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Spatial Omics Visualizations: Lessons Learned from Networks and Maps



visualization
design lab



EVERYBODY IS TALKING ABOUT SPATIAL OMICS!

FOCUS | TECHNOLOGY FEATURE

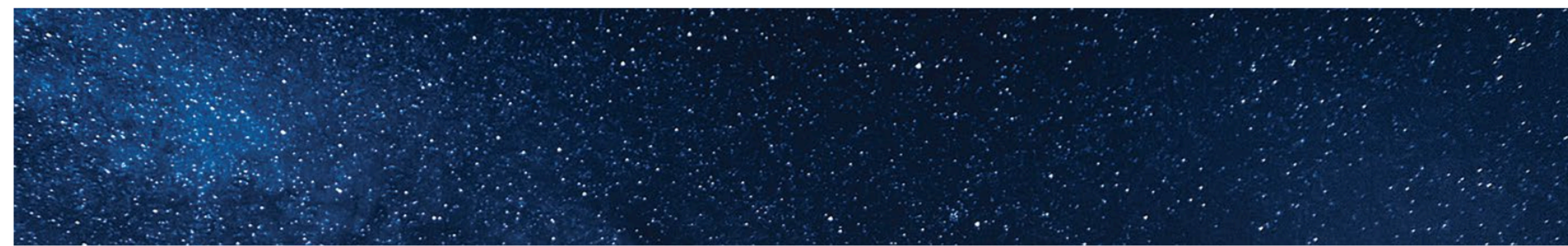


Method of the Year: spatially resolved transcriptomics

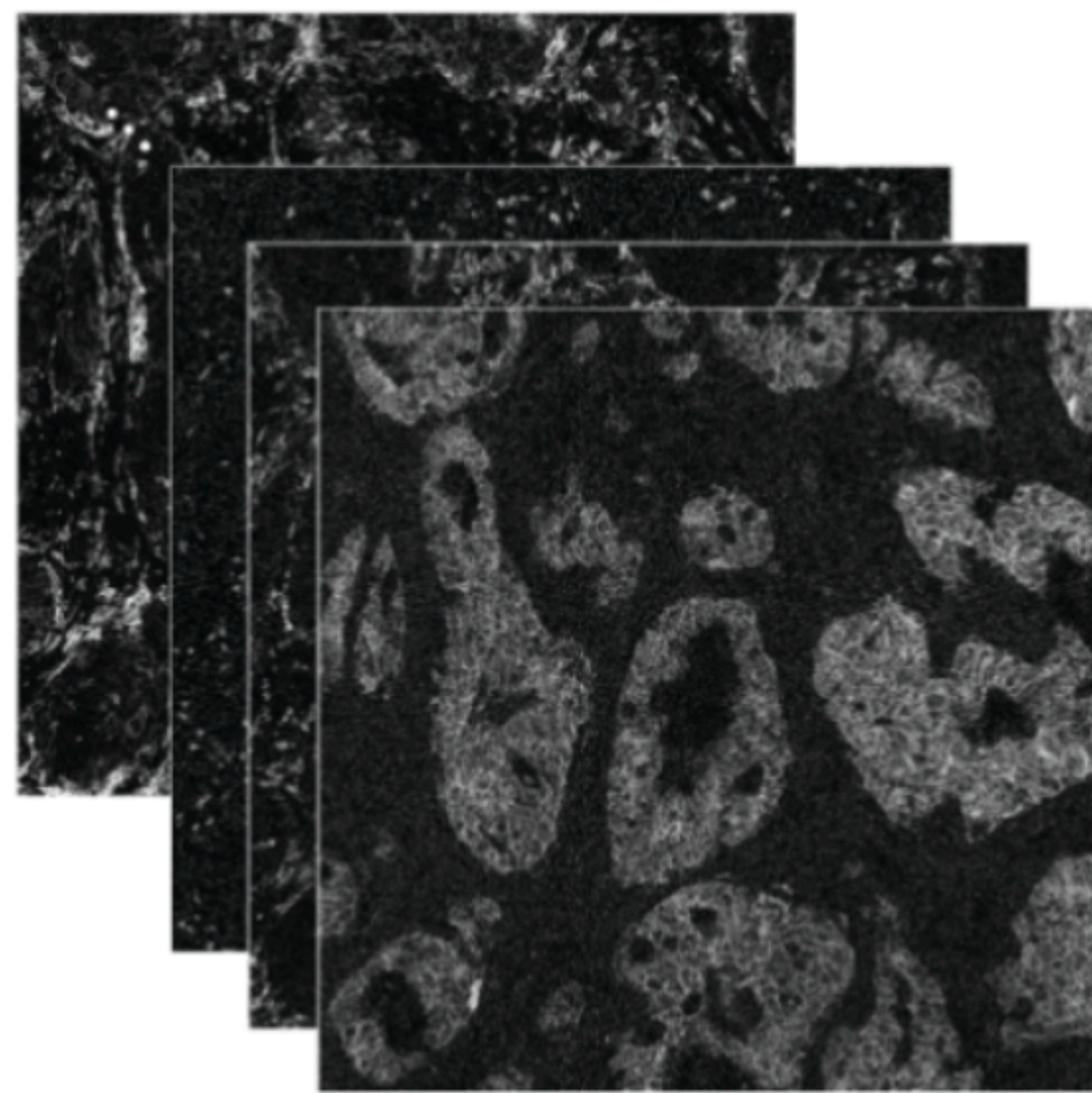
Nature Methods has crowned spatially resolved transcriptomics Method of the Year 2020.

Vivien Marx

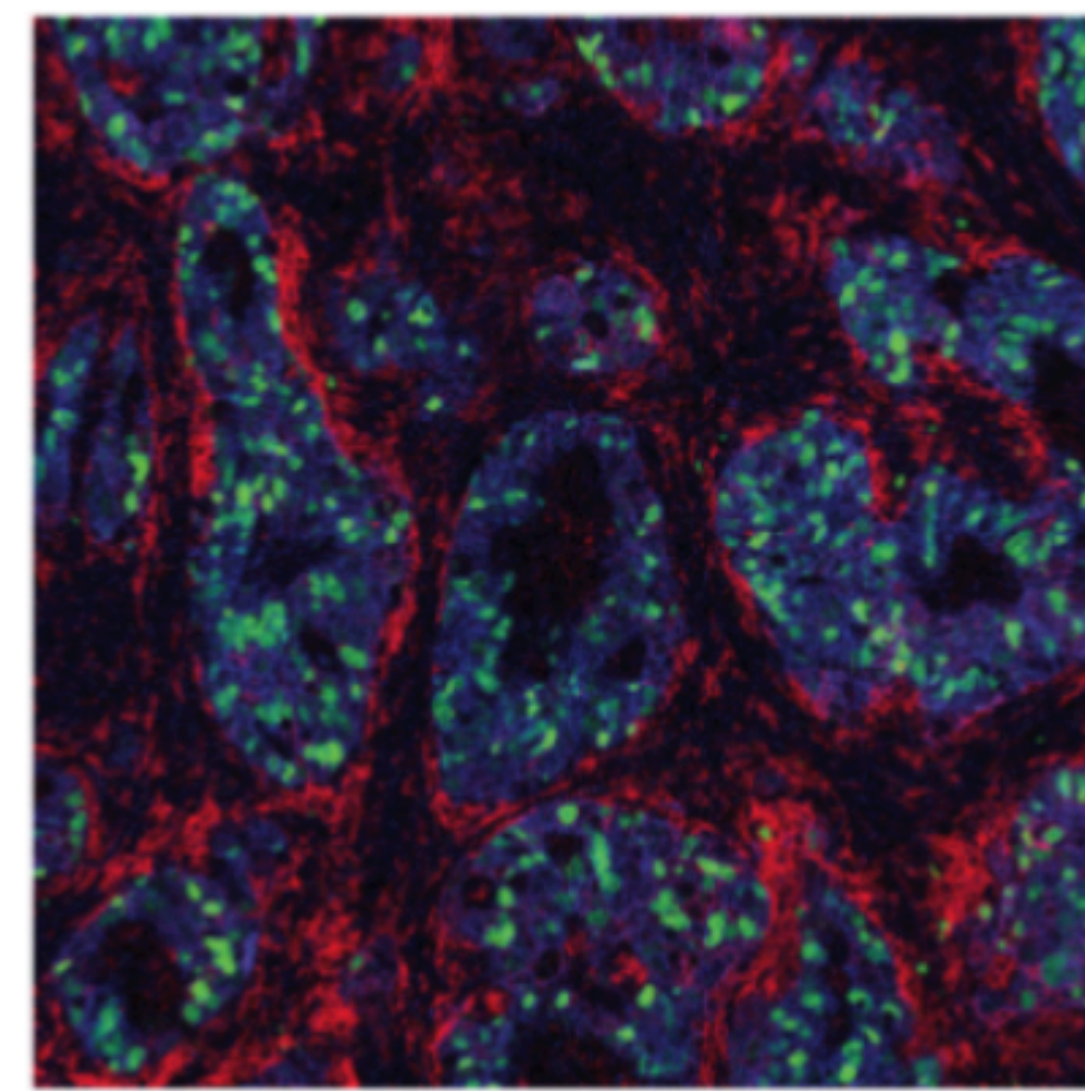
If a researcher is making a smoothie, it might be snack time. Or it could be the moment to prepare a sample for bulk RNA sequencing, in which tissue is homogenized and analyzed to yield



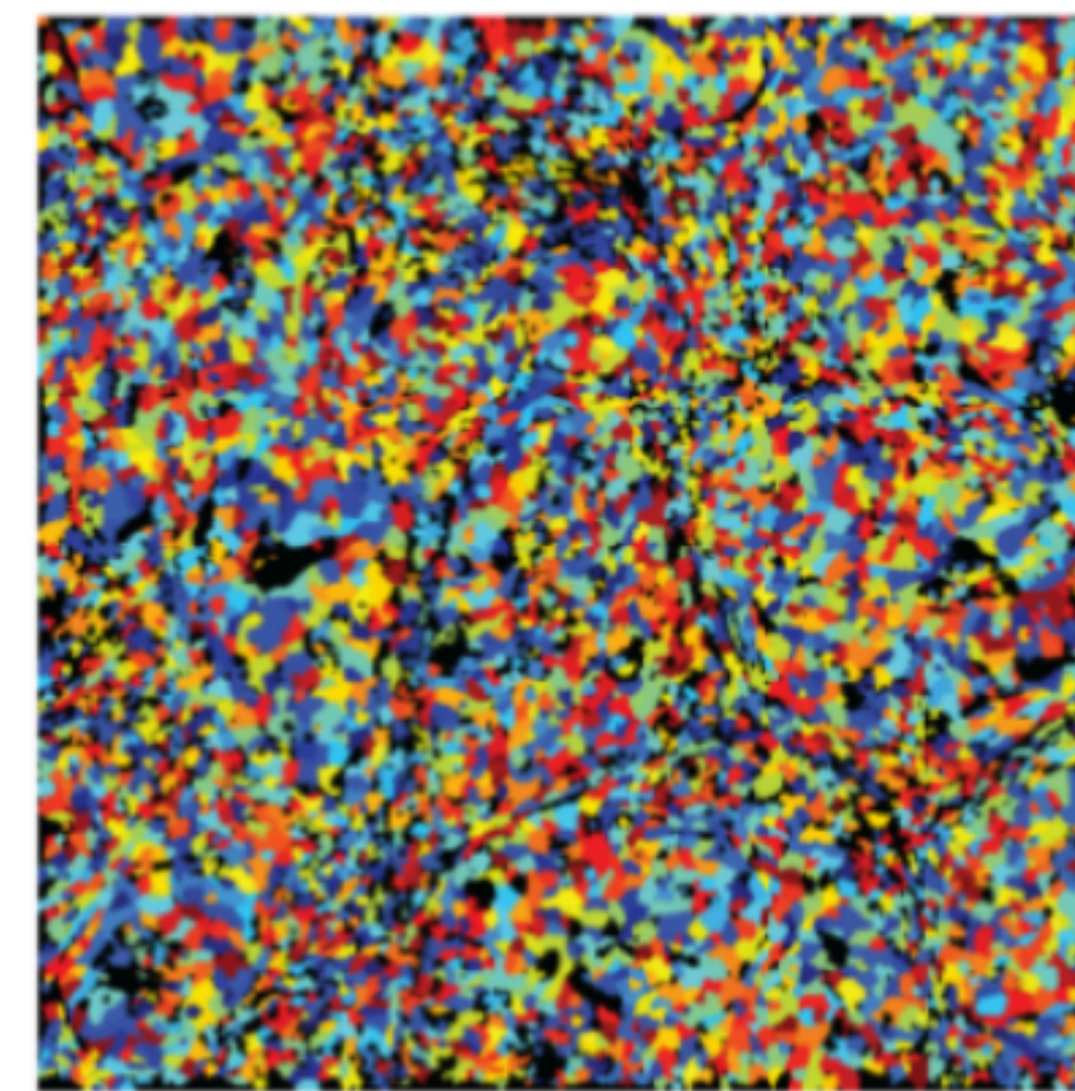
BIOVIS CHALLENGE: FROM IMAGES TO ANALYSIS



