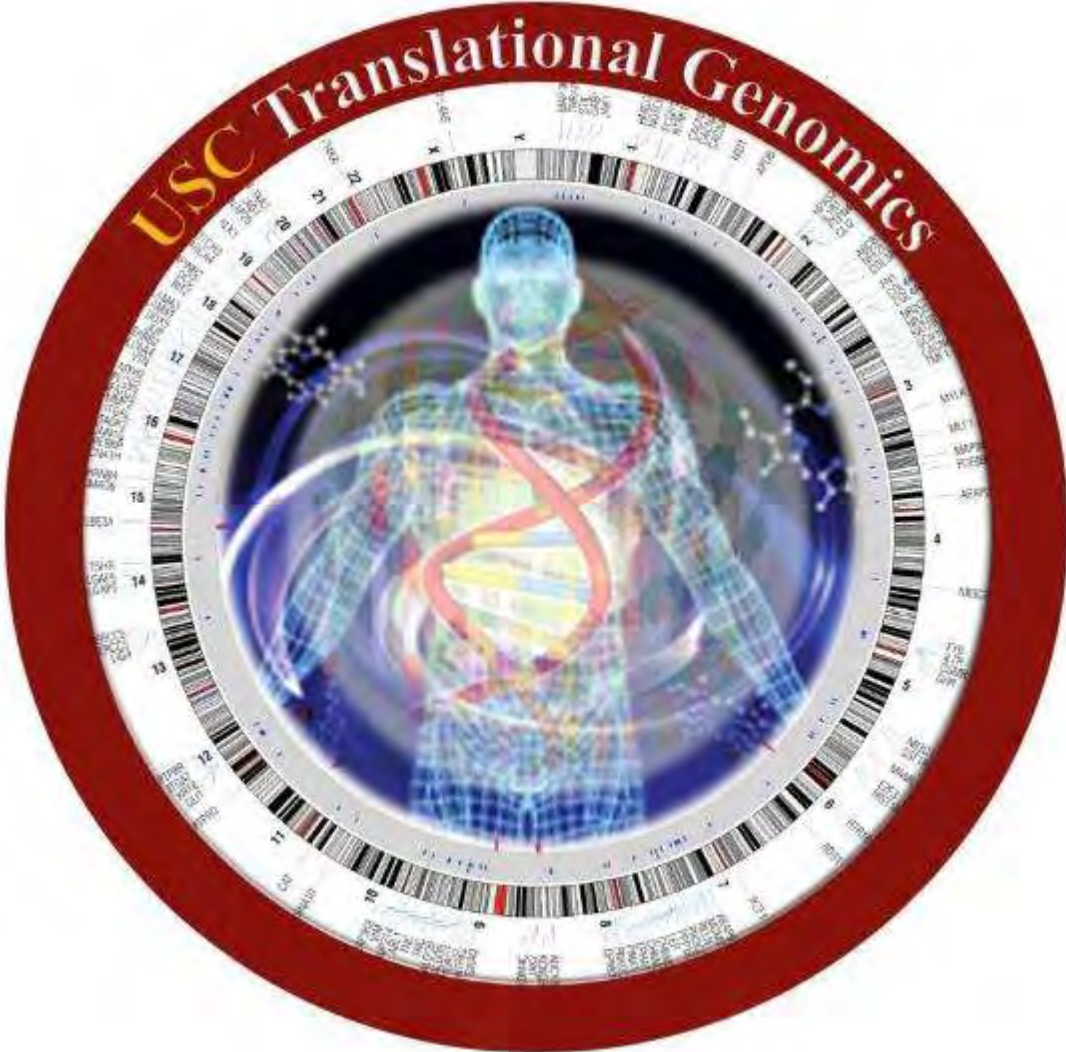


Accelerating improvements in healthcare at USC KSOM through intentional integration between basic discovery research and clinical science!



