



Spatial Multi-Omics for Cancer Systems Biology Virtual Workshop

February 12, 2024



**NATIONAL
CANCER
INSTITUTE**

<https://isbscience.org/spatial-multiomics-workshop/#agenda>

Jim Heath

President Institute for Systems Biology

Professor Univ of Washington

Conflicts: Scientific Advisory Board: AtlasXomics and Nanostring

Paid Consultant: Regeneron

Research Funding in past 5 years from Merck, Gilead



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Put your questions in the chat!

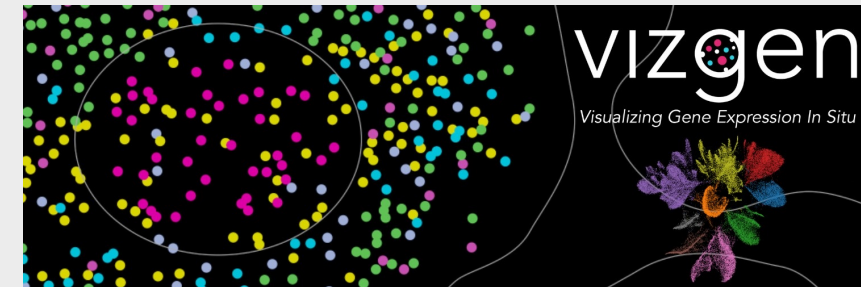


Dominic Lewis



Allison Kudla

Thanks to our sponsors



Session One: 8:30 PT

Experimental and Computational Methods I

Speaker: Rong Fan, PhD

Sequencing-based spatial multi-omics mapping

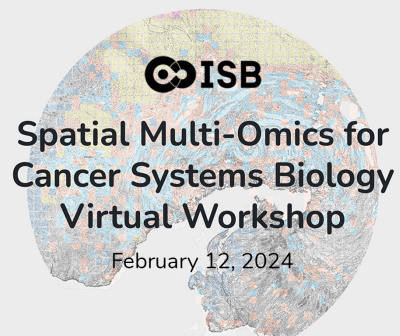
Speaker: Ruben Dries, PhD

The GIOTTO software tools

Speaker: Lu Wei, PhD

Spatial metabolic imaging

15 min Q&A roundtable



Session Two 10:00 PT

Experimental and Computational Methods II

Chair: Vésteinn Thorsson, PhD

Introduction and relating spatial features across assay Platforms

Speaker: Robert Krueger, PhD

Visual analytics for imaging-based spatial biology profiling

Speaker: Joseph Beechem, PhD

Spatial molecular imaging of FFPE cancer samples at any spatial-plex allows true systems biology understanding

Speaker: Lyla Atta

Evaluating normalization approaches for imaging-based spatial profiling technologies

15 min Q&A roundtable

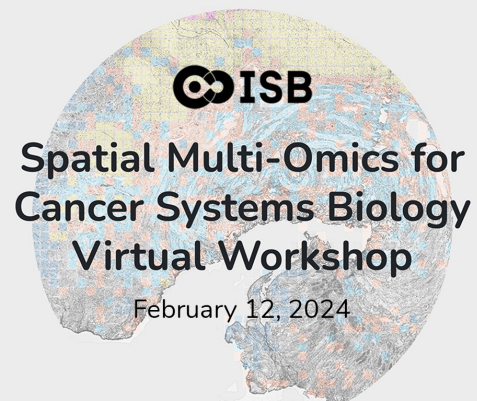
Session Three (12:00 PT):
Applications of Spatial Multi-Omics