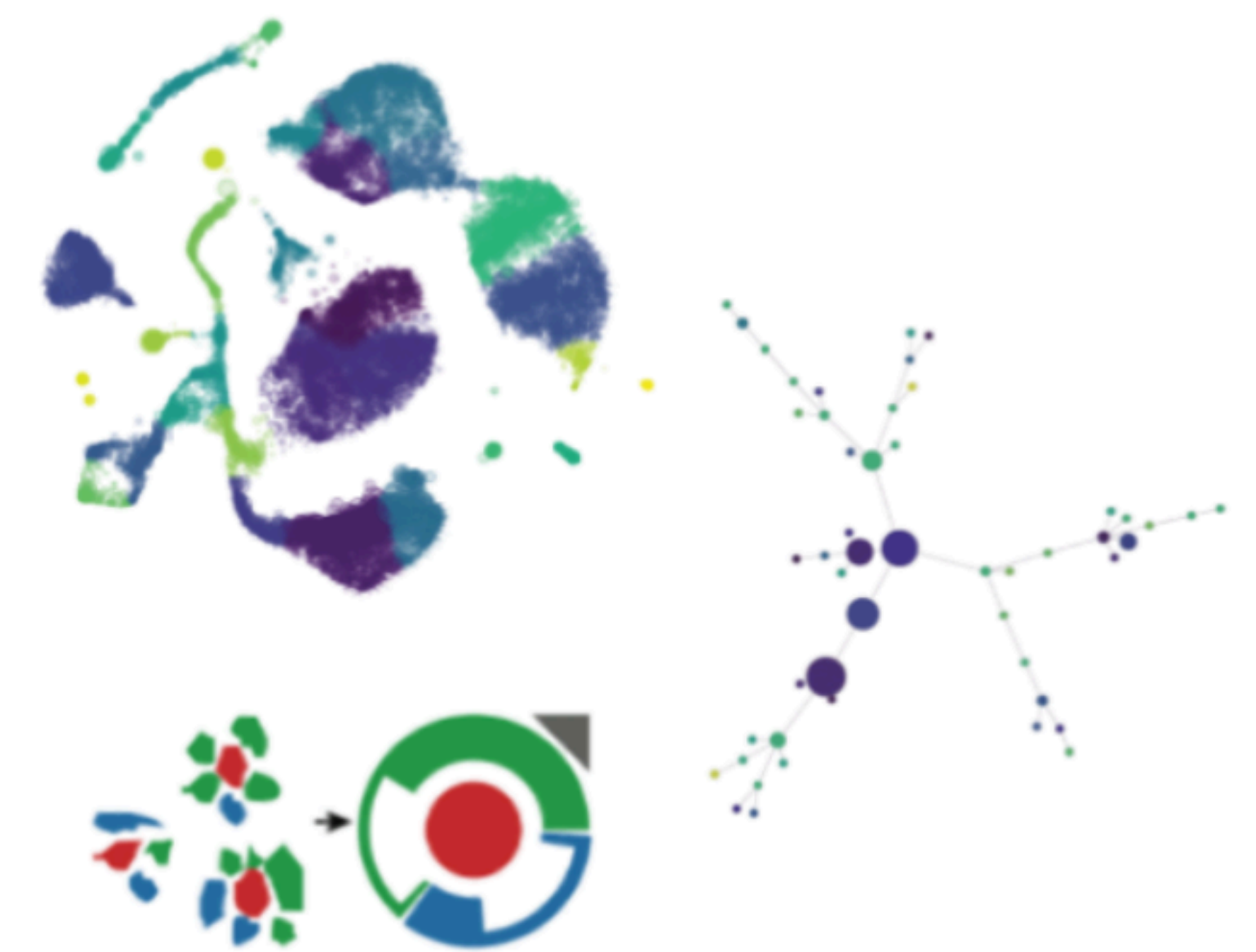
Stack of images, each corresponding to one protein, i.e., one channel of the multi-dimensional measurement



Basic visualization e.g., overlaying 3 proteins as RGB



Segmentation mask.
Cells indicated by random color



Downstream data analysis

SPATIAL OMICS VISUALIZATION CHALLENGES

High dimensional data

Similar to “classical” omics data (except for scale)

Spatial location / proximity is important

NOT encountered in “classical” omics data.

ANALOGY: MAPS!

Fixed location

Potentially high-D

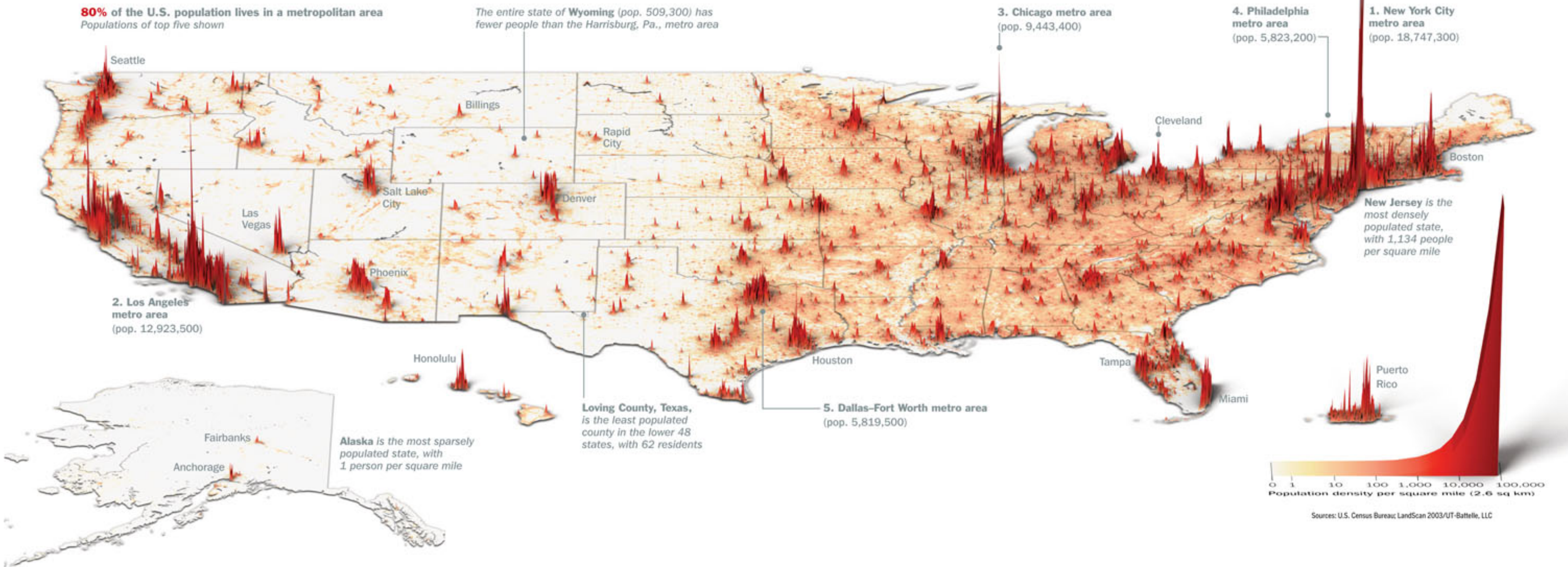
vector for each location

Where We Live...

Unlike many developed countries, the U.S. keeps growing. We are also moving south and west. But compared with China or India, the nation is a vast prairie

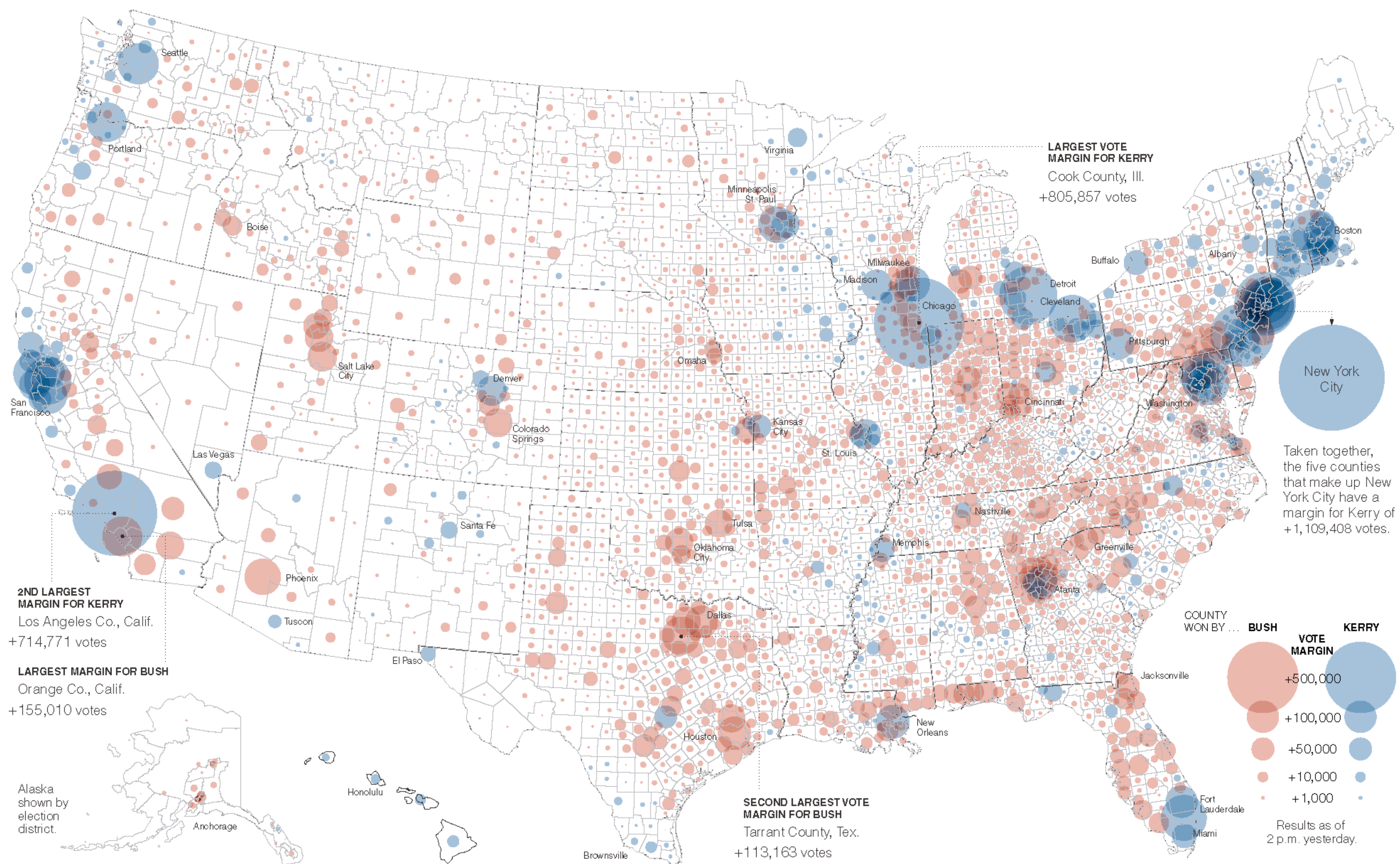
Our families are getting smaller—with one vital exception. Compared with those of Europe and Japan, the U.S. population is younger and more colorful because of the continued arrival of immigrants and their higher-than-average birthrates. Of the 100 million Americans who will join us in the next 37 years, half will be immigrants or their children. In the next few decades, 97% of the world's population growth will occur in the developing world; the U.S. is the largest developed country in the world that is still growing at a healthy clip. That matters, strategically, economical-

Ala.; Possum Trot, Ky.; or Lonelyville, N.Y. But they are all probably close to someone's idea of paradise. —By Nancy Gibbs



