology & genomics, 2018	Guest lecturer	DNA Damage Work- shop	
	Field Programmable Gate Arrays programming,2006	Guest lecturer		
Invited speaker	Droad institute Single call n	ti	.h.: d MA	2022

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Broad institute, Single cell meeting, Cambridge, MA.				
Cancer Center Symposium and Retreat. University of Colorado				
	Seven Bridge and Harvard Center of Mathematical Science and Applications, bridge, MA.	Cam- 2019		
	Dana-Farber Cancer Institute Center Genomic meetup(CFCE), Boston, MA.	2018		
	Immunotherapy response & resistance Weill Cornell Medical College	2017		

Synthetic rescue and immunotherapy resi	istance Tango Therapeutics	2017
Personalized medicine	Weizmann Institute	2016
Personalized medicine	Metabomed Inc. Israel	
Cardiovascular diseases Center for Bioinf 2015	formatics and Computational Biology Ser	minar
Personalized medicine	Tel Aviv University	2015
Epigenetics Center for Bioinformatics	and Computational Biology Seminar	2013
Enhancers associated with hypertension	National human genome research Inst. $$	2012
Hierarchical and multipass segmentation	University of Washington, Seattle	2012
Normalization of ChIP-Seq data	Pacific Science Biocomputing	2012

Consulting

- Lead Pharma(Ongoing)
- Checkmate Pharmaceuticals (Ongoing)
- C-Reveal Therapeutics(Ongoing)
- Metabomed
- Pangea Therapeutics

ADDITIONAL PROFESSIONAL

STMicroelectronics, Imaging Dept, Grenoble, France

PROFESSIONAL Design Architect
EXPERIENCE

May 2007 to August 2009

Responsibilities (Optical image stabilizations, JPEG, Video stabilization):

• Formulation of Architecture & specification of an IP.

Professional Memberships

• Society for Immunotherapy of Cancer (STIC)	since 2020
• International Society for Computational Biology (ISCB)	since 2010
\bullet American Association for Cancer Research (AACR)	since 2015
• Swiss Institute of Bioinformatics (SIB)	since 2010

Professional Services

• Ad hoc reviewer for journals:

Nature

Nature Biotechnology

Cell

Nature Method Bioinfomatics Nature Medicine Nature Communica

Nature Communications Genome Research

Nucleic Acids Research BMC Bioinformatics

IEEE/ACM Transactions on Computational Biology and Bioinformatics

PUBLICATIONS

*Co-corresponding Author,(Reverse chronological)

Appear as "Avinash Das" or "Avinash Sahu" due to a name change

Journal

- Avinash D.*, Xiaoman W., ..., Keith F., Data-driven discovery of targets for bipotent anticancer drugs identifies Estrogen Related Receptor Alpha. Cancer Discovery. (In press) https://www.biorxiv.org/content/10.1101/2021.10.25.465724v1
- Mike M, ..., Avinash D, ..., Benjamin V. A Community Challenge to Predict Clinical Outcomes After Immune Checkpoint Blockade in Non-Small Cell Lung Cancer. Journal of Clinical Oncology. https://doi.org/10.1101/2022.12.05.518667. 2022.
- Xiaoman W., ..., Avinash D.*, INCISOR: An algorithm to identify synthetic rescue mediators of resistance to targeted and immunotherapy. Springer, Methods in Molecular Biology; 2021.
- Elise A., ..., Avinash D., ..., Roberta Z., Supporting the next generation of scientists to lead cancer immunology research. Cancer Immunology Research, 2021.
- David L., ..., Avinash D., ..., Genevieve B., Evolution of Resistance to Immunotherapy in a Melanoma Exceptional Responder. Nature Medicine; 2021.
- Shengqing G,..., Avinash D., ..., X. Shirley L., Therapeutically increasing MHC-I expression potentiates immune checkpoint blockade. Caner Discovery. 2021.
- Bernard P., Cristopher C., ..., Avinash D., ..., Dejan J., Cell-free DNA captures tumor heterogeneity and driver alterations in rapid autopsies with pre-treated metastatic cancer. Nature Communication. 2021 May 27
- Zexian Z., ..., Avinash D., ..., X. Shirley L., Cross-site concordance evaluation of tumor DNA and RNA sequencing platforms for the CIMAC-CIDC network. Clinical Cancer Research. 2020
- Shengqing G., ..., Avinash D., ..., X. Shirley L., Clonal tracing reveals diverse patterns of response to immune checkpoint blockade. Genome Biology. 2020
- Jian Z., .., Avinash D., ..., X. Shirley L. Immune receptor repertoires in pediatric and adult acute myeloid leukemia. Genome Medicine. 2020
- Ziyi L., ..., Avinash D.,.., X. Shirley L. CRISPR Screens Identify Essential Cell Growth Mediators in BRAF Inhibitor-resistant Melanoma. Genomics, Proteomics & Bioinformatics. 2020
- Gaurav P.,..., Avinash D., ..., Zeev Ronai, Translational Reprogramming Marks Adaptation to Asparagine Restriction in Cancer. Nature Cell Biology. 2019
- Alicia H., ..., Avinash D. ..., Sridhar H. Allele-specific enhancers mediate associations between LCAT and ABCA1 polymorphisms and HDL metabolism. PLoS one. 2019.
- Cristopher C., Avinash D., ..., Dejan J., Plasma sequencing demonstrates that breast cancer patients have a higher prevalence of clonal and multiple PIK3CA mutations than other solid tumor patients. CANCER RESEARCH.
- Yan L., ..., Avinash D., ..., Zeev R., Gut microbiota dependent anti-tumor immunity restricts melanoma growth in Rnf5-null mice. Nature Comm. 2019.
- Joo Sang L., Avinash D., ..., Eytan R. Harnessing synthetic lethality to predict clinical outcomes of cancer treatment. *Nature Communication*, 2019.