Chair: Wei Wei, PhD
Introduction to applications of spatial multiomics

Speaker: Katie Campbell, PhD
Applications of spatial multiomics tools

Speaker: Sandro Santagata, MD, PhD
Applications of spatial multiomics tools

15 min Q&A roundtable



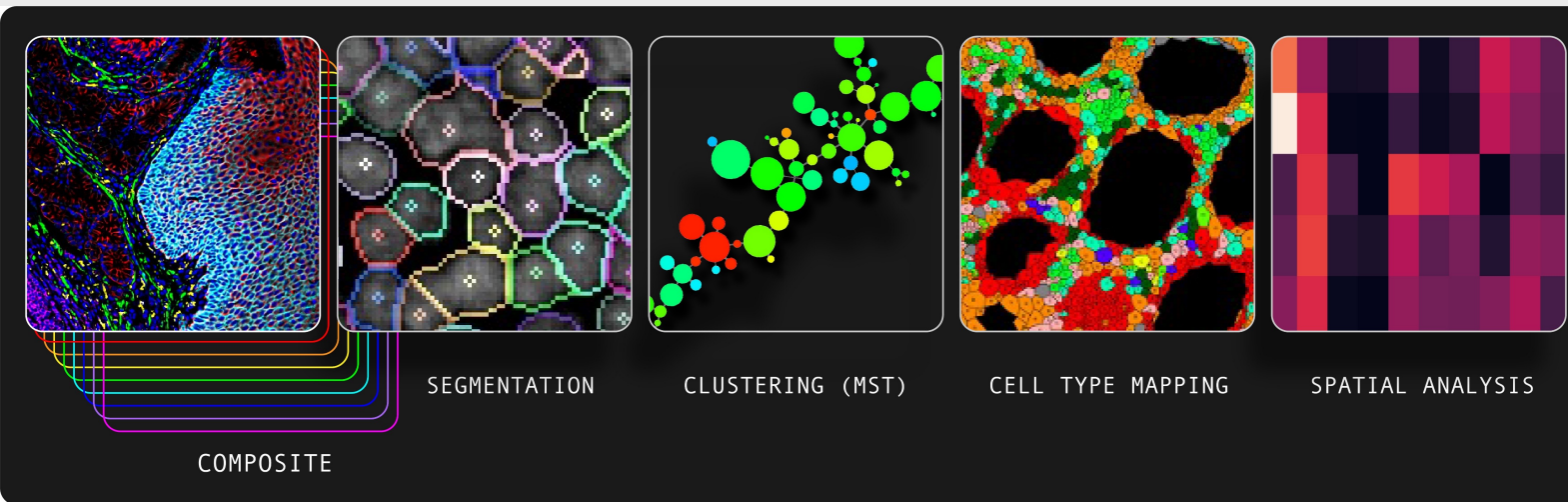
Session Four (13:15 PT):
Practical Methods in Spatial Omics

Rong Fan his group members from Yale University
Shuozhen Bao
Alev Baysoy
Zhiliang Bai

lead a practical tutorial with Jupyter notebooks and GitHub software, etc., to teach 'hands-on' experimental methods and computational tools for spatial omics and data analysis

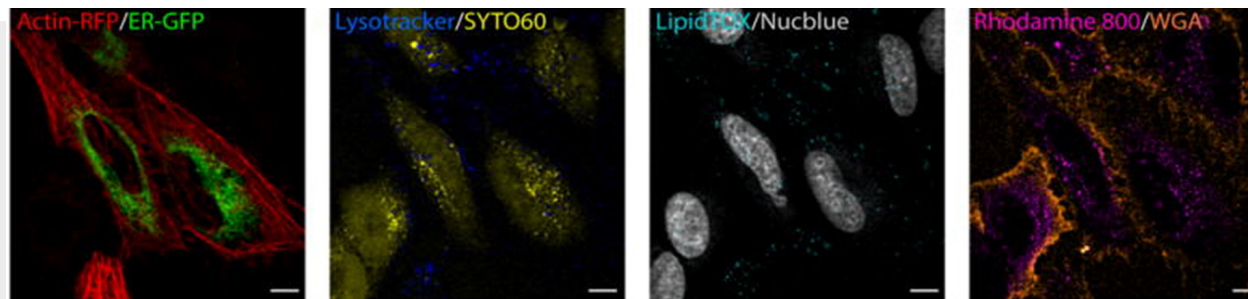
Multiplex antibody Immunofluorescence (example here is CODEX)

Black, et al. (Nolan group) Nat Protocols (2021), Gotsev,... Nolan, Cell (2018)



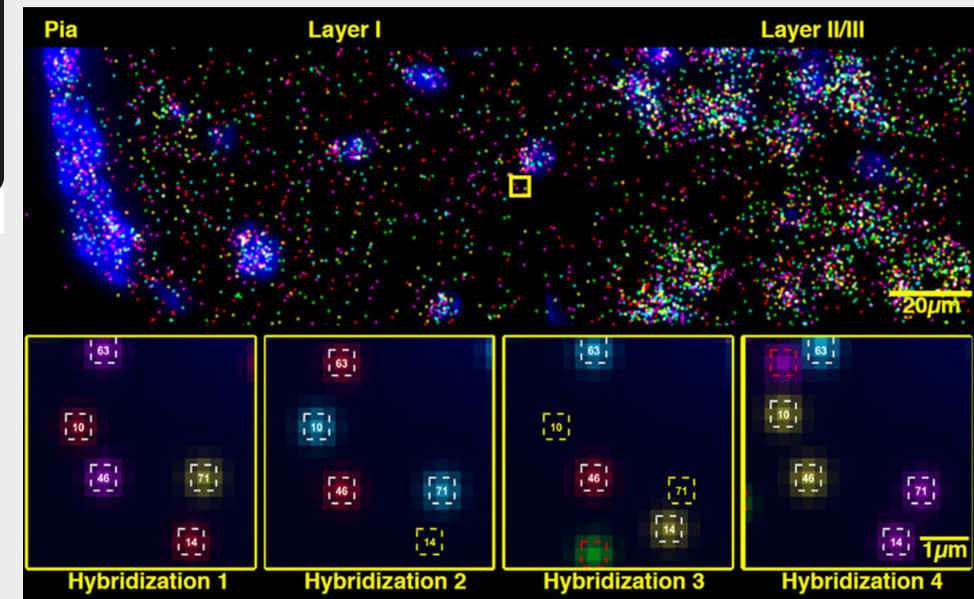
Single cell in situ

highly multiplex proteomics and transcriptomics and metabolomics and epigenomics tools are 5-10 years old.