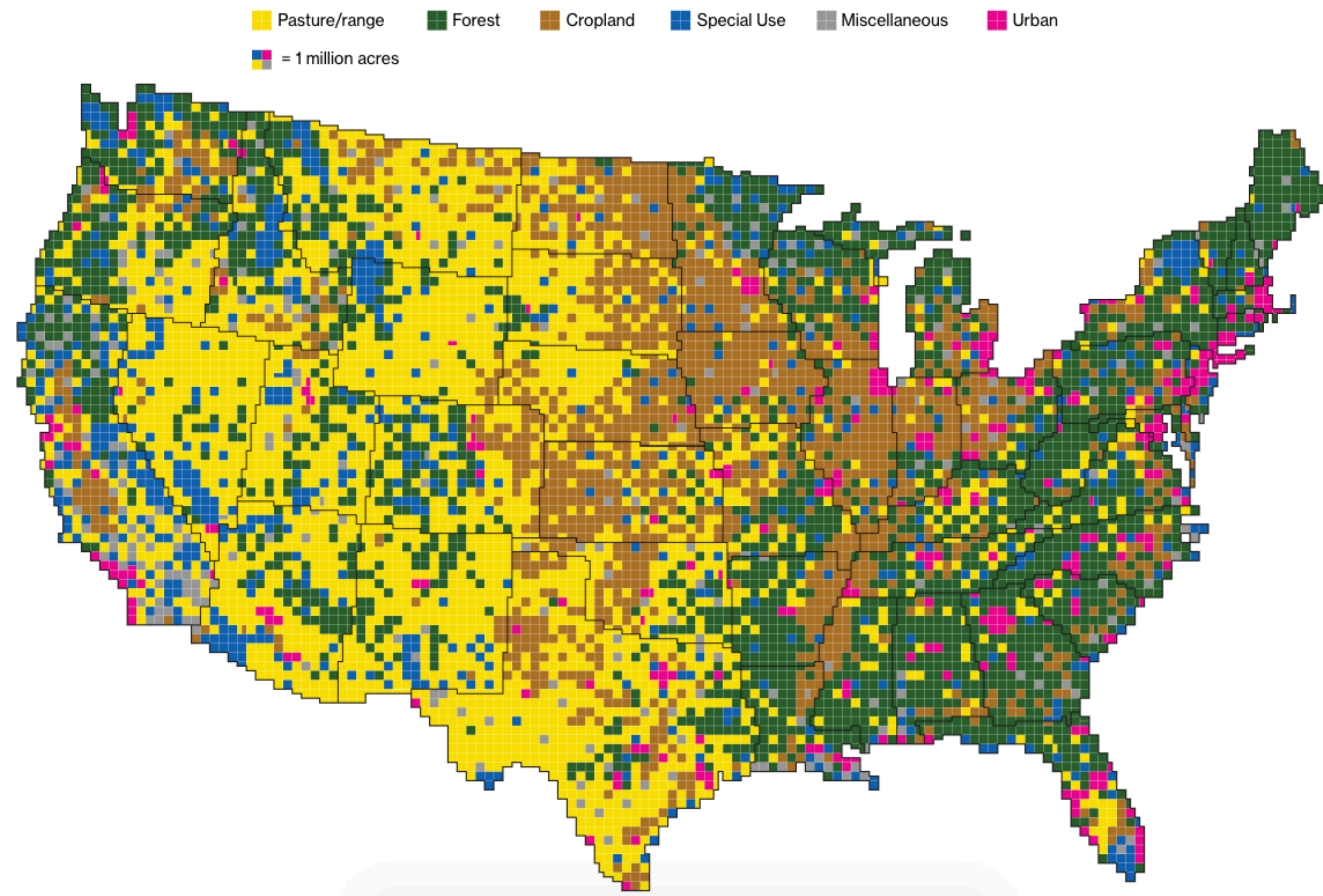
SINGLE DATA VALUE + SPATIAL LOCATION

Glyphs - Size



Matthew Ericson, NY Times

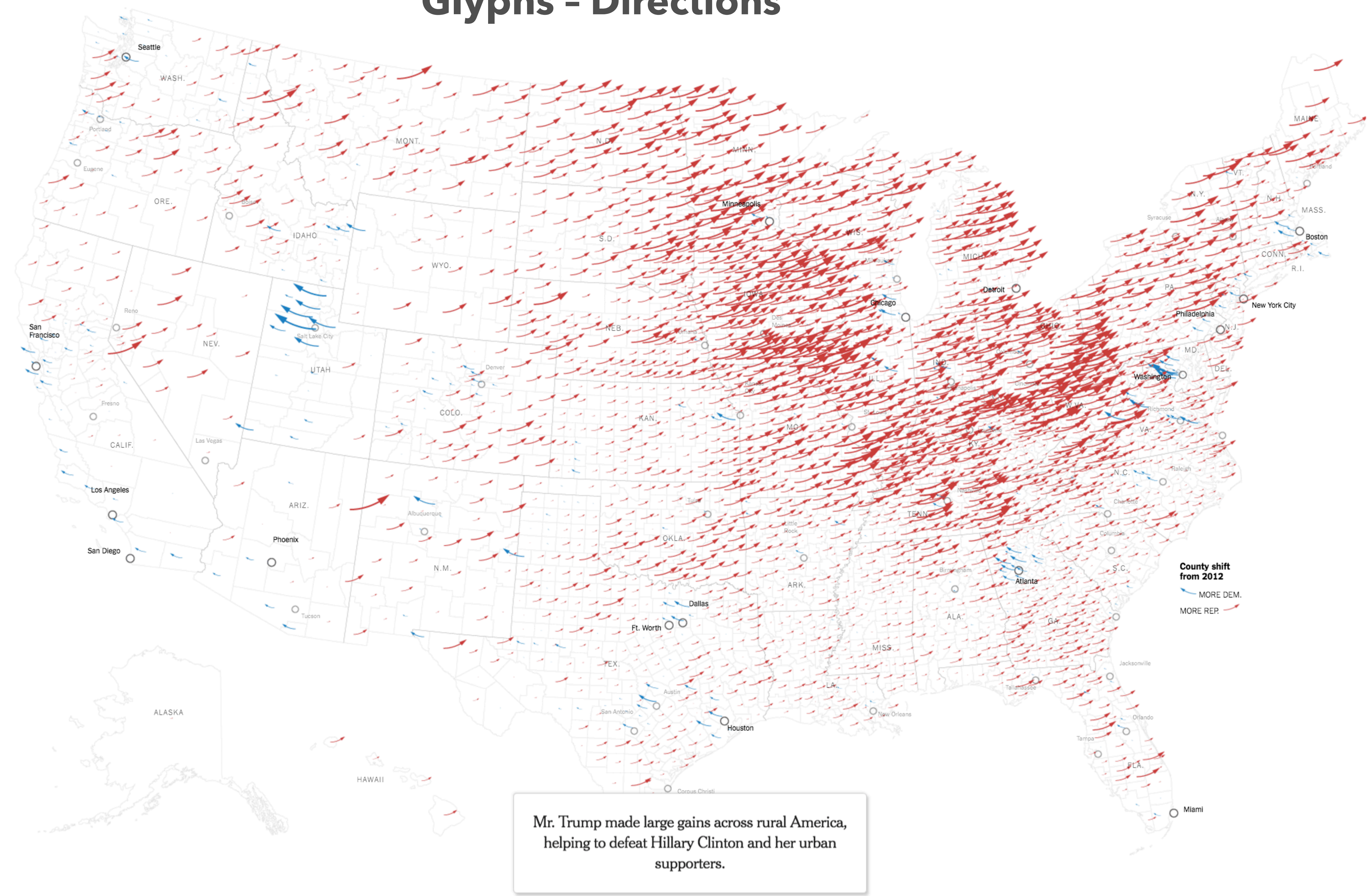
Color



Blomberg

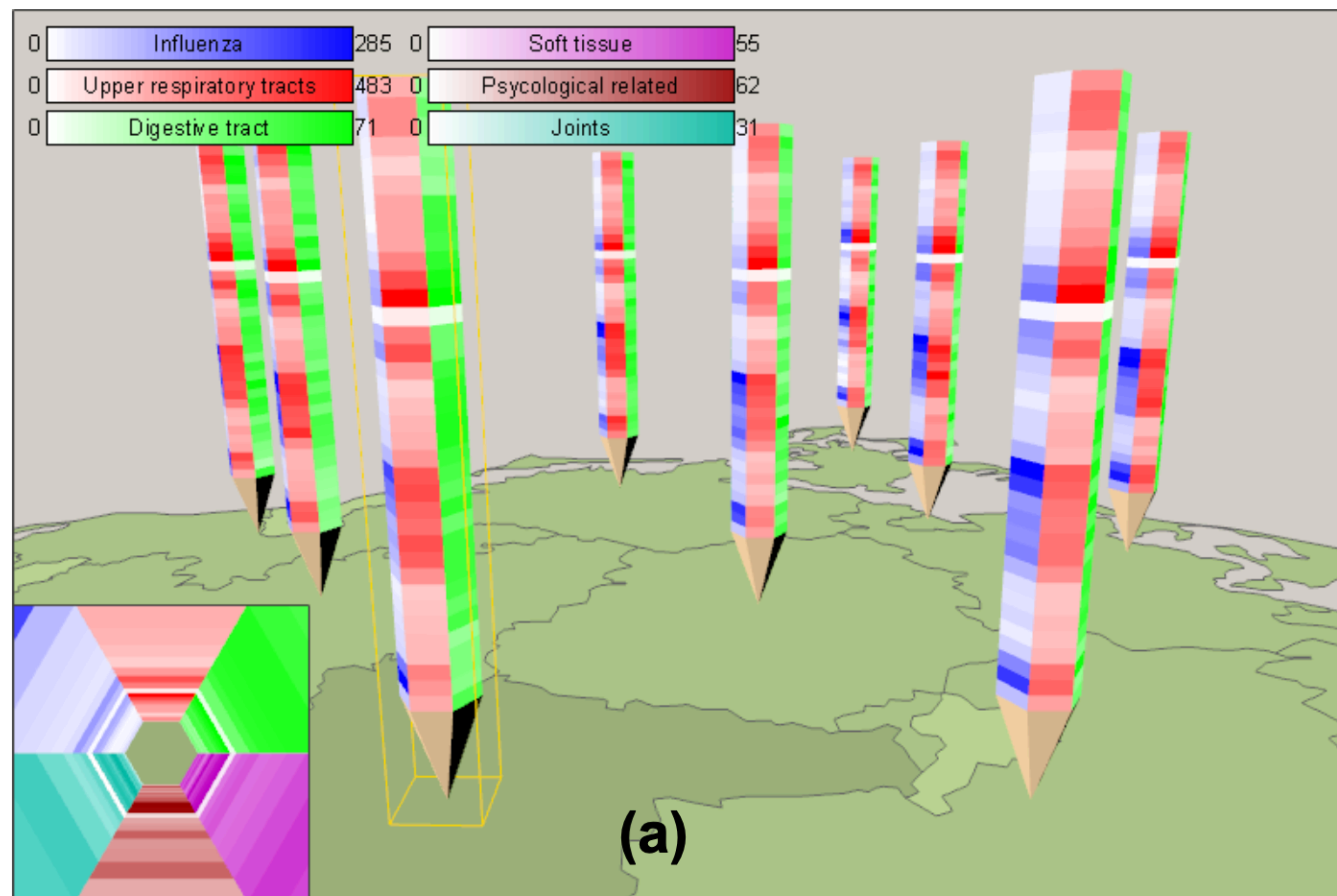
SINGLE DATA VALUE + SPATIAL LOCATION

Glyphs - Directions



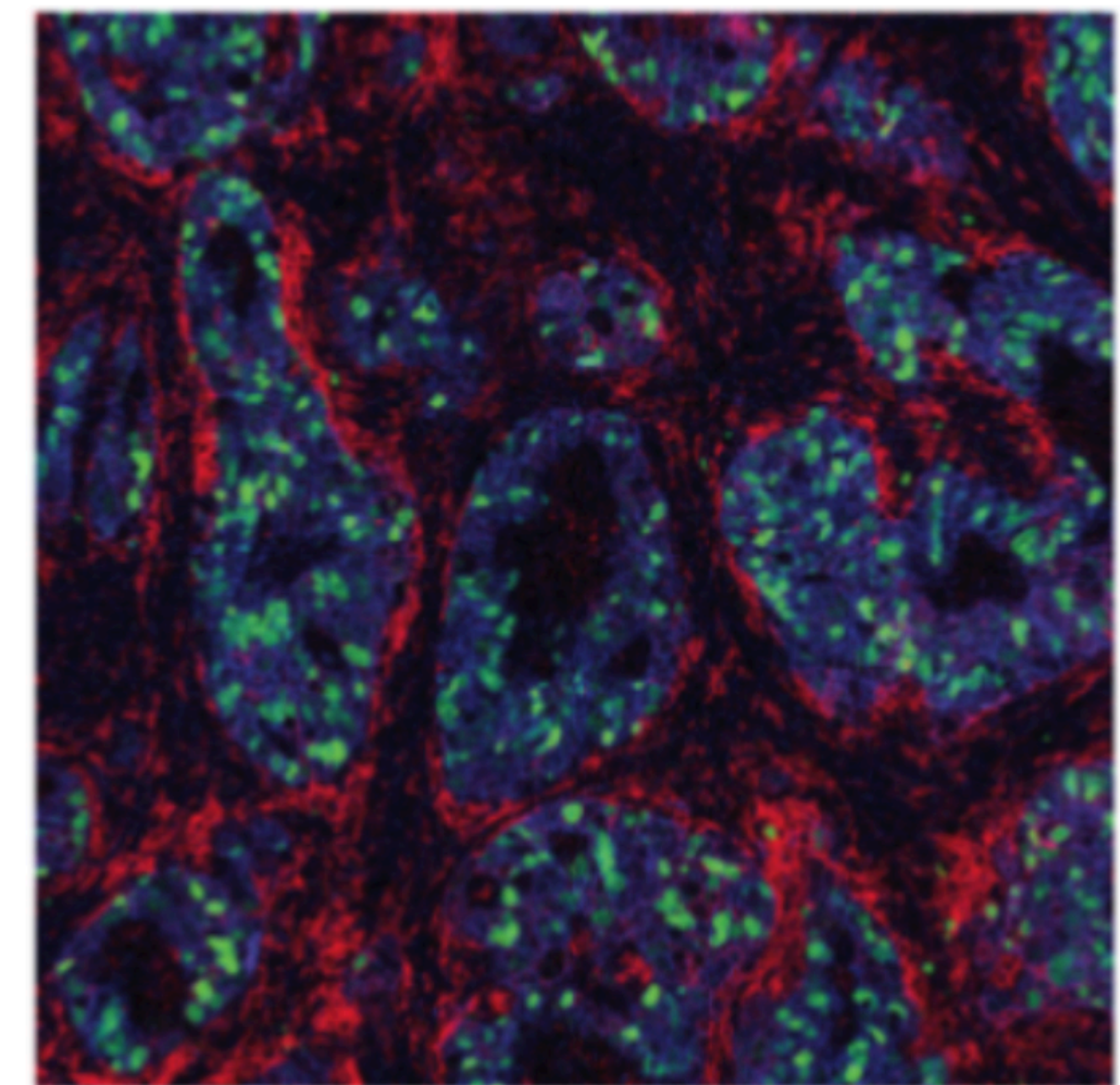
BUT WHAT IF YOU HAVE MORE THAN ONE DATA VALUE?