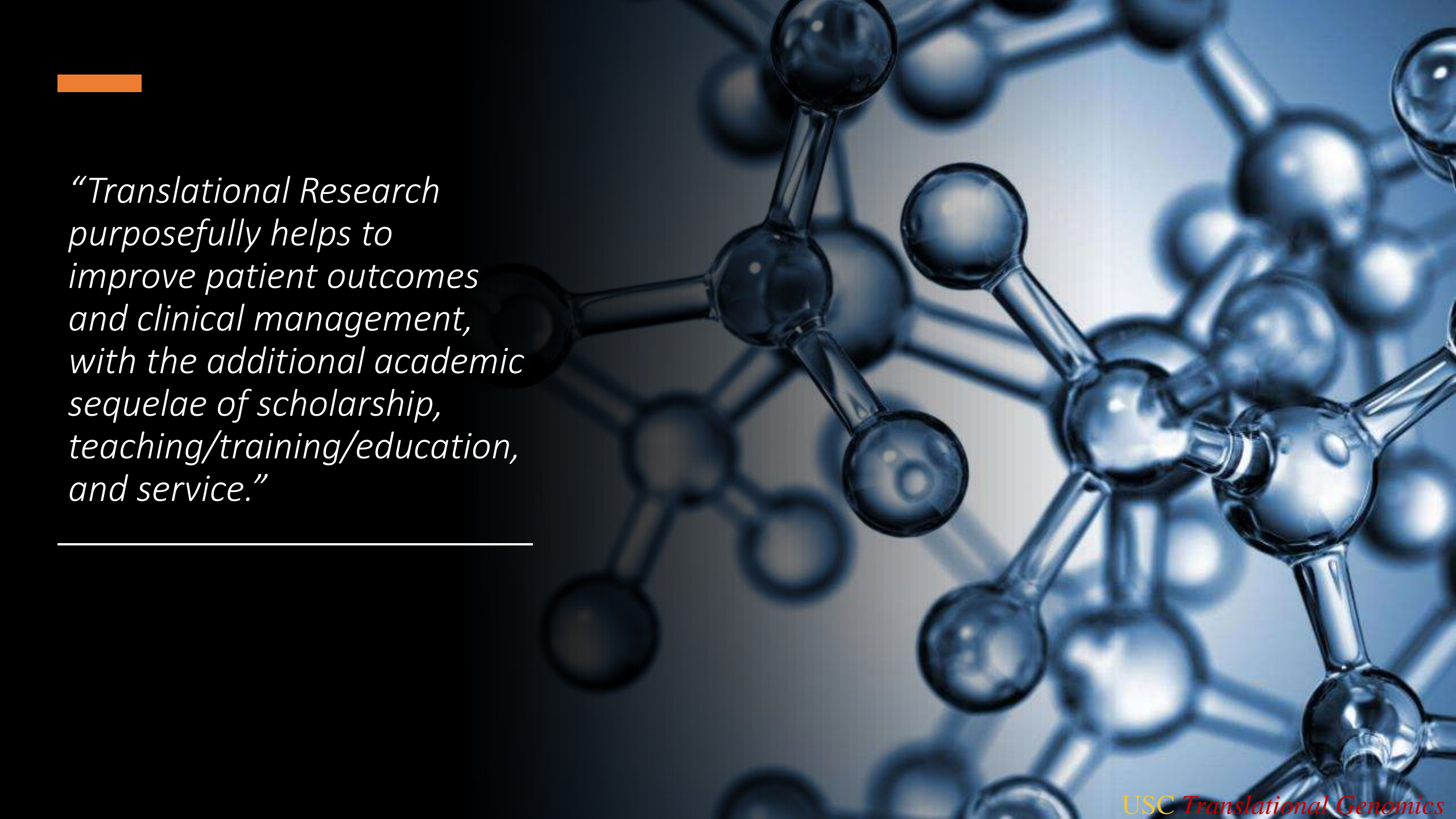
USC Translational Genomics Leadership



John D. Carpten, PhD
Chair



David W. Craig, PhD
Vice Chair



“Translational Research purposefully helps to improve patient outcomes and clinical management, with the additional academic sequelae of scholarship, teaching/training/education, and service.”

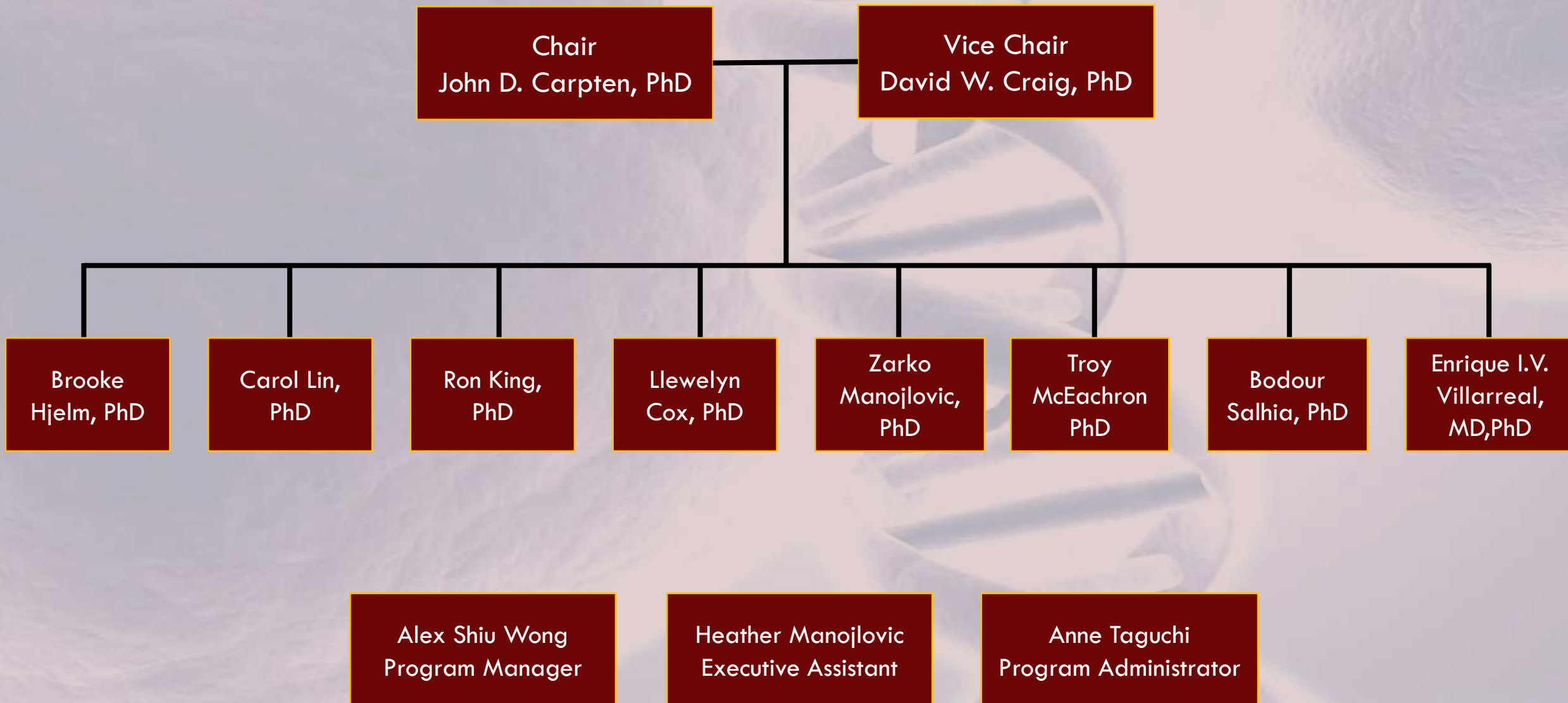
Mission Statement

Drive genome science towards a deeper understanding of human disease processes to improve patient management and outcomes.