Super Multiplex vibrational imaging in in single cells—L. Wei,... W. Min Nature 2017

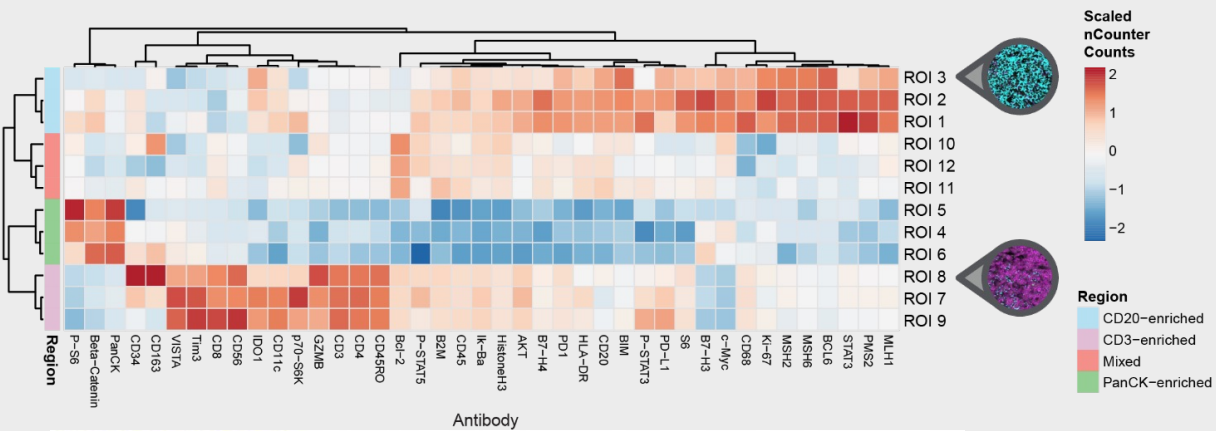
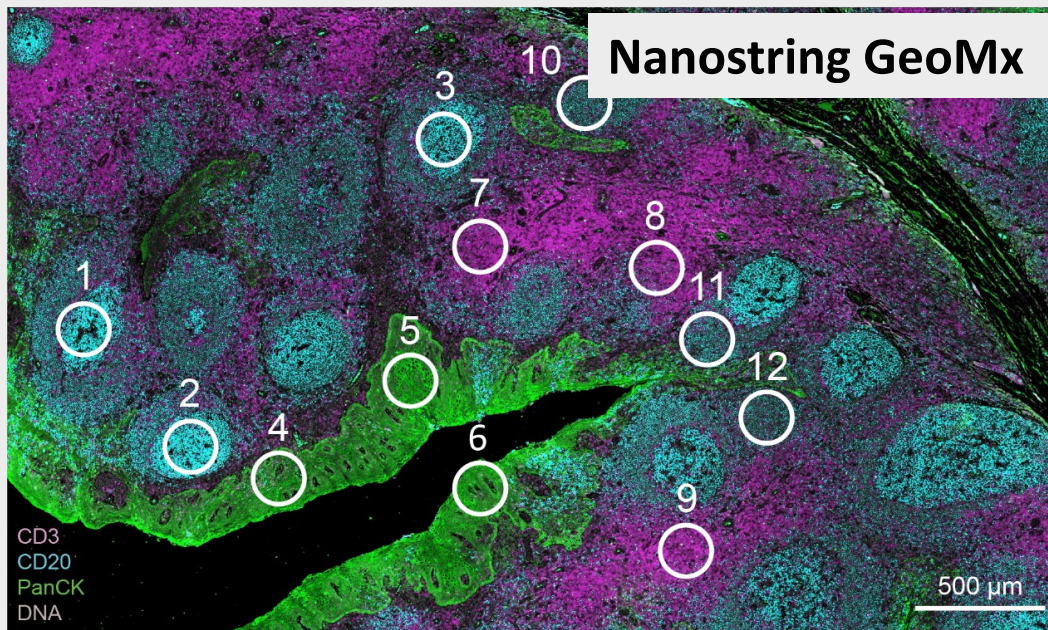
Spatial Transcriptomics –
Nature Method of the Year 2020