Glyphs?



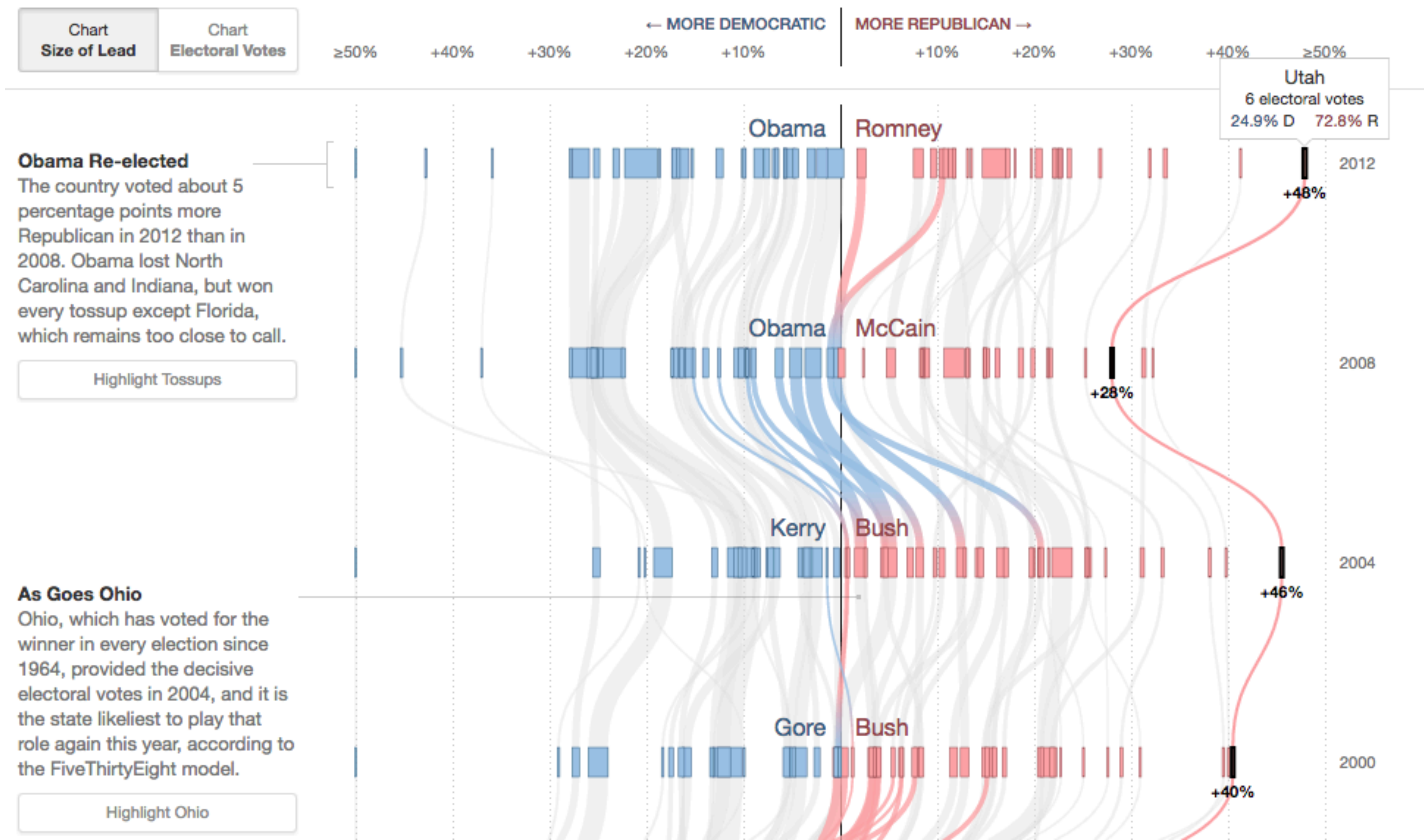
Scalability? [Tominski]

Don't treat color channels as separate visual channels



Basic visualization e.g.,
overlaying 3 proteins as RGB

DO WE REALLY NEED A MAP?

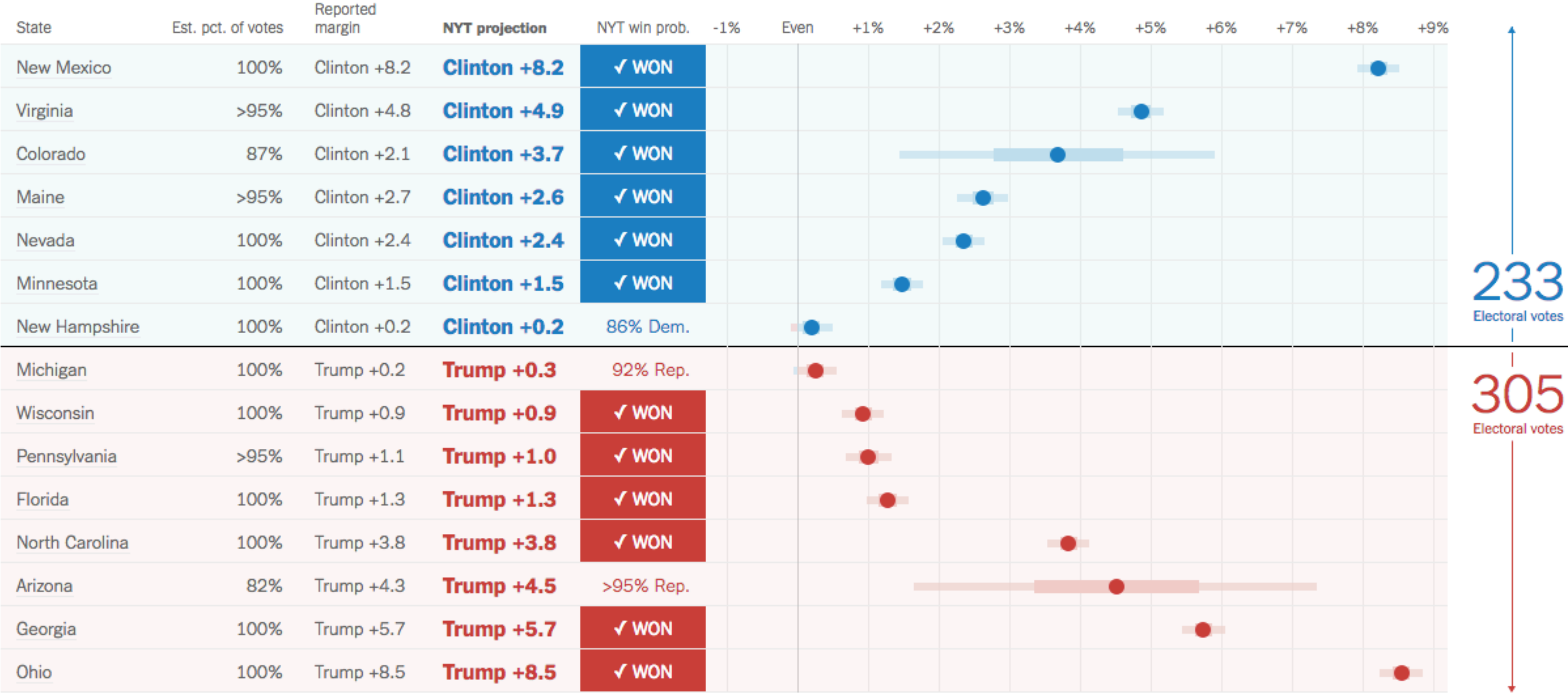


DO WE REALLY NEED A MAP?

It's hard to do more complex things with maps

Is the spatial context paramount?

Is the spatial context a proxy for something?