- Basic discovery and clinical translational research.
- Training and education.
- Grow status of the KSOM and NCCC in the area of genome sciences.
- Strong commitment to diversity



Organizational Chart



Research Platform

– Genome science

- Germline genetics and risk (Cancer and Rare Genetic Disorders)
- Somatic cancer interrogation (discovery and clinical relevance)

– Bioinformatics

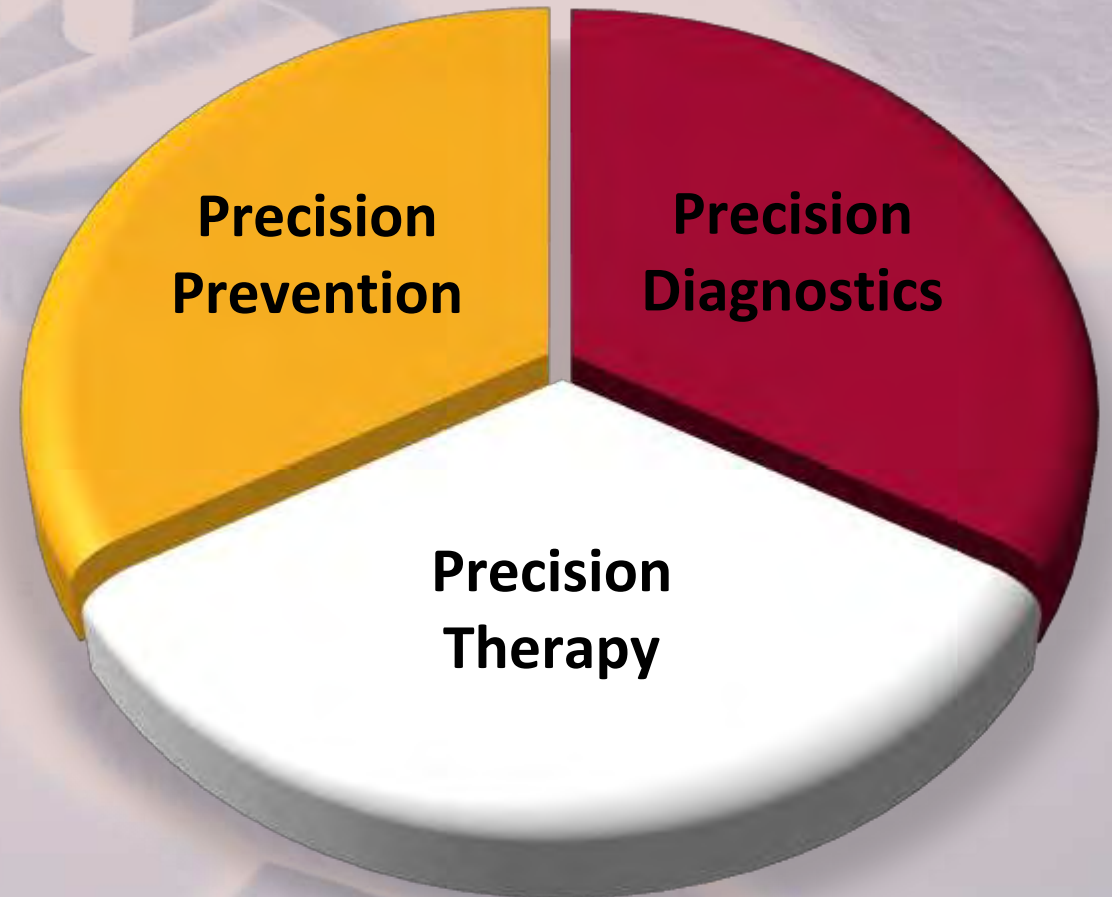
- Molecular Genomic Data Processing and Analysis
- Integration with Oncology Pharmacopeias
- Integration with Clinical Data Parameters

– Cell biology and Functional genomics

- Target validation
- Functional and cell-based screens
- PDX model development

– Applied Epigenetics

- Developmental regulation
- Prediction of metastasis



Educational and Training Platform

- **Translational Biotechnology Masters Degree Program**
- **Translational Biomedical Informatics Masters Degree Program**
- **Undergraduate Student Research Training**
- **Medical Student Education**
- **PIBBS PhD student Training**
- **Fellowship Training**
 - **Postdoctoral Fellows**
 - **Clinical Fellows**
 - **Fulbright Fellows**

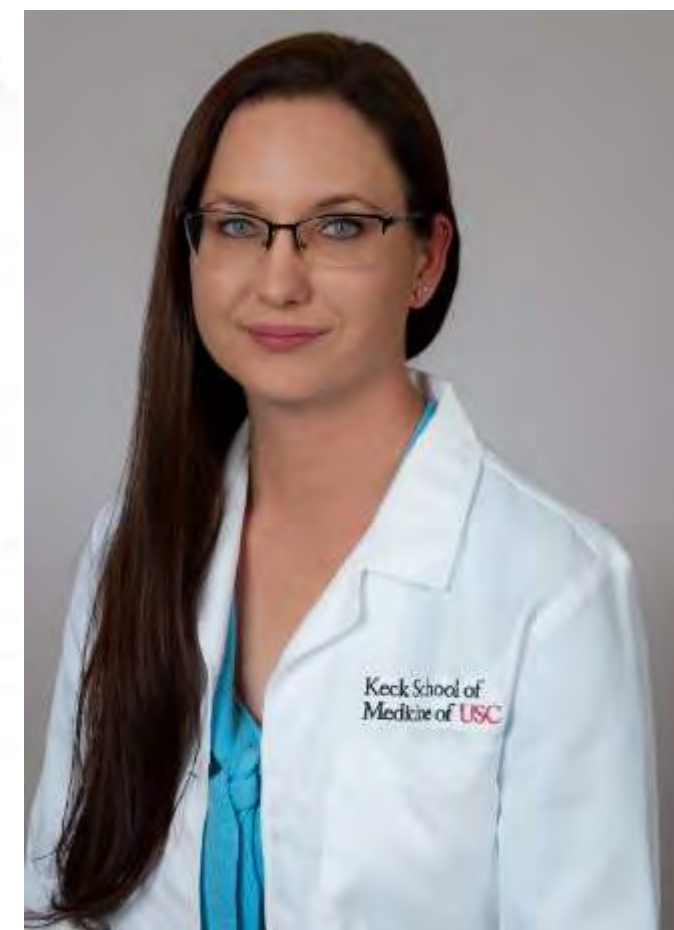


Advanced Genomics Technologies





Scholarship



Brooke E. Hjelm, PhD
Assistant Professor

Splice-Break: exploiting an RNA-seq splice junction algorithm to discover mitochondrial DNA deletion breakpoints and analyses of psychiatric disorders