Avinash D*, ...,Eytan R* Genome-wide prediction of synthetic rescue mediators of resistance to immunotherapy. Molecular Sys. Bio., 2019. *Corresponding author

- Assaf M, Avinash D., ..., Sridhar H. Charting the landscape of pairwise gene expression states associated with survival in cancer. Cell Reports, 2019.
- Avinash D., Joo S.L, Eytan Ruppin. Computer System And Methods For Harnessing Synthetic Rescues And Applications Thereof. 2019
- Peng J., ..., Avinash D., .., X. Shirley L., Signatures of T-cell dysfunction and exclusion predict cancer immunotherapy response. Nature Medicine. 2018
- Nishanth N., Avinash D., ..., Eytan R., Cell migration is a stronger predictor of patient survival in breast cancer than cell proliferation. Scientific Report. 2018
- Nishanth N., Avinash D., .., Sridhar Hannehalli, Putative functional genes in idiopathic dilated cardiomyopathy. Scientific Reports, 2018
- Gaurav P.,... Avinash D., ..., Zeev R. Targeting the Warburg Effect via LDHA inhibition engages ATF4 signaling for cancer cell survival. The EMBO, 2018.
- David L., ..., Avinash D., .., Genevieve B., Phylogenetic analysis of longitudinal melanoma samples to reveal convergent evolution and markers of immunotherapy resistance. *Journal of Clincal Oncology*, 2018.
- Kun W, ..., Avinash D., ..., Sridhar H., Comprehensive map of age-associated splicing changes across human tissues and their contributions to age-associated diseases. Scientific reports, 2018
- Shrutii S, Avinash D., ..., Sridhar H. Distal CpG islands can serve as alternative promoters to transcribe genes with silenced proximal-promoters. *Genome Research*, 2017.
- Uri Amit, ..., Avinash D., ..., Jonathan L., A New Role for Interleukin-13 Receptor in Myocardial Homeostasis and Heart Failure. *Journal of the American Heart Association*, 2016.
- P Cardro,..., Avinash D.,.., Euan A., A community overlap strategy reveals central genes and networks in heart failure. https://doi.org/10.1101/038174, BioArxiv.
- Mahashweta Basu, ..., Avinash D., Sridhar Hannenhalli. Prediction and Subtyping of Hypertension from pan-tissue transcriptomic and genetic analyses. *Genetics*, 2016.
- Kun W., Avinash D., ..., Sridhar H. "Phenotype-dependent coexpression gene clusters: application to normal and premature ageing." *Computational Biology and Bioinformatics*, *IEEE/ACM Transactions* on 12, no. 1 (2015): 30-39.
- Nishanth N., ..., Avinash D., ..., Bernard M, Study of cell differentiation by phylogenetic analysis using histone modification data. BMC bioinformatics. 2014

Avinash D*, M Morley, Moravec CS, Tang WHW, Hakonarson H, Margulies KB et al. Bayesian integration of genetics and epigenetics detects causal regulatory SNPs underlying expression variability. *Nature Communications*. 2016 Oct 12;6. 8555.