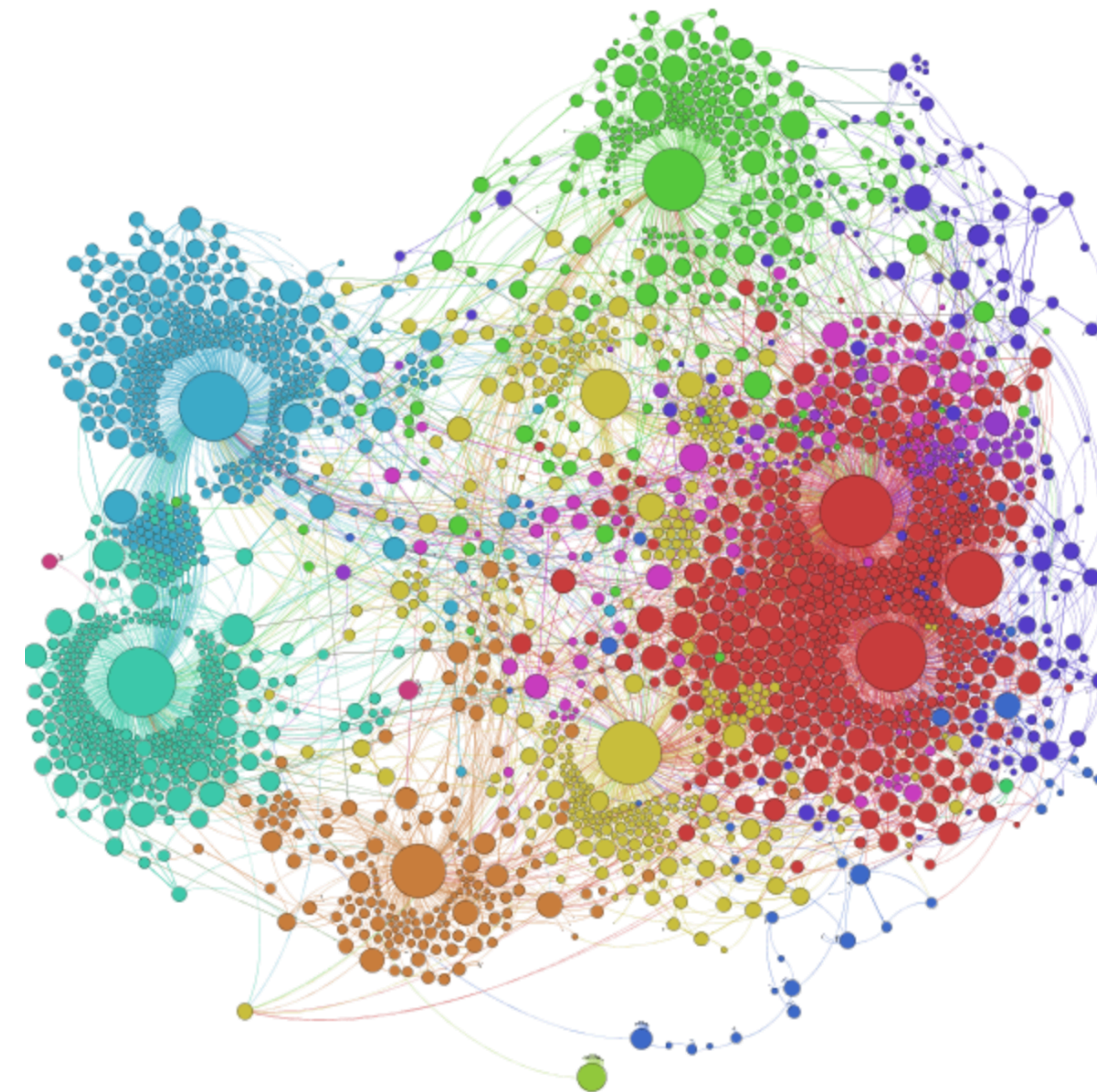


ANALOGY: MULTIVARIATE NETWORKS

Lots of attributes for nodes and edges

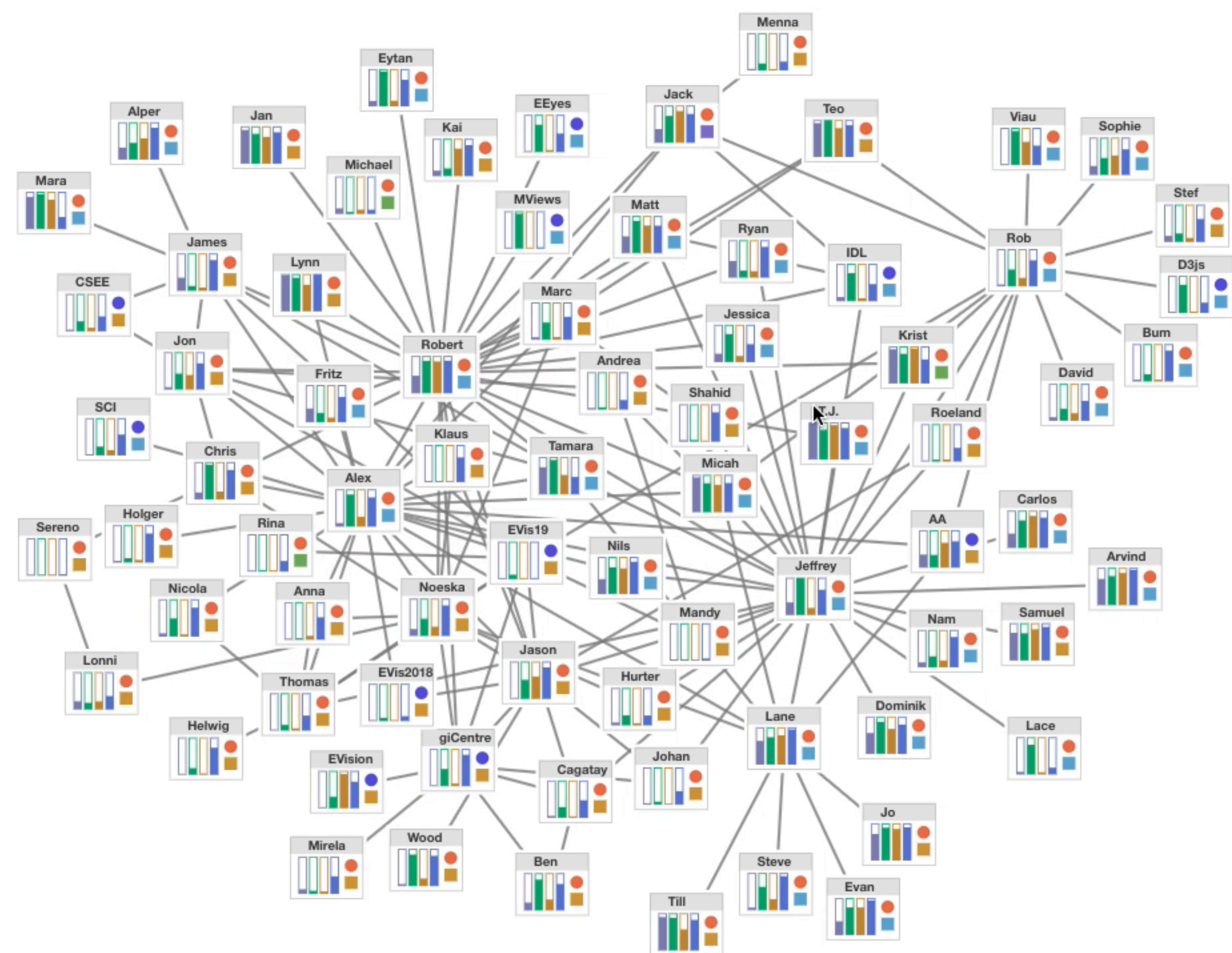
**Location doesn't matter, but
connectivity does**

Can't choose location freely



ON-NODE ENCODING?

Maybe if you zoom in
Still limited



The State of the Art in Visualizing Multivariate Networks

C. Nobre¹, M. Meyer¹, M. Streit², and A. Lex¹

¹University of Utah, Utah, USA
²Johannes Kepler University Linz, Austria

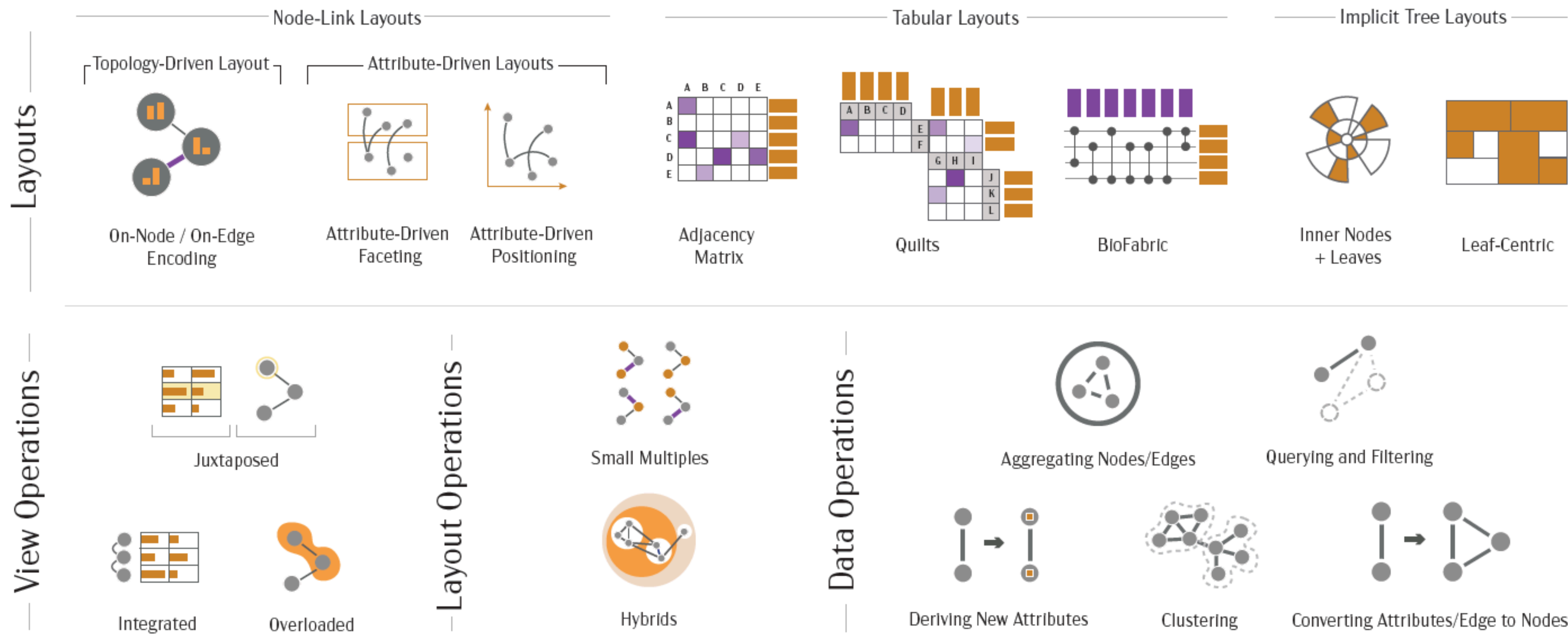


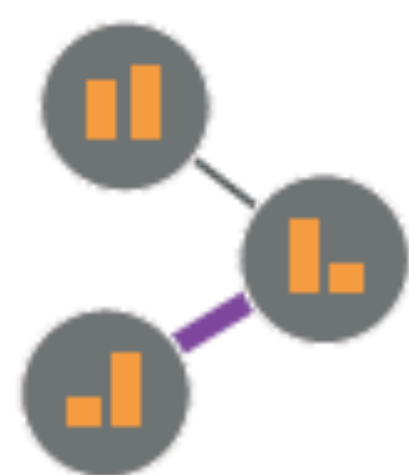
Figure 1: A typology of operations and layouts used in multivariate network visualization. *Layouts* describe the fundamental choices for encoding multivariate networks. *View Operations* capture how topology and attribute focused visualizations can be combined. *Layout Operations* are applied to basic layouts to create specific visualization techniques. *Data Operations* are used to transform a network or derive attributes before visualizations. The colors reflect node attributes (orange), edge attributes (purple), and topology (grey).

Abstract
Multivariate networks are made up of nodes and their relationships (links), but also data about those nodes and links as attributes. Most real-world networks are associated with several attributes, and many analysis tasks depend on analyzing both, relationships and attributes. Visualization of multivariate networks, however, is challenging, especially when both the topology of the network and the attributes need to be considered concurrently. In this state-of-the-art report, we analyze current practices and classify techniques along four axes: layouts, view operations, layout operations, and data operations. We also provide an analysis of tasks specific to multivariate networks and give recommendations for which technique to use in which scenario. Finally, we survey application areas and evaluation methodologies.