Brooke E. Hjelm^{1,2,*}, Brandi Rollins¹, Ling Morgan¹, Adolfo Sequeira¹, Firoza Mamdani¹, Filipe Pereira³, Joana Damas⁴, Michelle G. Webb², Matthieu D. Weber¹, Alan F. Schatzberg⁵, Jack D. Barchas⁶, Francis S. Lee⁶, Huda Akil⁷, Stanley J. Watson⁷, Richard M. Myers⁸, Elizabeth C. Chao⁹, Virginia Kimonis⁹, Peter M. Thompson¹⁰, William E. Bunney¹ and Marquis P. Vawter^{1,7}

¹Department of Psychiatry and Human Behavior, University of California-Irvine (UCI), Irvine, CA 92697, USA, ²Department of Translational Genomics, Keck School of Medicine of USC, University of Southern California (USC), Los Angeles, CA 90033, USA, ³Interdisciplinary Centre of Marine and Environmental Research (CIIMAR), University of Porto, Matosinhos 4050-123, Portugal, ⁴The Genome Center, University of California-Davis, Davis, CA 95616, USA, ⁵Department of Psychiatry and Behavioral Sciences, Stanford University, Stanford, CA 94305, USA, ⁶Department of Psychiatry, Weill Cornell Medical College at Cornell University, New York, NY 10085, USA, ⁷The Molecular and Behavioral Neuroscience Institute, University of Michigan, Ann Arbor, MI 48109, USA, ⁸HudsonAlpha Institute for Biotechnology, Huntsville, AL 35806, USA, ⁹Division of Genetics and Genomic Medicine, Department of Pediatrics, UCI, Irvine, CA, USA and ¹⁰Southwest Brain Bank, Department of Psychiatry, Texas Tech University Health Sciences Center (TTUHSC), El Paso, TX 79905, USA

Received January 29, 2019; Editorial Decision February 25, 2019; Accepted February 28, 2019

My research focuses on gene expression and DNA variation in complex neurological disorders. Our lab works on both **neurodegenerative disorders** like Alzheimer's Disease and Parkinson's Disease, as well as **psychiatric disorders** like Schizophrenia and Major Depressive Disorder. We have a strong focus in **mitochondrial genetics** and how mitochondrial DNA (mtDNA) structural variation contributes to both aging and disease. In addition, we utilize **spatial transcriptomics** and **single-cell sequencing methods** to evaluate postmortem human brain tissue from subjects with these debilitating diseases to assess gene expression and DNA variation in specific neuron/glia cell types and/or cortical layers of the brain.



Can changes in brain energy pathways cause depression?


New research has identified mutations in the DNA code that may affect energy metabolism. It also found a link to major depressive disorder.



The World Health Organization (WHO) describe depression as "the leading cause of disability worldwide".

It affects more than 300 million people around the world.

ORIGINAL RESEARCH

 Check for updates

Profiling targetable immune checkpoints in osteosarcoma

Troy A McEachron^{a,b,c}, Timothy J Triche^{b,d}, Laurie Sorenson^a, David M Parham^d, and John D Carpten^{a,b}

^aDepartment of Translational Genomics; ^bNorris Comprehensive Cancer Center; ^cDepartment of Pediatrics, Keck School of Medicine of the University of Southern California, Los Angeles, CA, USA; ^dDepartment of Pathology, Children's Hospital Los Angeles, Los Angeles, CA, USA



Troy McEachron, PhD
Assistant Professor of Research

nature
genetics

BRIEF COMMUNICATIONS

Somatic histone H3 alterations in pediatric diffuse intrinsic pontine gliomas and non-brainstem glioblastomas

St. Jude Children's Research Hospital–Washington University Pediatric Cancer
Genome Project

My research focuses on two separate yet complimentary areas: (1) the utilization of next generation genomic, transcriptomic, and proteomic technologies to profile recurrent and/or refractory cancer patients for clinical decision making; (2) the use of functional genomics to identify and interrogate the developmental and therapeutic aspects of cancers, with a focus on sarcomas, which predominantly arise in the pediatric and AYA populations. Moreover, our laboratory is interested in uncovering and understanding the immune microenvironment of cancers using novel molecular technologies in hopes of exploiting these data to identify smarter approaches for the therapeutic management of cancer.



RESEARCH ARTICLE

Comprehensive molecular profiling of 718 Multiple Myelomas reveals significant differences in mutation frequencies between African and European descent cases

Zarko Manojlovic^{1,2*}, Austin Christofferson², Winnie S. Liang², Jessica Aldrich², Megan Washington², Shukmei Wong², Daniel Rohrer³, Scott Jewell³, Rick A. Kittles⁴, Mary Derome⁵, Daniel Auclair⁵, David Wesley Craig¹, Jonathan Keats², John D. Carpten^{1,2}

“The present moment is filled with joy and happiness. If you are attentive you will see it.”

-Thich Nhat Hanh



Innovative Tools and Methods | Full Access

Cancer transcriptomic profiling from rapidly enriched circulating tumor cells

Gareth J Morrison, Alexander T Cunha, Nita Jojo, Yucheng Xu, Yili Xu, Eric Kwok, Peggy Robinson, Tanya Dorff, David Quinn, John Carpten, Zarko Manojlovic, Amir Goldkorn



Zarko Manojlovic, PhD
Assistant Professor of Research
Director Keck Genomics Platform

As Director of Keck Genomics Platform (high-throughput sequencing center), my goal is to provide cutting-edge genomic service to Keck School of Medicine, greater USC, and its affiliates with complete high throughput next-generation sequencing workflows. In this role, my focus is to build a collaborative environment and provide team-driven research support within the scope of the next-generation technologies. In my role as an Assistant Professor of Research, my primary interest is in delineating the molecular interplay of tumor heterogeneity and microenvironments in metastatic disease to help guide clinical decision-making. Furthermore, building on my fellow training, I am interested in interrogating the ancestry effects on cancer progression and outcomes in multi-ethnic prostate tumors.