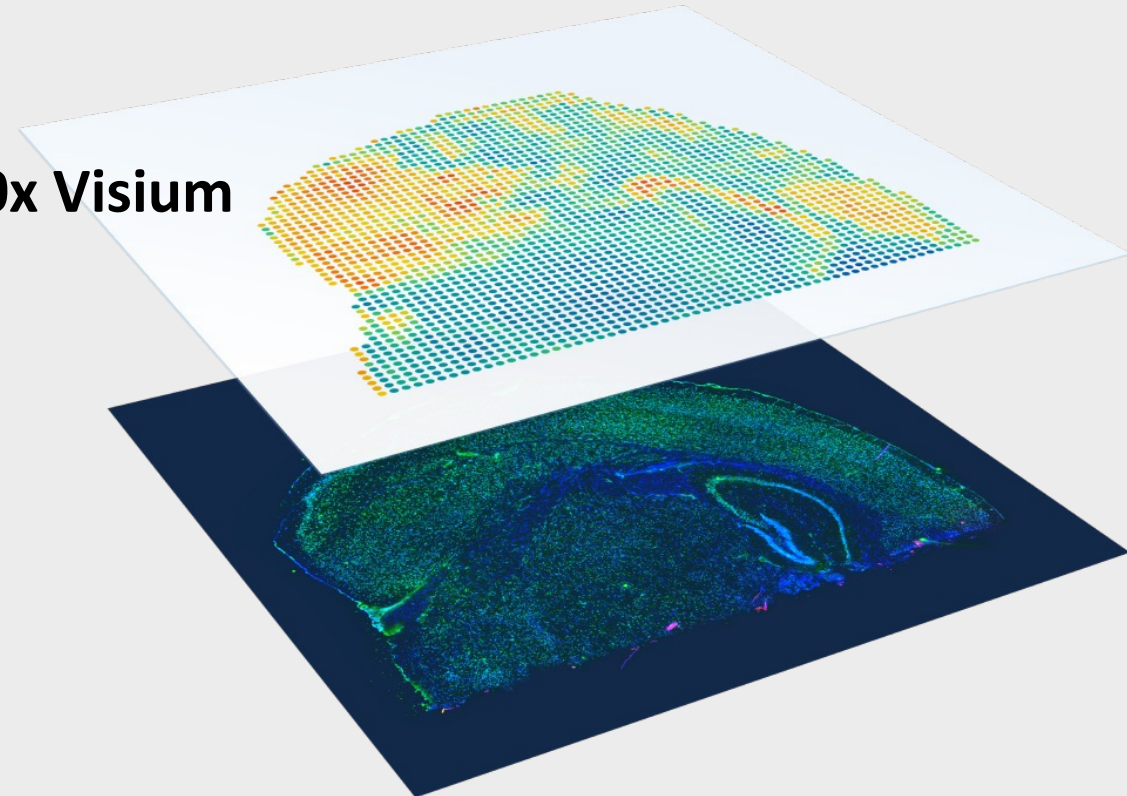
seqFISH+ images 10^4 mRNAs in single cells—with high accuracy and sub-diffraction-limit resolution

Long Cai group, Neuron 2016, Nature 2019

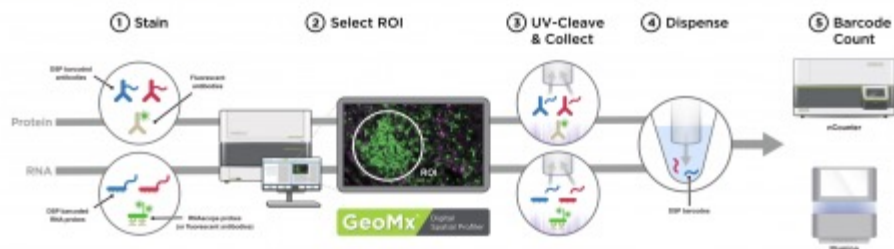
**Proteins and Transcripts at high
 (multi-cellular) resolution
 With targeted probes and next
 gen sequencing**