Layouts

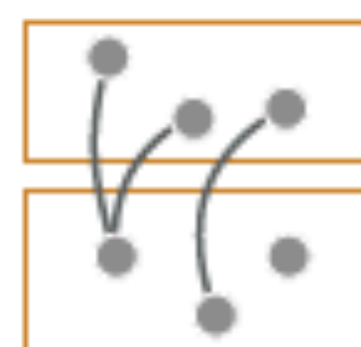
Node-Link Layouts

Topology-Driven Layout

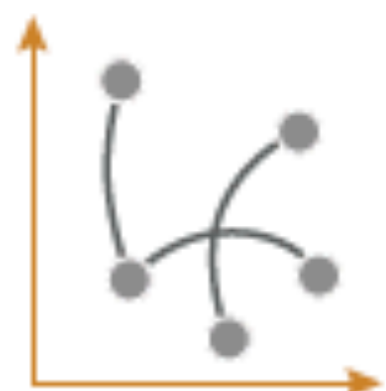


On-Node / On-Edge
Encoding

Attribute-Driven Layouts



Attribute-Driven
Faceting

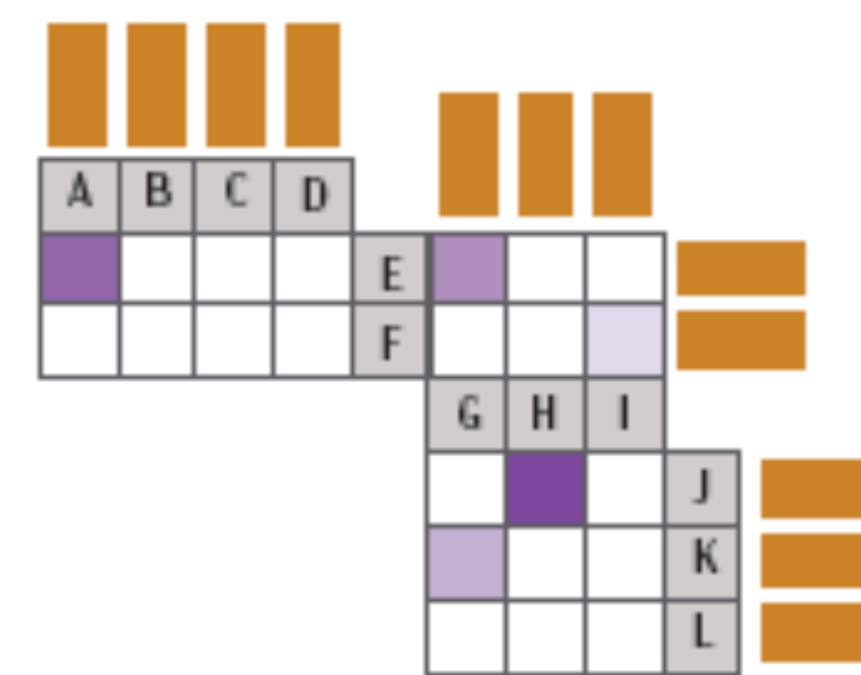


Attribute-Driven
Positioning

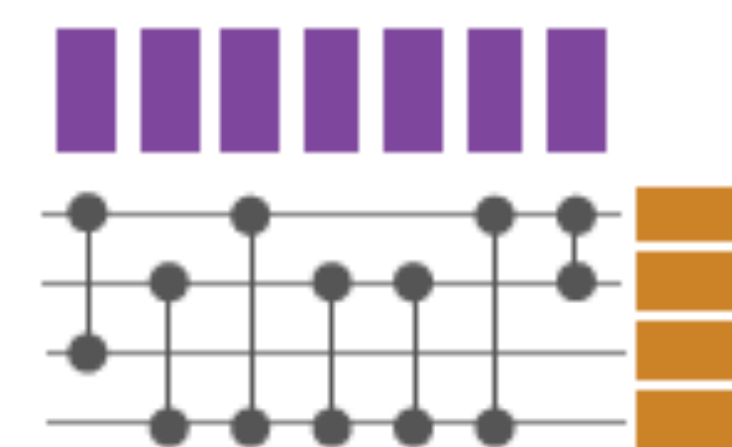


Adjacency
Matrix

Tabular Layouts



Quilts

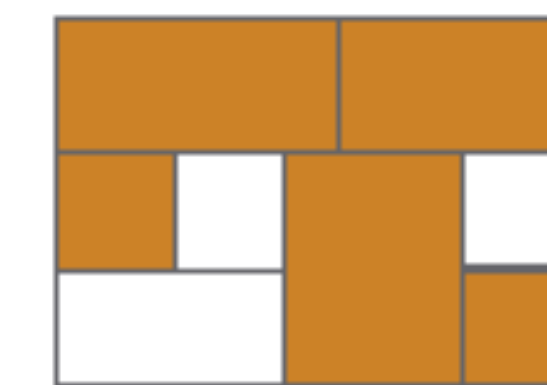


BioFabric

Implicit Tree Layouts



Inner Nodes
+ Leaves



Leaf-Centric

View Operations

Juxtaposed



Integrated

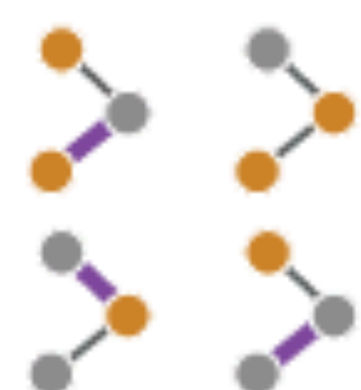


Overloaded



Layout Operations

Small Multiples



Hybrids



Data Operations

Aggregating Nodes/Edges



Deriving New Attributes



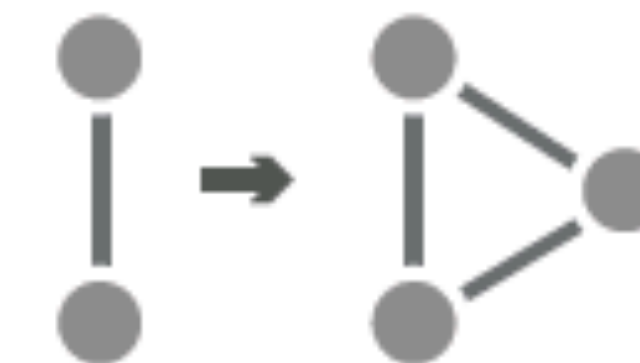
Clustering



Querying and Filtering



Converting Attributes/Edge to Nodes



**SO WHAT ELSE CAN
YOU DO?**

INTERACTION!

Two Paths:

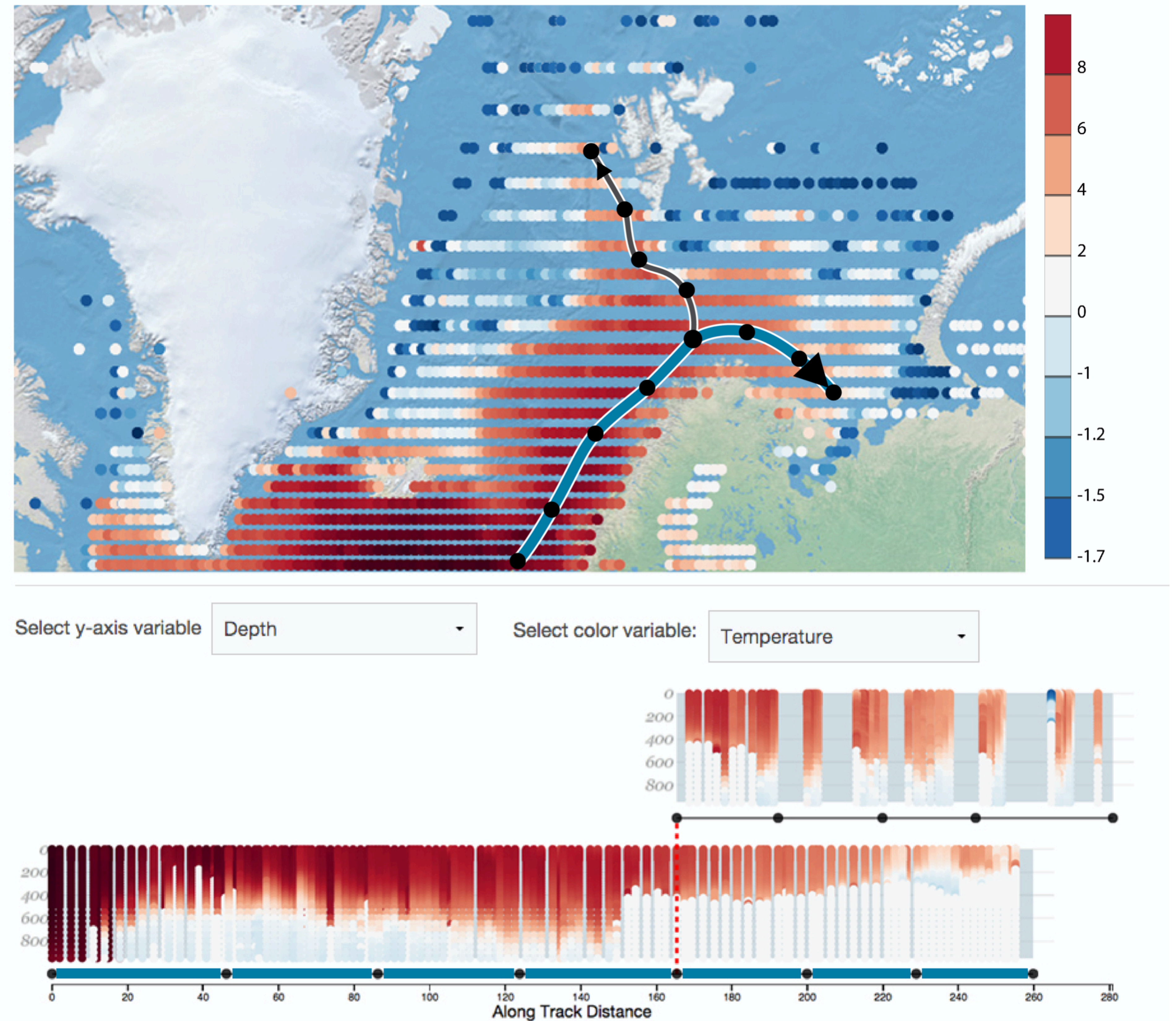
- 1. Select regions to show**
- 2. Select / derive data to show**

SELECTING REGIONS TO SHOW

PRINCIPLE

Select one or multiple items/regions
Show rich data about them in
separate view

SELECT A PATH IN A MAP

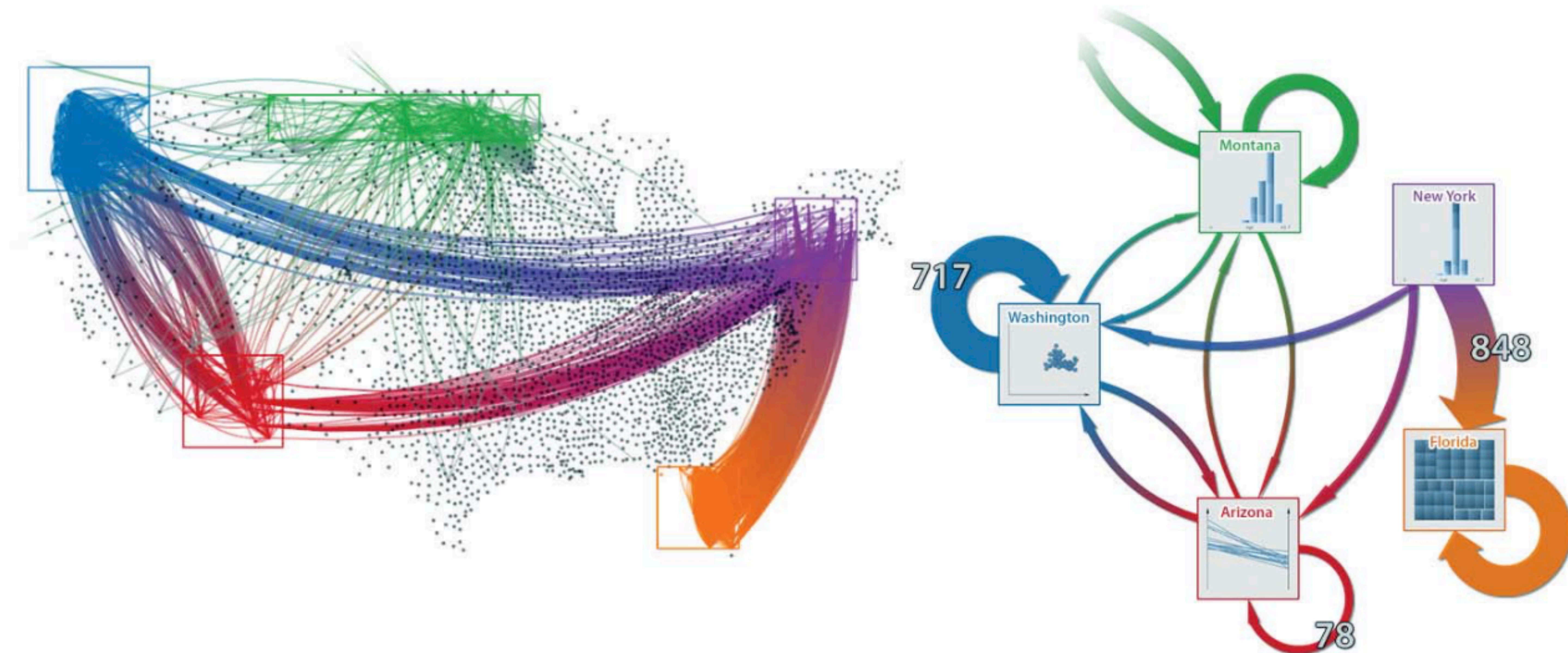


Carolina Nobre, Alexander Lex

OceanPaths: Visualizing Multivariate Oceanography Data

Proceedings of the Eurographics Conference on Visualization (EuroVis '15) - Short Papers, doi:10.2312/eurovisshort.20151124, 2015.

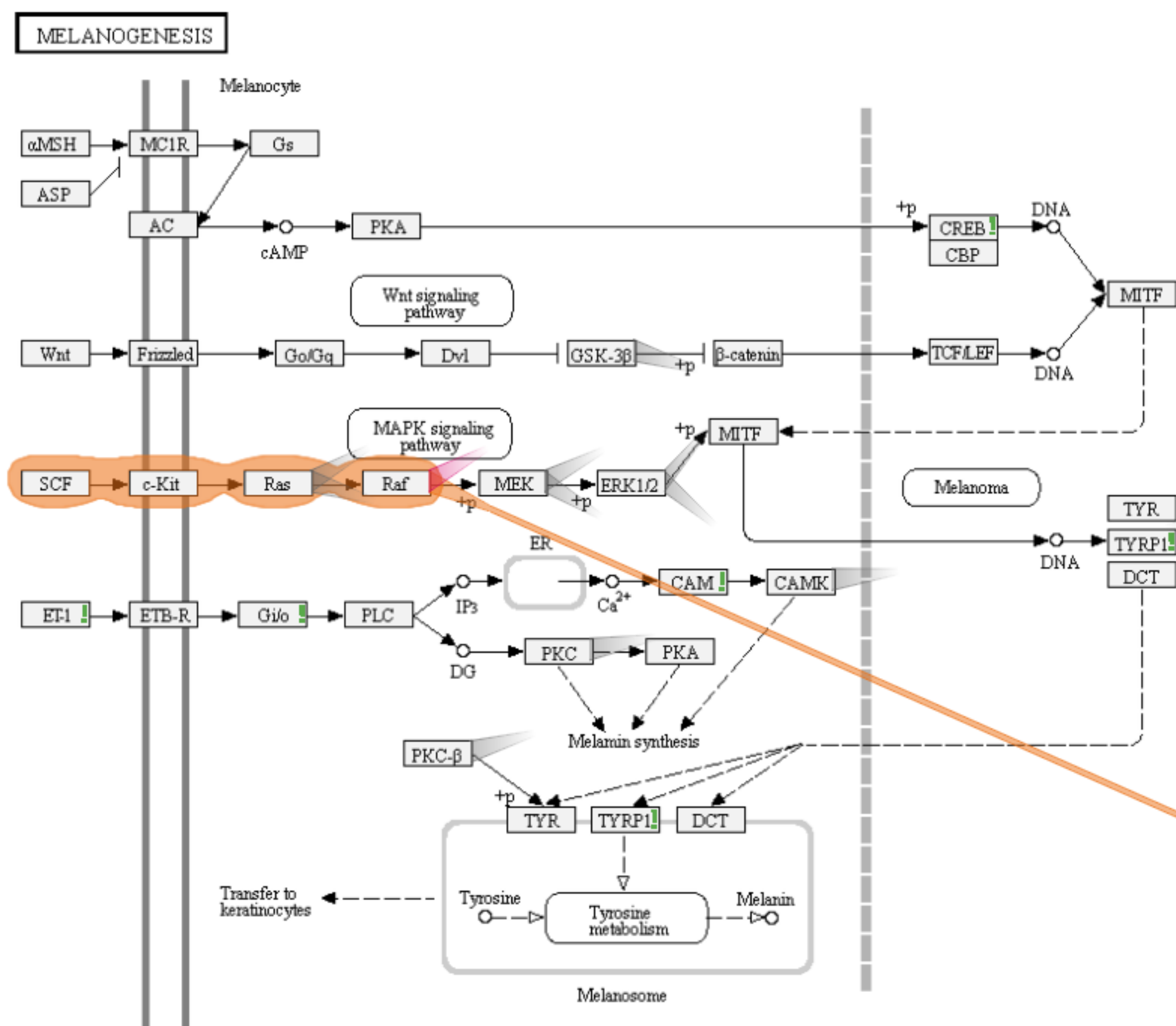
SELECT A REGION AND AGGREGATE!