Bodour Salhia, PhD
Associate Professor

Legendre et al. *Clinical Epigenetics* (2015) 7:100
DOI 10.1186/s13148-015-0135-8



CLINICAL
EPIGENETICS

RESEARCH

Open Access



Whole-genome bisulfite sequencing of cell-free DNA identifies signature associated with metastatic breast cancer

Christophe Legendre[†], Gerald C. Gooden[†], Kyle Johnson, Rae Anne Martinez, Winnie S. Liang and Bodour Salhia^{*}

70

Neuro-Oncology

22(1), 70–83, 2020 | doi:10.1093/neuonc/noz137 | Advance Access date 21 August 2019

Patient-derived xenografts of central nervous system metastasis reveal expansion of aggressive minor clones

Ben Yi Tew, Christophe Legendre, Mark A. Schroeder, Tim Triche Jr, Gerald C. Gooden, Yizhou Huang, Loren Butry, Daniel J. Ma, Kyle Johnson, Rae Anne Martinez, Mariaelena Pierobon, Emanuel F. Petricoin, Joyce O'Shaughnessy, Cindy Osborne, Coya Tapia, David N. Buckley, Jennifer Glen, Mark Bernstein, Jann N. Sarkaria, Steven A. Toms, and Bodour Salhia

My research centers around the development and application of cutting-edge epigenomic technologies. I have been working on developing liquid biopsies using cell-free DNA methylation for the indications of minimal residual disease in breast cancer, bladder cancer and for the early detection of ovarian cancer. My lab has also been studying the underlying genomic underpinnings of brain metastasis and developed preclinical models to study novel treatment interventions.

RESEARCH



NEUROGENOMICS

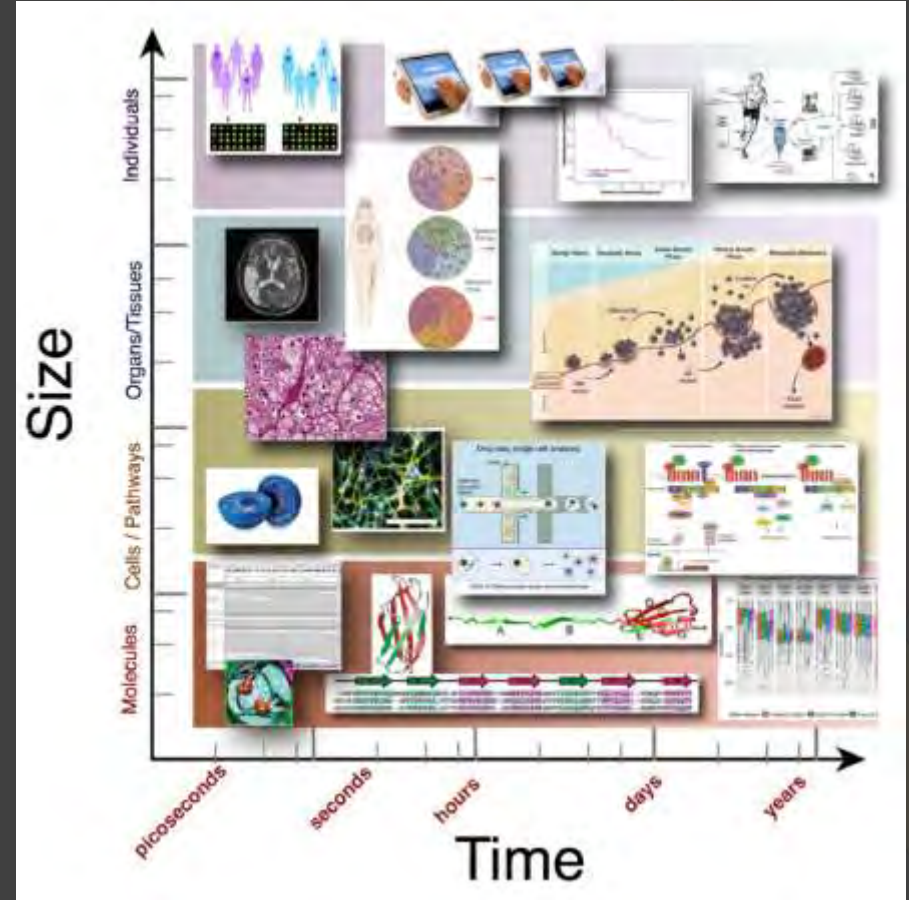
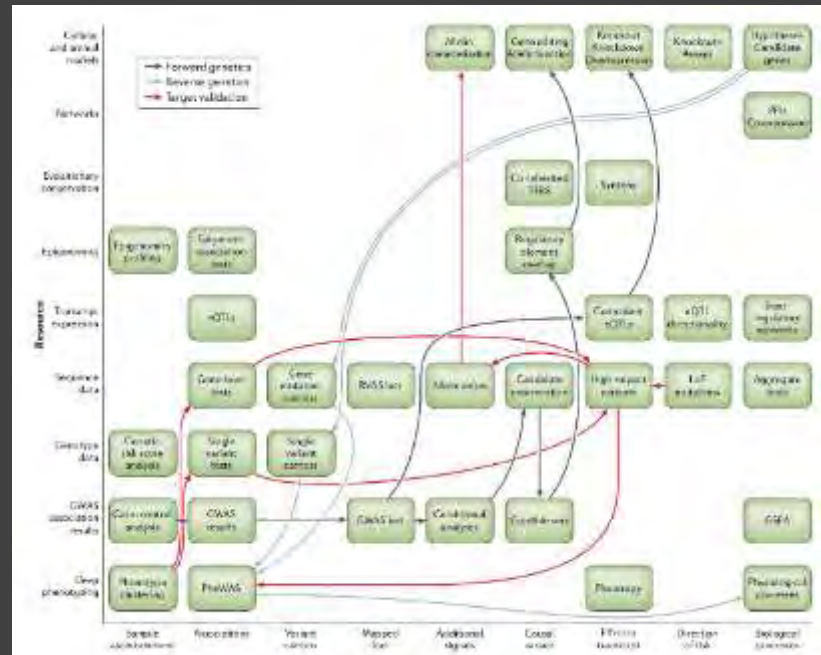
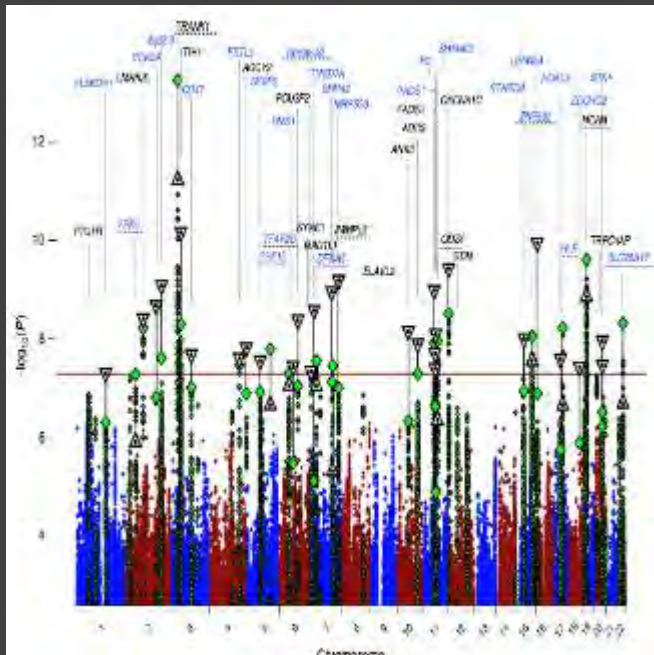
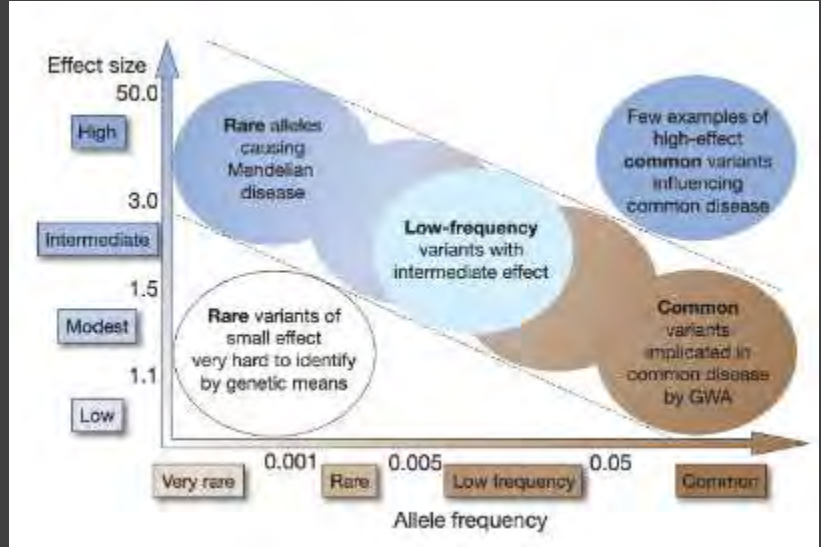
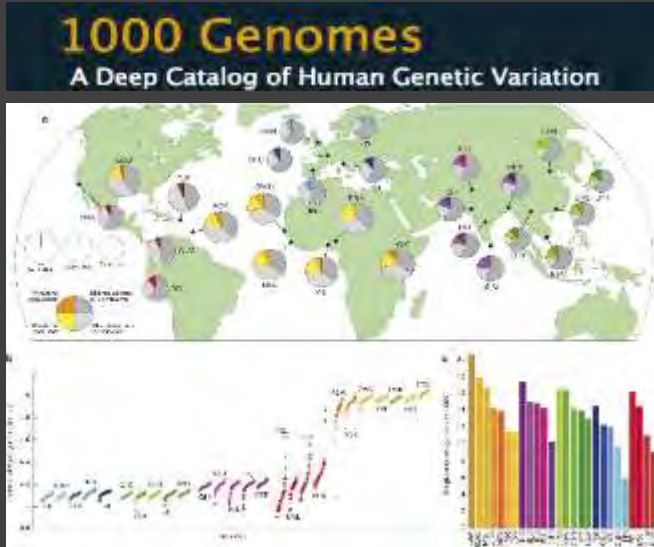
ONCOLOGY GENOMICS