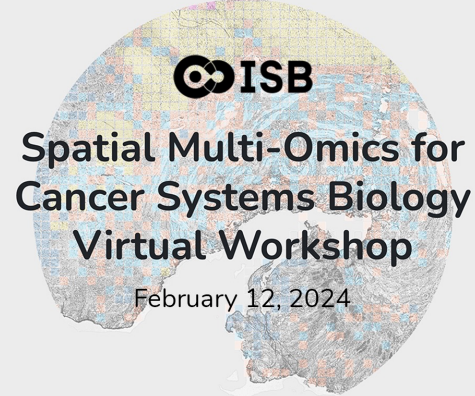
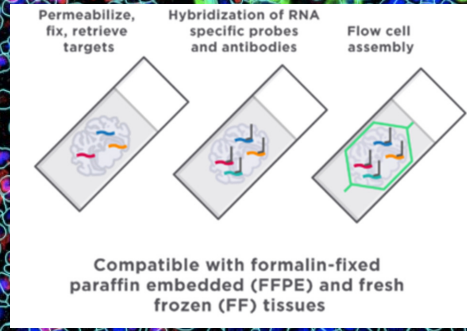
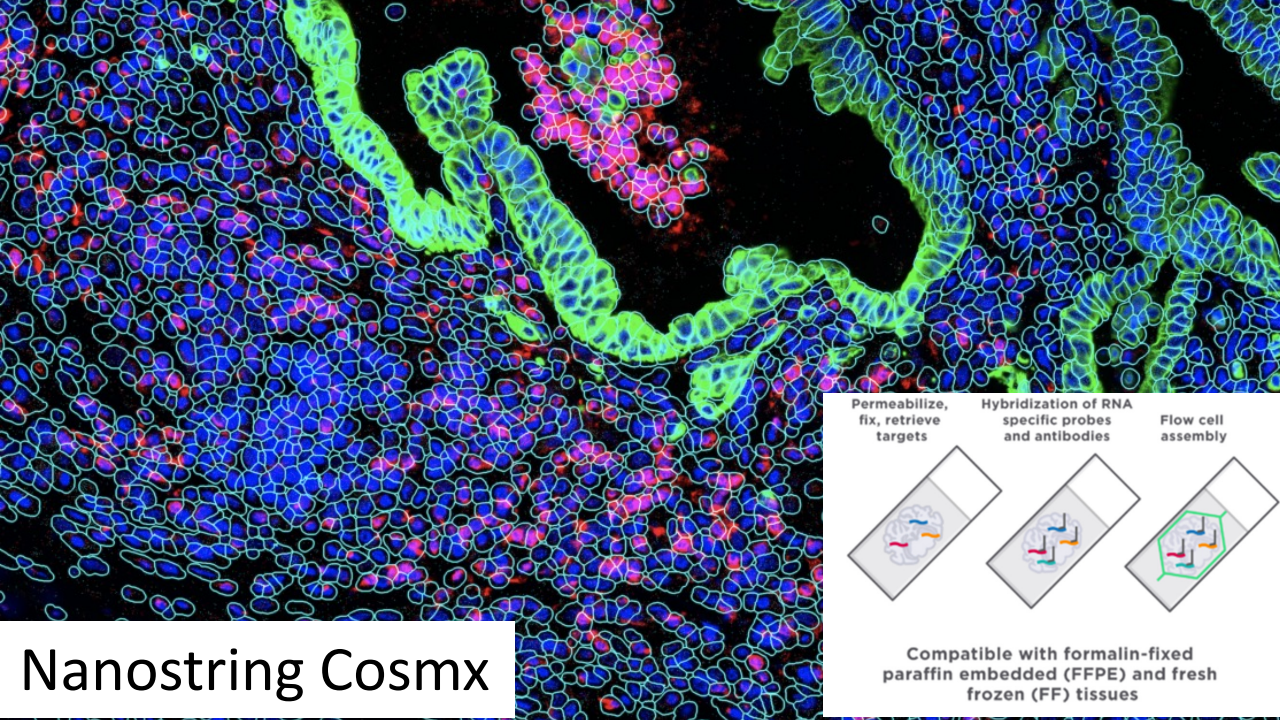