S. van den Elzen and J. J. van Wijk, "Multivariate Network Exploration and Presentation: From Detail to Overview via Selections and Aggregations," IEEE Transactions on Visualization and Computer Graphics (InfoVis '14), vol. 20, no. 12, pp. 2310-2319, 2014, doi: 10.1109/TVCG.2014.2346441.

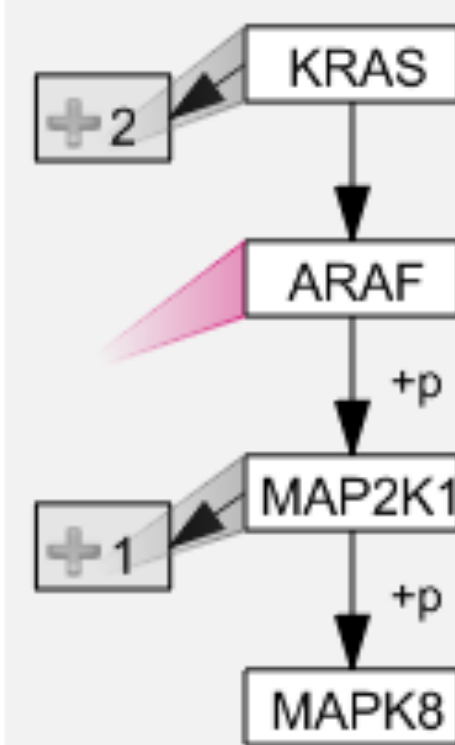
ENROUTE – PATH SELECTION

Melanogenesis

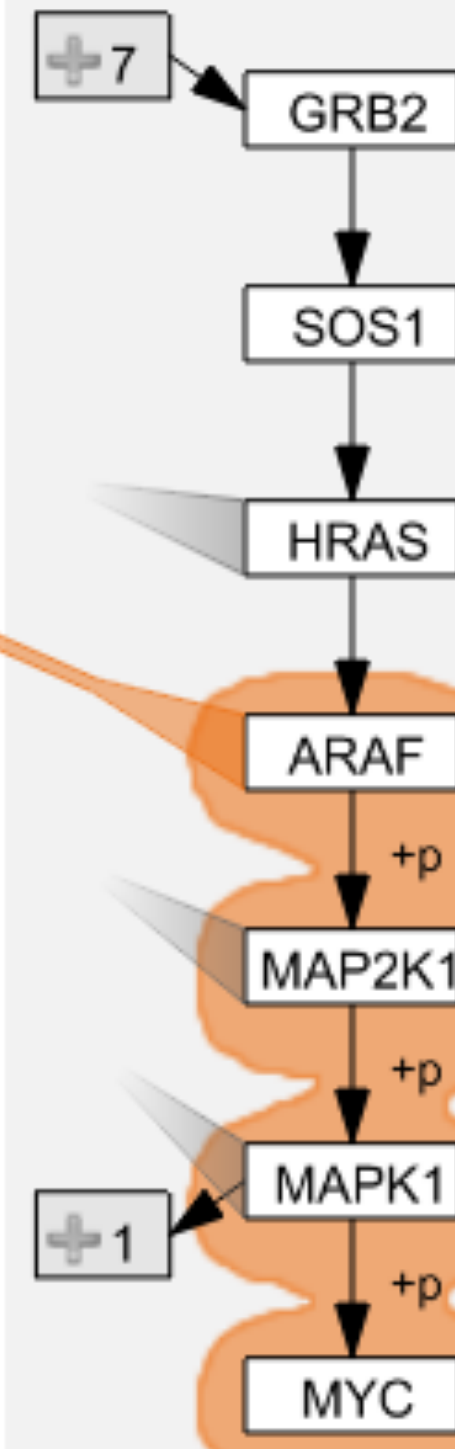


04916 10/16/12
(c) Kanehisa Laboratories

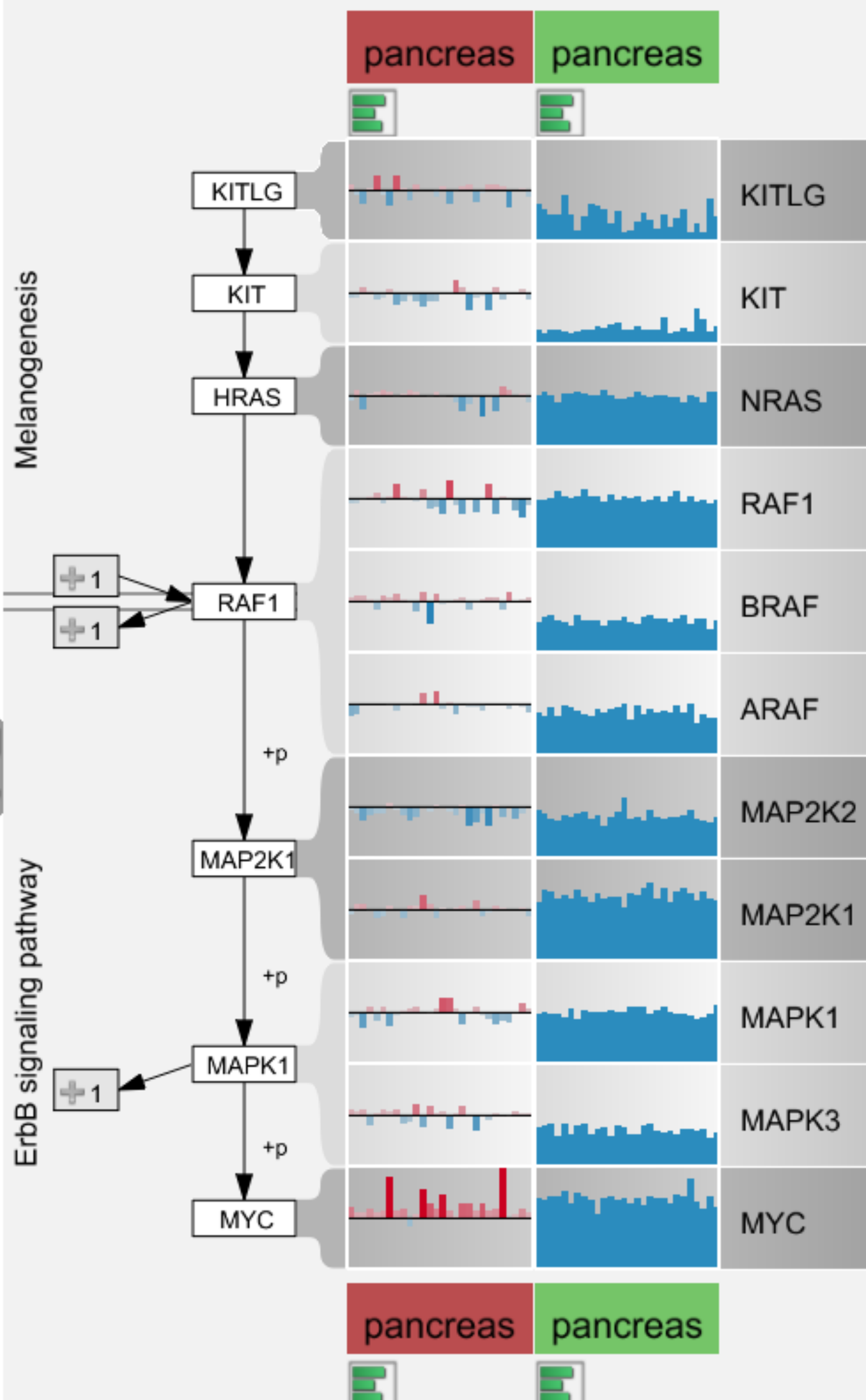
Pancreatic cancer



ErbB signaling pathway



Selected Path



**SELECT / DERIVE DATA
TO SHOW**

**SHOWING RAW DATA
IS HOPELESS!**

But do you need to?

Show average expression (etc)