David W. Craig, PhD
Professor and Vice Chair

APPLIED BIOINFORMATICS

GENOMIC TECHNOLOGY

TRANSLATION GENETIC BASIS OF DISEASE; TRANSLATING DNA -> RNA



Role of Integrative Multi-Scale Analysis Of RNA



David W. Craig, PhD
Professor and Vice Chair



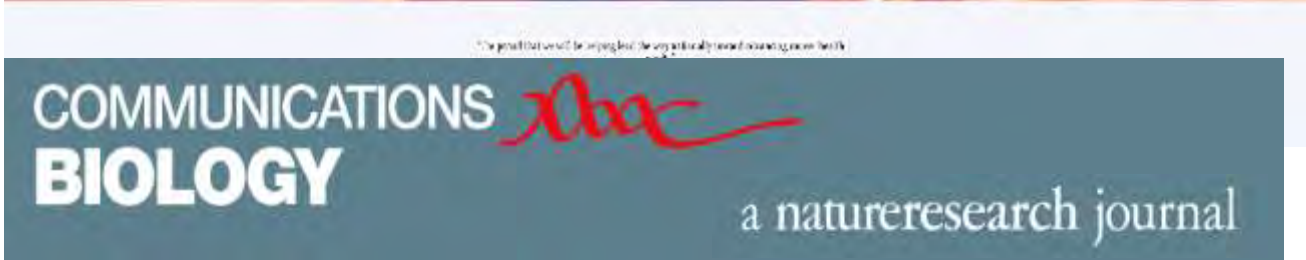
Carmel McCullough, B.S.

Utilizing RNA and Outlier Analysis to Identify an Intronic Splice-Altering Variant in AP4S1 in a Sibling Pair with Progressive Spastic Paraplegia

McCullough et al., Human Mutation. 2020 Feb;41(2):412-419.