- Avinash D.,..., Sridhar H., Epigenomic model of cardiac enhancers with application to Genome wide association studies. *PSB*, 2013.
- Kun W., Avinash D., ... Sridhar H.: Identification of gene clusters with phenotype-dependent expression with application to normal and premature ageing. ACM BCB, 2013.
- Nishanth N., Avinash D., ..., Bernard Moret. ChIPnorm: A statistical method for identifying interesting regions in histone modification libraries. *PLoS ONE* 2012.
- ENCODE Project Consortium. Integrative analysis of the human genome by the ENCODE Project. Nature, 2012.

Non-peer reviewed Conferences

- Gaurav P., ..., Avinash D., ..., Zeev R., Co-targeting MAPK signaling and asparagine metabolism in pancreatic cancer, Gastroenterology, 2020.
- Avinash D., Xiaoman W, ..., X. Shirley L. Identification of drugs that synergize with immunotherapy. AACR Annual Meeting, 2020.
- Genome-wide prediction of synthetic rescue mediators of resistance to targeted and immunotherapy. Cancer Research 2019.
- Signatures of T-cell dysfunction and exclusion predict cancer immunotherapy response. Cancer Immunology Research 2019.
- Harnessing genetic interactions to advance whole-exome precision cancer treatment. ISMB 2019.
- Genome-wide prediction of synthetic rescue mediators of resistance to targeted and immunotherapy. AACR Annual Meeting 2019.
- Harnessing genetic interactions to advance whole-exome precision cancer treatment. AACR Annual Meeting 2019.
- Avinash D., Data-driven approach to detecting novel gene interactions in cancer with applications to drug response prediction and cancer stratification. *Cancer Research* 2017.
- Harnessing synthetic lethality to predict clinical outcomes of cancer treatment. *Molecular Cancer Therapeutics* 2017.
- Cell migration is a stronger predictor of patient survival in breast cancer than cell proliferation. *Molecular Targets Conference* 2017.
- Avinash D., et al. "Genome-wide identification of gene determinant of cancer resistance" AACR Annual Meeting (2017).
- Avinash D., et al. "Tracing synthetic rescue reprogramming to counteract cancer resistance." *Molecular Cancer Therapeutics* 14.12 Supplement 2 (2015): B56-B56.
- Assaf M., Avinash D., Sridhar H., Data-driven approach to detecting novel gene interactions in cancer with applications to drug response prediction and cancer stratification. *Cancer Research*, 2017

Avinash D., Eytan R., Harnessing synthetic rescues to evaluate and mitigate resistance to cancer therapy. 2017, Cancer Research.

- Y Li, ..., Avinash D., ..., Zeev R., Coordinated regulation of gut microbiota and immune checkpoint activity by the ubiquitin ligase RNF5 controls melanoma growth. FEBS JOURNAL, 2017.
- Paula D, ..., Avinash D., ..., Raya A., Co-expression of immune checkpoint mRNAs and miR-15a and miR-15b in bladder cancer—highlighting potential novel targets and approaches for immunotherapy. 2017.
- Avinash D., Sridhar H.. Dirichlet process model for joint haplotype inference and GWAS. BMC Bioinformatics. 2012
- Manish P., Avinash D., Mohammed H., Graph Balancing: A special case of scheduling on unrelated machines. UMD Annals. 2012
- Avinash Sahu, Sridhar Hannenhalli. Bayesian integration of genetics and epigenetics detects causal regulatory SNPs underlying the cardiac failure. *Gordon Research Conference*. Human Nucleotide Polymorphism and Diseases. 2014 Aug.
- Avinash D. Sahu, Ludovic Chotard. Architecture and hardware implementation of Relative Motion Vector estimation of the image. *IP based Electronic System Conference & Exhibition*. Grenoble 2008.

Manuscript in preparations

- *Corresponding Author
- Phillip Nicol., ..., Rafael I*, Aviansh D.*. Robust cell prioritization method for single-cell transcriptome revealed role of LEF1+ T helper cells in anti-PD1 response (In revison, Nature Biotech, available upon request).
- Murad M., Avinash D.,..., Alexander M., Genome-wide CRISPR screens reveal metabolic and transcriptional regulation of butyrophilin 3A and susceptibility to $V\gamma9V\delta2$ T cell targeting. (In review Nature)
- Avinash D.*, Songpeng Zu, ..., X. Shirley L., Probablistic inference of functional role of immune populations in patient prognosis reveals $\gamma\delta$ T cells essential for healthy immune system.

Patents

- Avinash D., Joo Sang L., Eytan R. Computer system and methods for harnessing synthetic rescues and applications thereof. 2016
- Avinash D., Ludovic C. Architecture and hardware implementation of Relative Motion Vector estimation of the image.