10x Visium



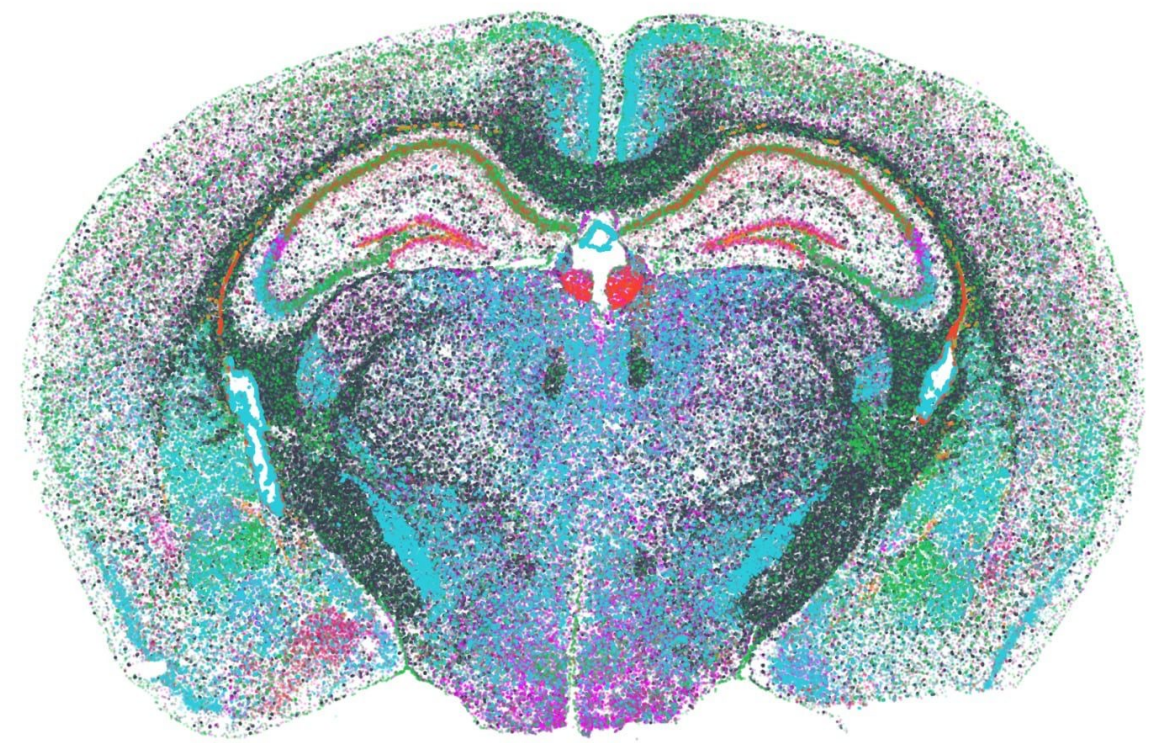
GeoMx DSP Workflow



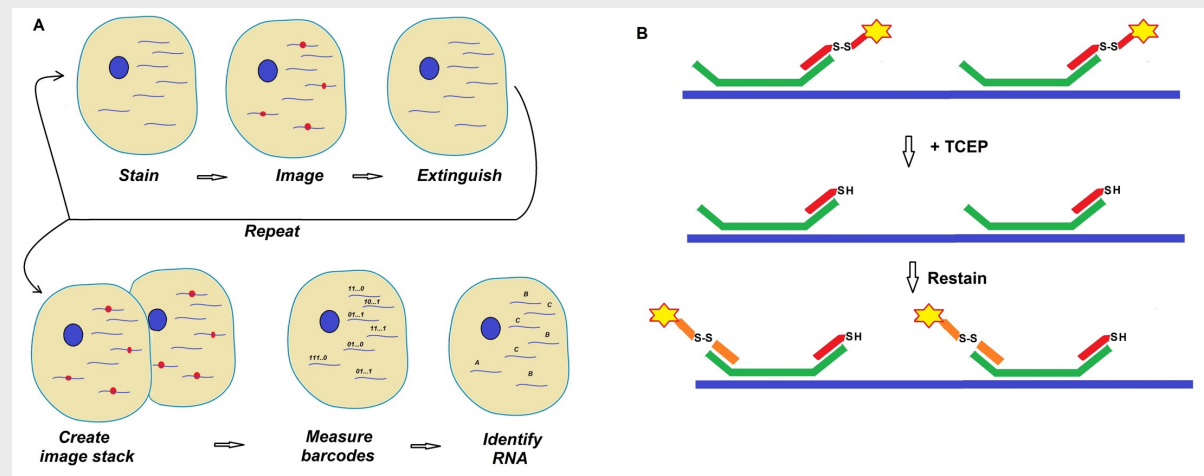


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Subcellular resolution from multiple toolsets



Vizgen MerFISH probes



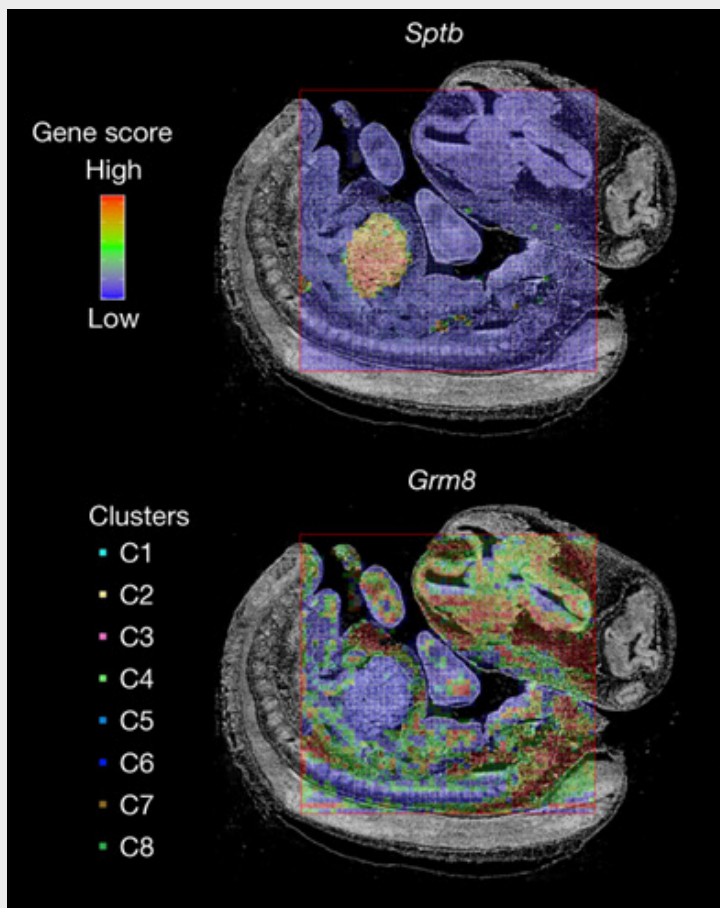
Epigenetics

Metabolomics

Spatial ATAC-seq & other Multiomics

(Dbit-seq methods) R. Fan Group, Yale
AtlasXomics, Inc.