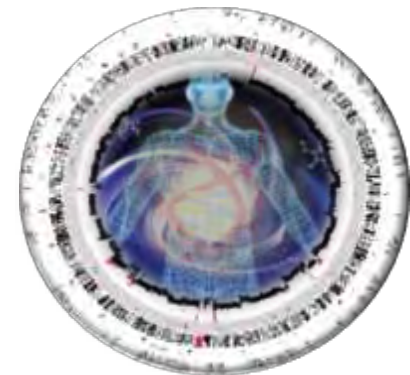
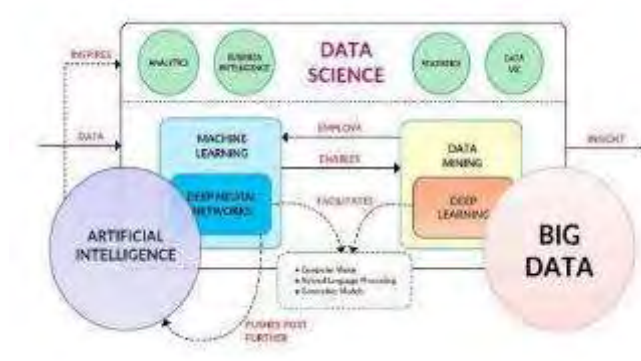
Whole exome sequencing with the inclusion of mRNA-seq allowed for the identification of a previously missed splice-altering variant, and thereby expands the mutational spectrum of AP-4 Deficiency Syndrome to include impacts to some tissue-dependent isoforms.





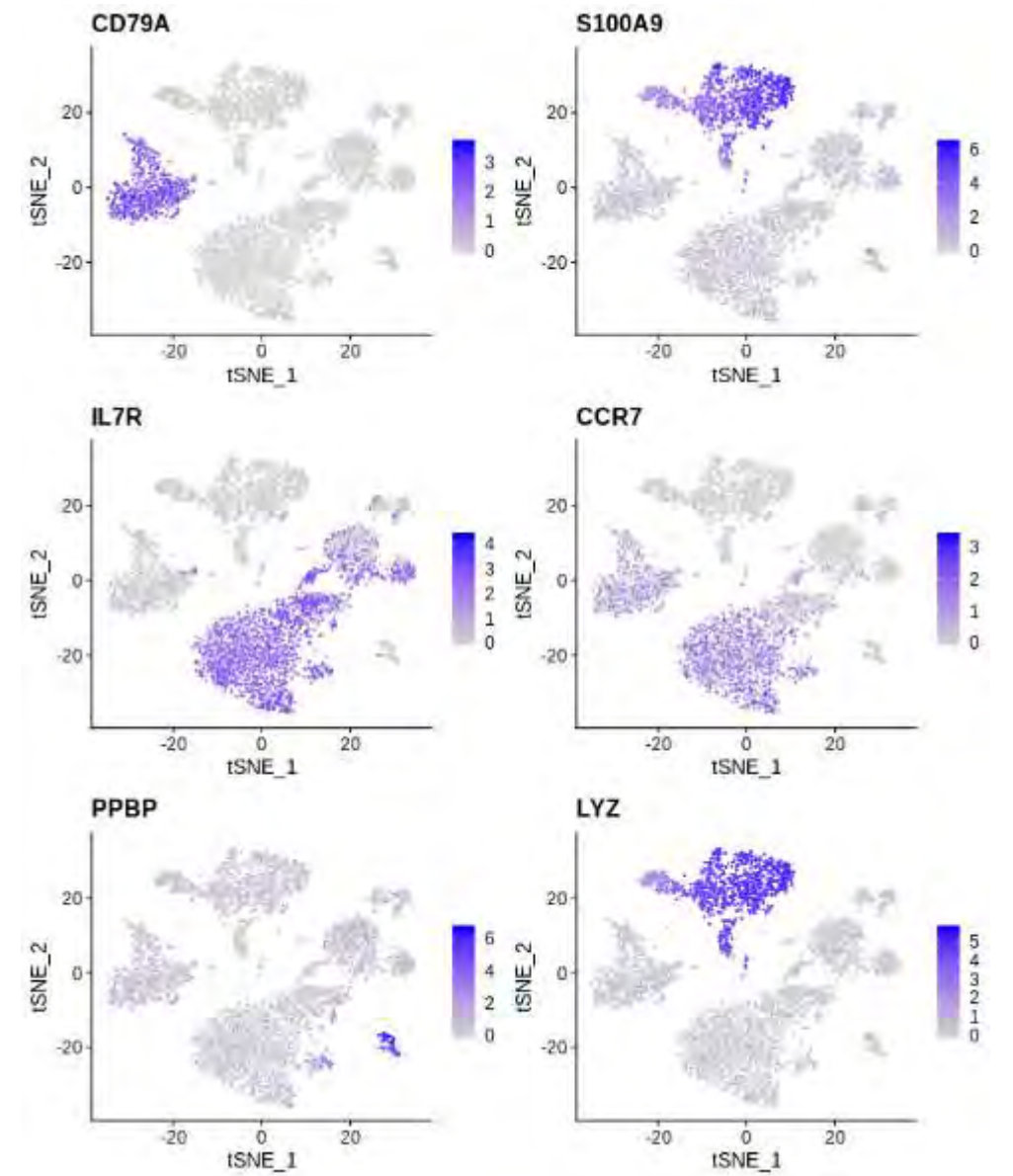
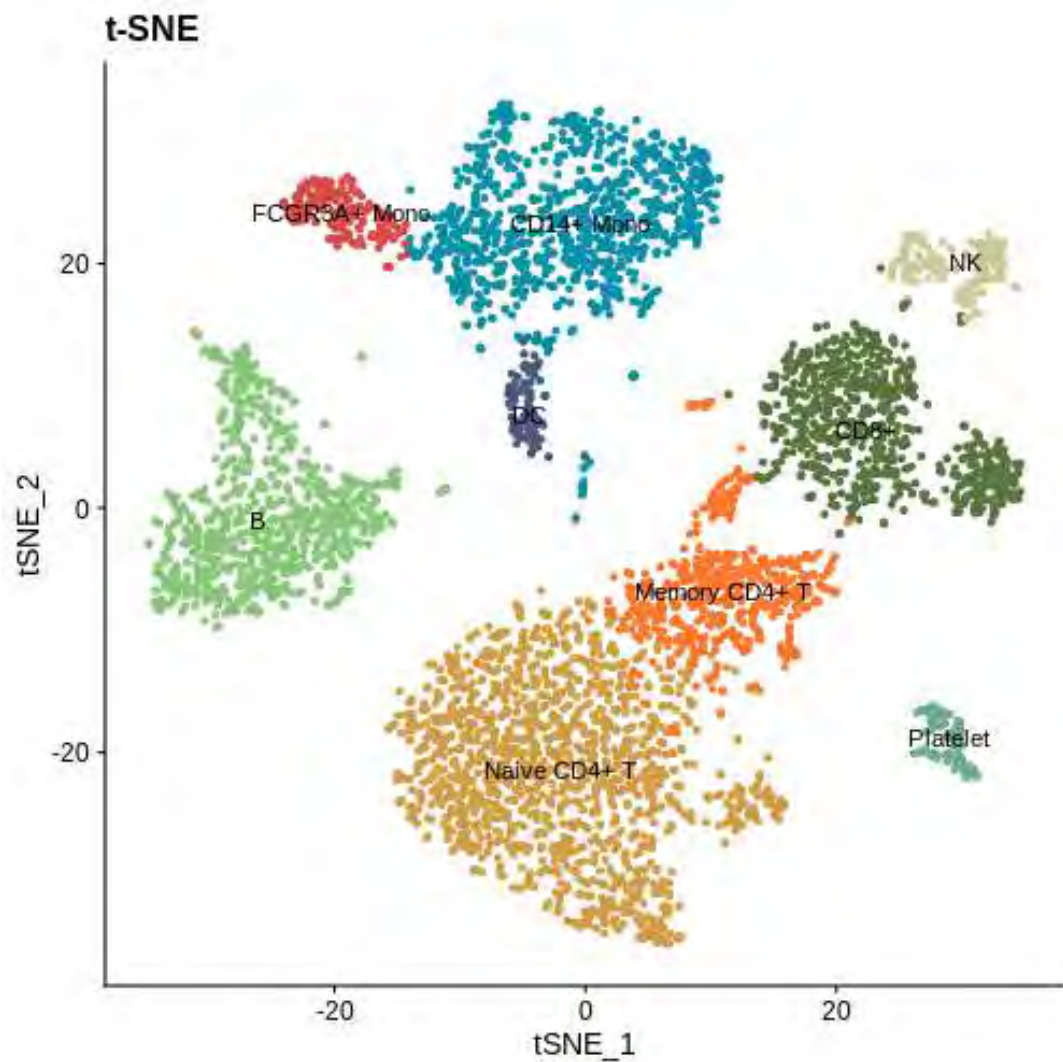
Single-cell sequencing of genomic DNA resolves sub-clonal heterogeneity in a melanoma cell line

My research involves two main components. First, it performs statistical and computational genomics analyses, also by developing novel bioinformatics methods, using cutting-edge genomic technologies with emphasis in single cell to identify molecular/genomic alterations in cancer health disparities, with strong emphasis on gynecological cancers from Latinos and African American populations. Second, it integrates clinical and genomic data by using Big Data science management systems and Artificial Intelligence/Machine Learning to uncover clues for treating cancer, immunological and neurological diseases.



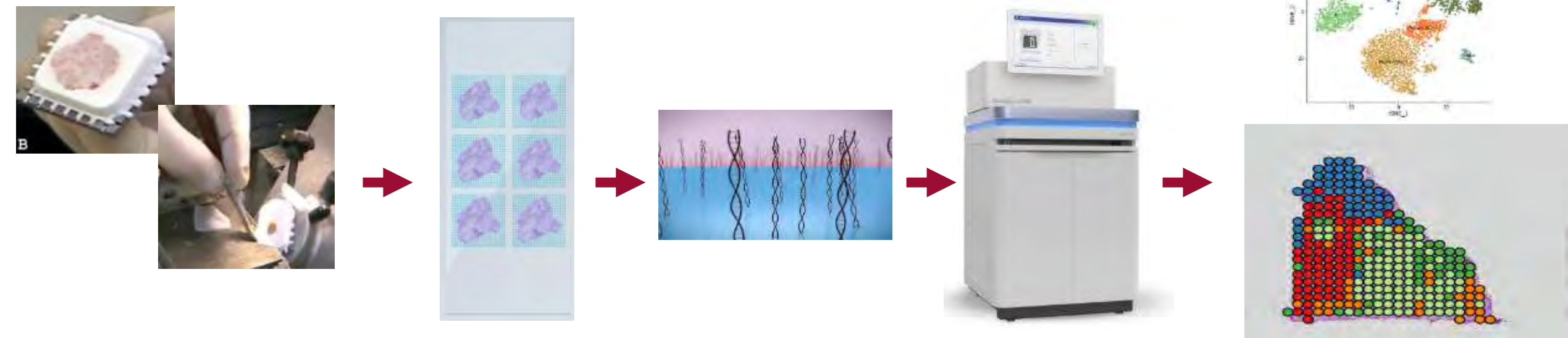
Enrique I. Velazquez Villarreal, MD, PhD, MPH, MS
Assistant Professor

Single Cell Technologies



Gene expression level shown on t-SNE plot

Spatial Transcriptomics





Lynda Roman



John Carpten



David Craig



Lee Gibbs



Diana Da Silva



Troy McEachron



Laila Muderspach

Saloni Walia
Moli Chen
Alex Trana
David Sands
USC GTFR



Heather Miller



Solomon Rotimi



Rania Bassiouni



Yifeng Yin



Neil Weisenfeld



Nigel Delaney



Stephen Williams



Erik Borgstrom



Amy Tam



Edwin Hauw

Studies underway:

- Ovarian Cancer
- Brain Tissue/Neurological
- Triple Negative Brain Cancer (African American)
- Brain Tumors