Y. Deng, ... R. Fan. *Nature* 2022

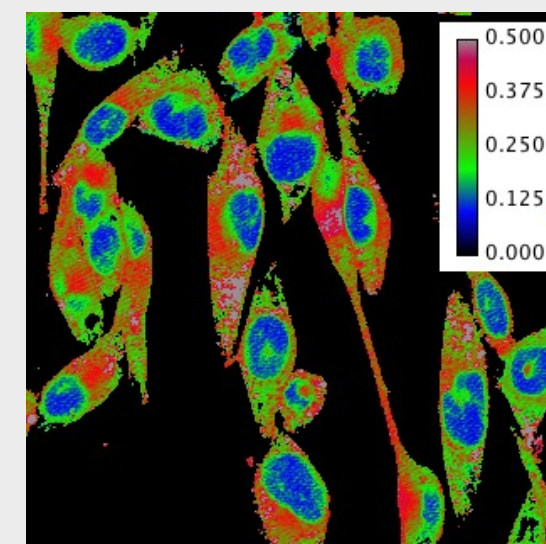
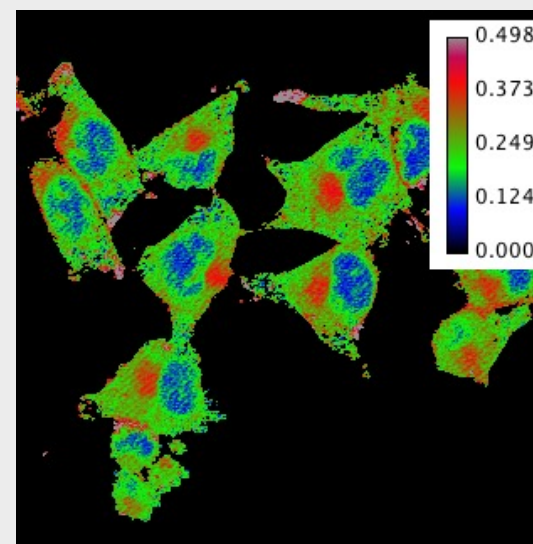
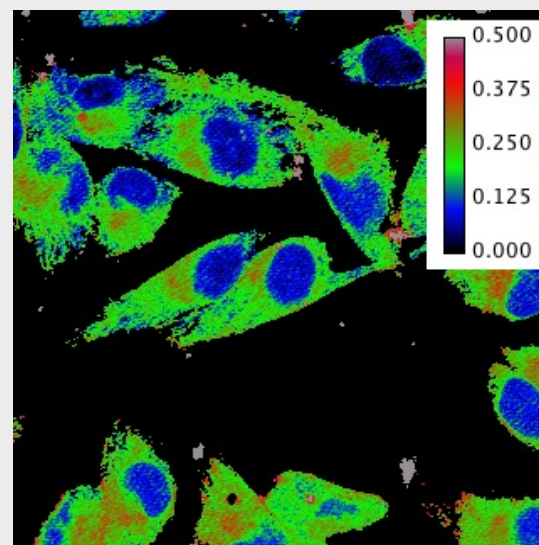


Raman (metabolic) Imaging: No labels, live cells, near video rate

M409 ■ Neural crest

M229 ■ plastic

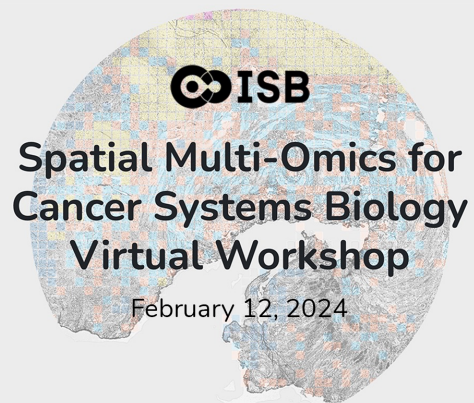
M262 ■ Melanocytic



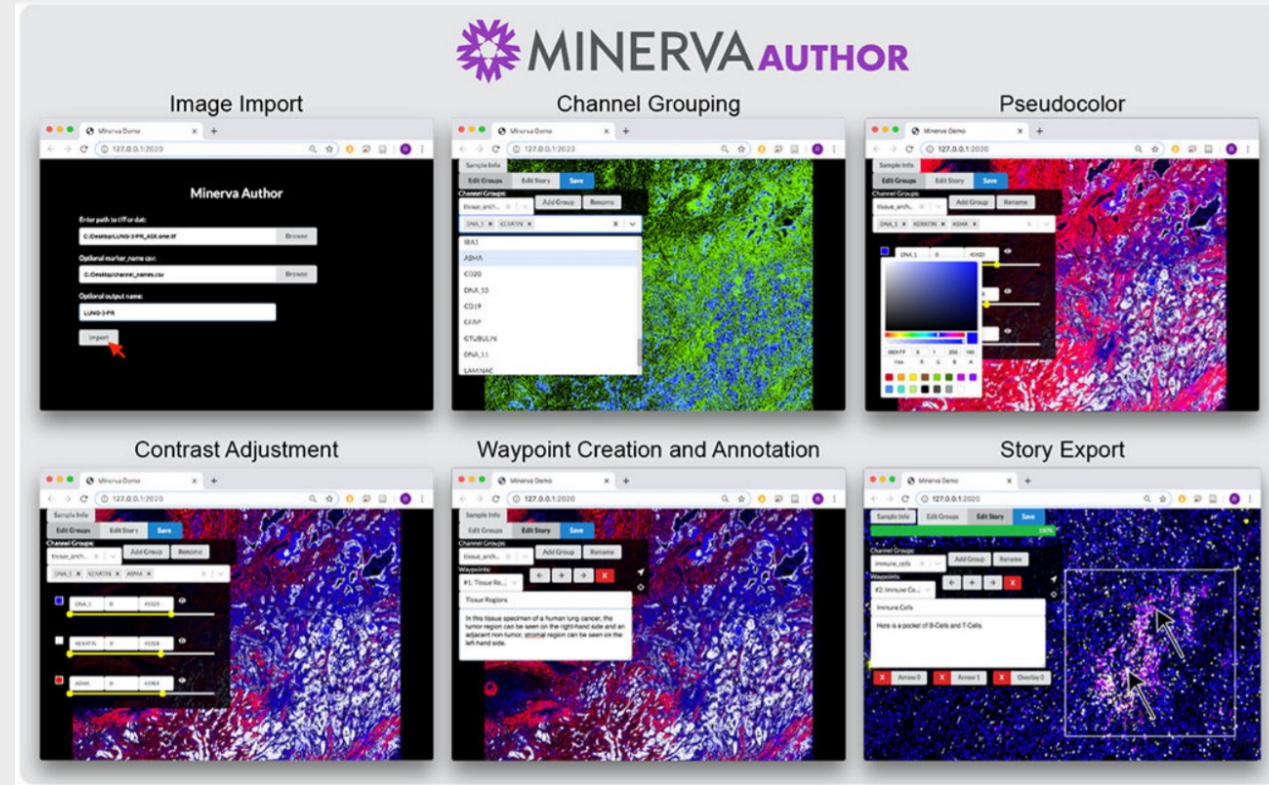
J. Du, .. L Wei *Nat. Comm* 2019



R. Dries Group, Gen. Bio (2021)



Sorger Group, J. Open Source Software (2020).

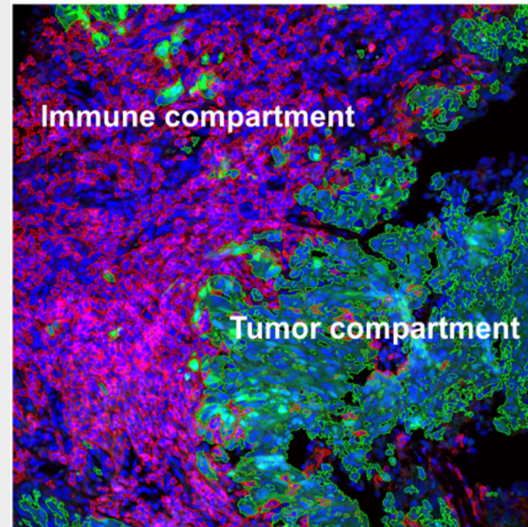


Why it's worth making computational methods easy to use. J. Fan Nat 2022

In any new field, confounding factors can limit the transportability of results from one platform to another, or across labs

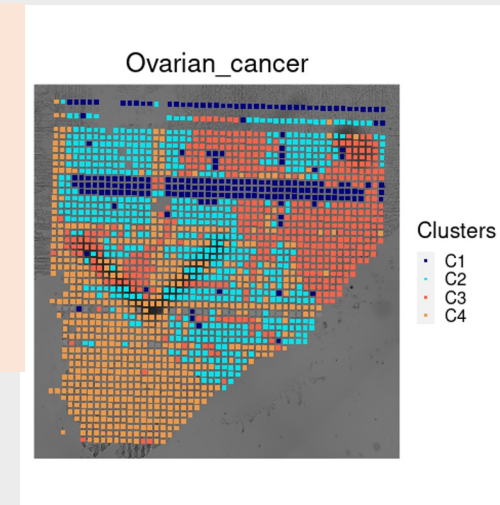
Tissue Types

- live cells
- FFPE
- fresh frozen
- size
- prep details



Data Quality

- spatial variations
- # of transcripts/pixel
- # fragments/pixel
- alignment with ground truth



Ground Truth References

- nuclear staining?
- H&E images?
- Immunofluorescence?
- Algorithms?

Data Integration

- tissue specific cell atlases
- integration across omics
- lab-to-lab variations and standards
- benchmarking algorithms