**Show average expression for
pathway of interest**

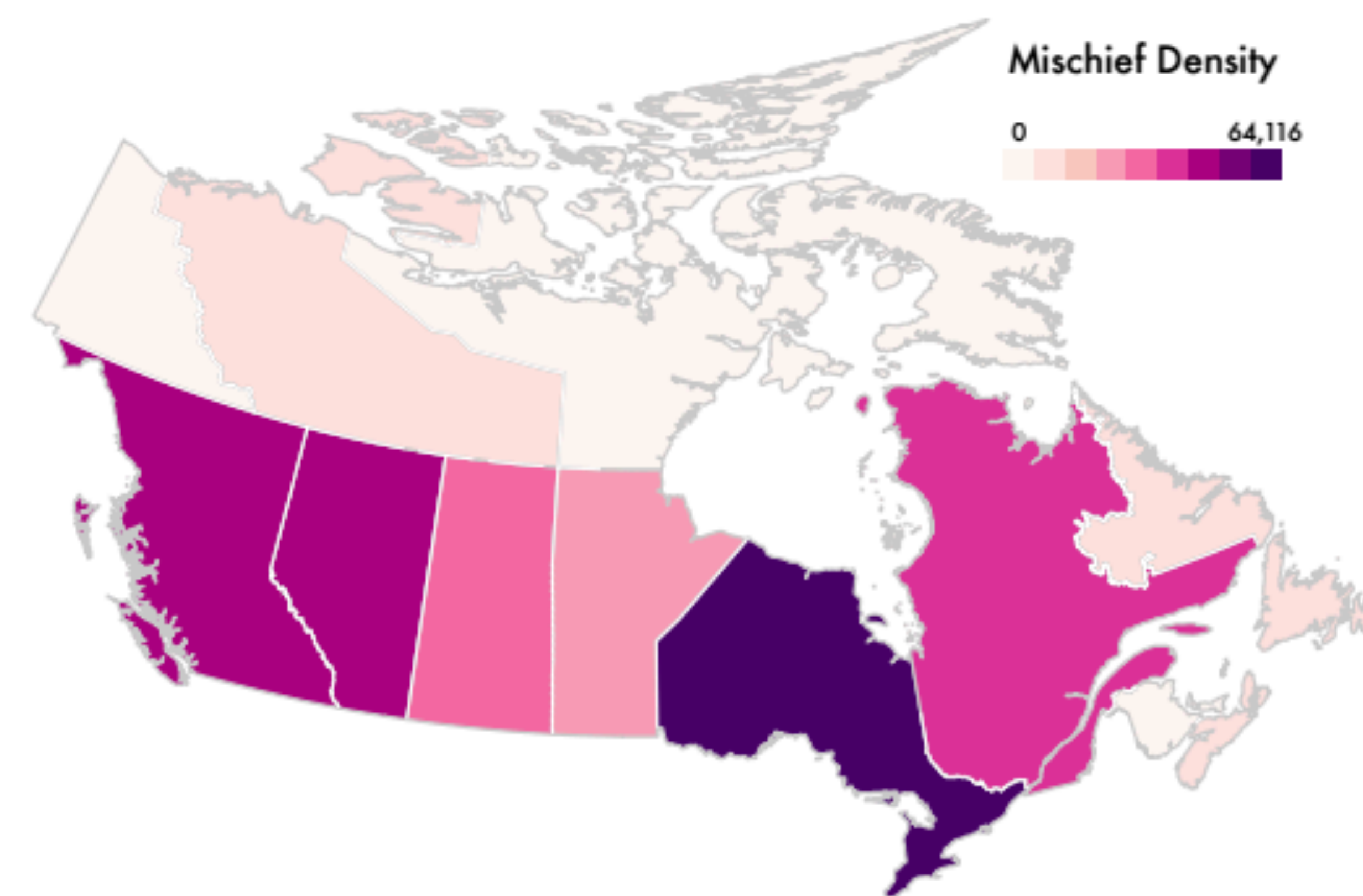
Filter out uninteresting items

Create domain specific scores

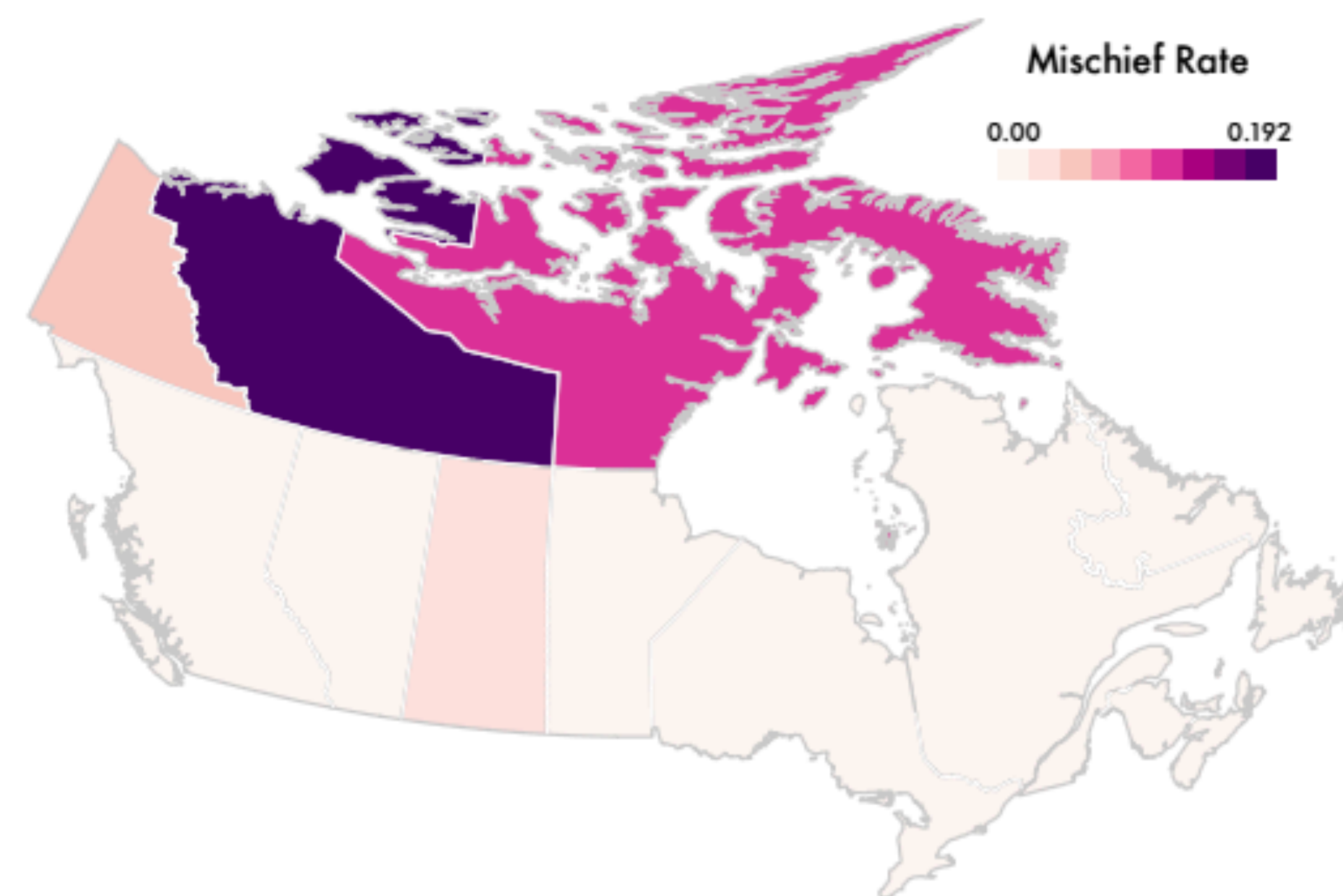
Create clusters/archetypes

...

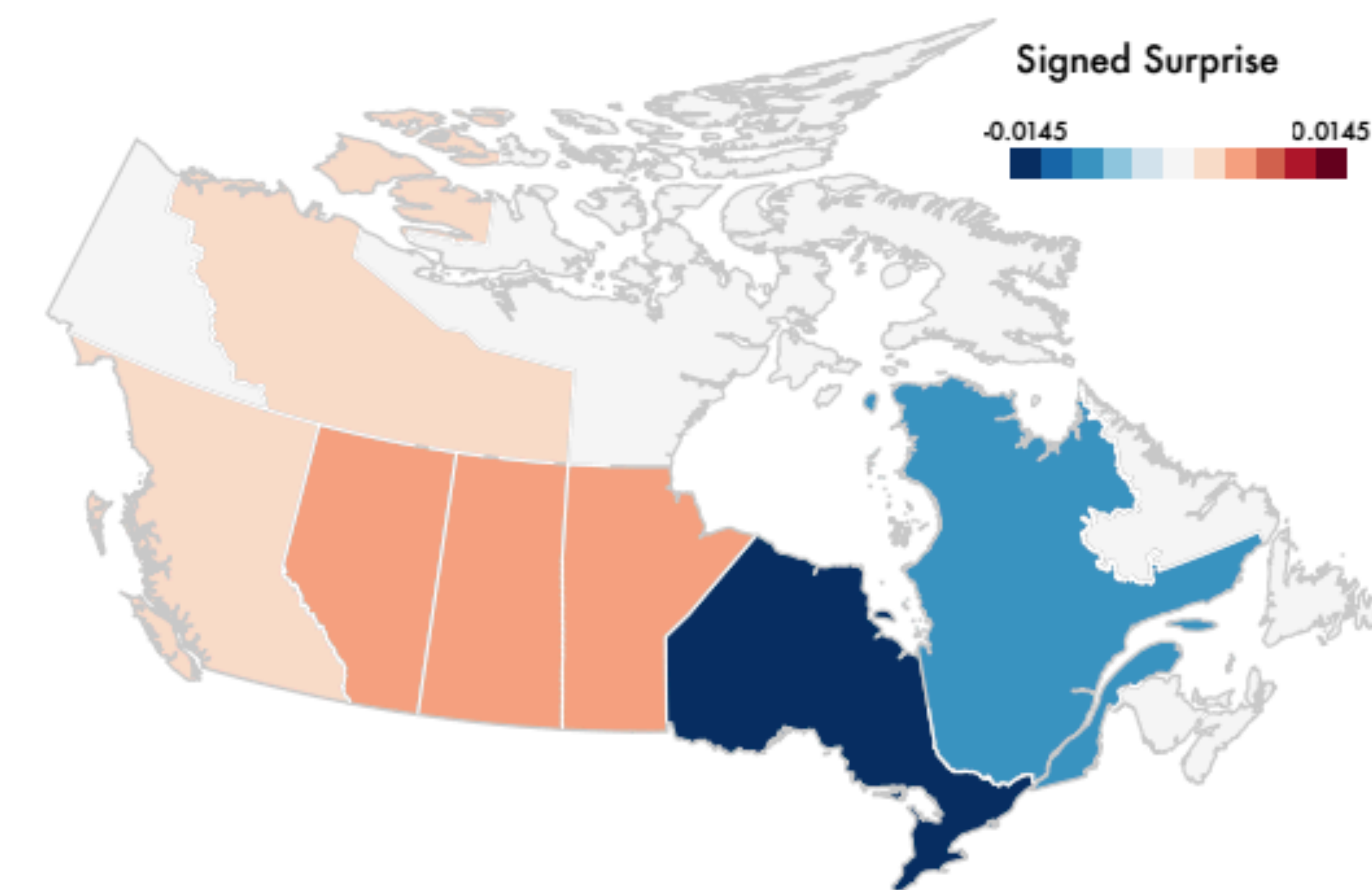
APPROACH: USE A PRIOR, SHOW DIFFERENCE.



(a) The **Event Density** of “mischief” in Canada.



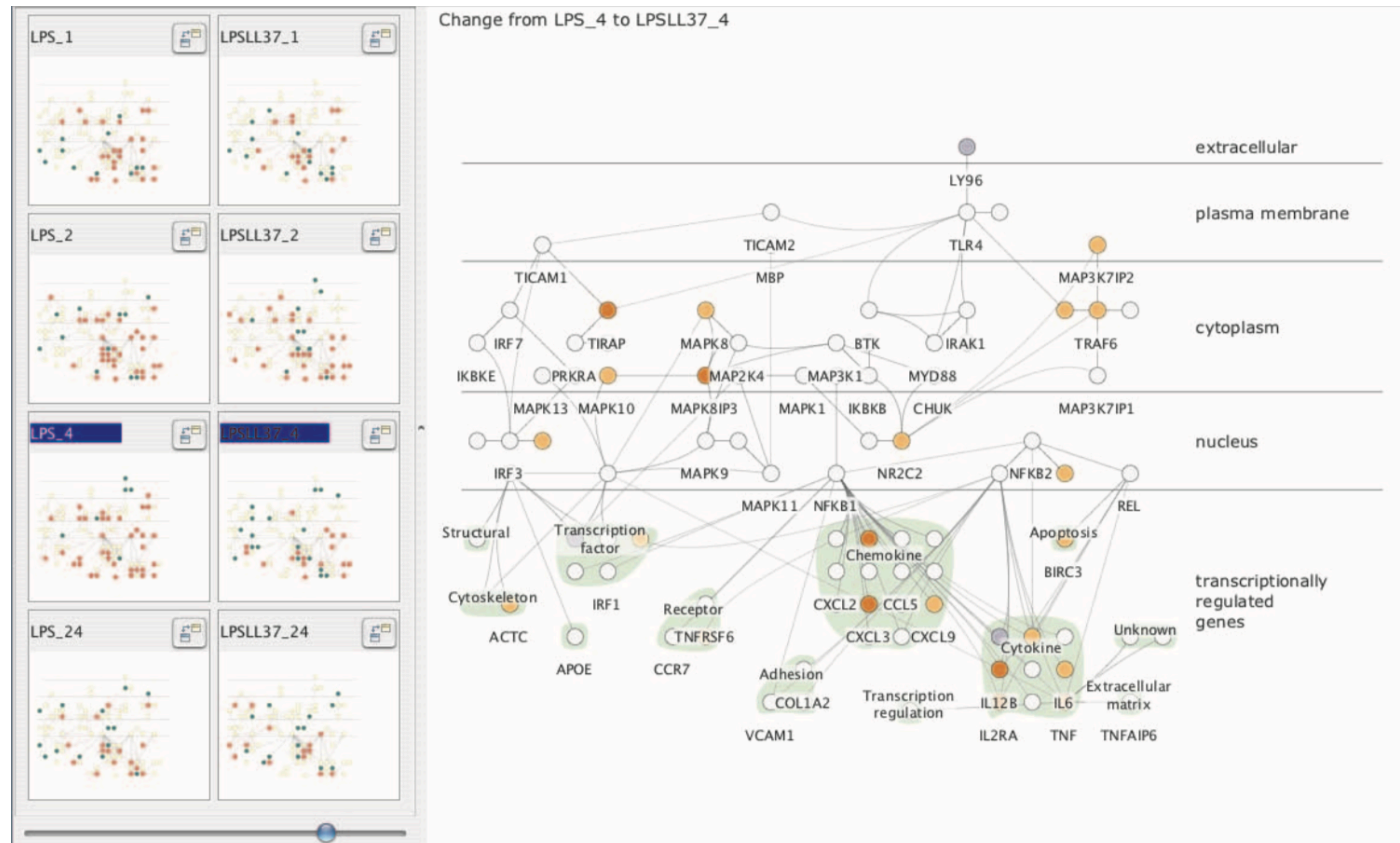
(b) The per-capita **Event Rate** of mischief.



(c) The **Surprise Map** of mischief.

model of population density +
accounting for variability when
analyzing small numbers

HAVE A HANDFUL OF SCORES? VIS FEASIBLE!



CLUSTERGRAMMER JUST EARLIER AT BIOVIS: DYNAMIC FILTERS

Clustergrammer: an interactive tool to visualize

Tissue Location:

Example with public dataset from 10X Genomics
Visium spatial RNA transcriptomics (mouse brain)
Visium Tissue: Zvint

<https://clustergrammer.org>

Sacha Gnajatic
Nick Fernandez
Avi Ma'ayan

Cell Type

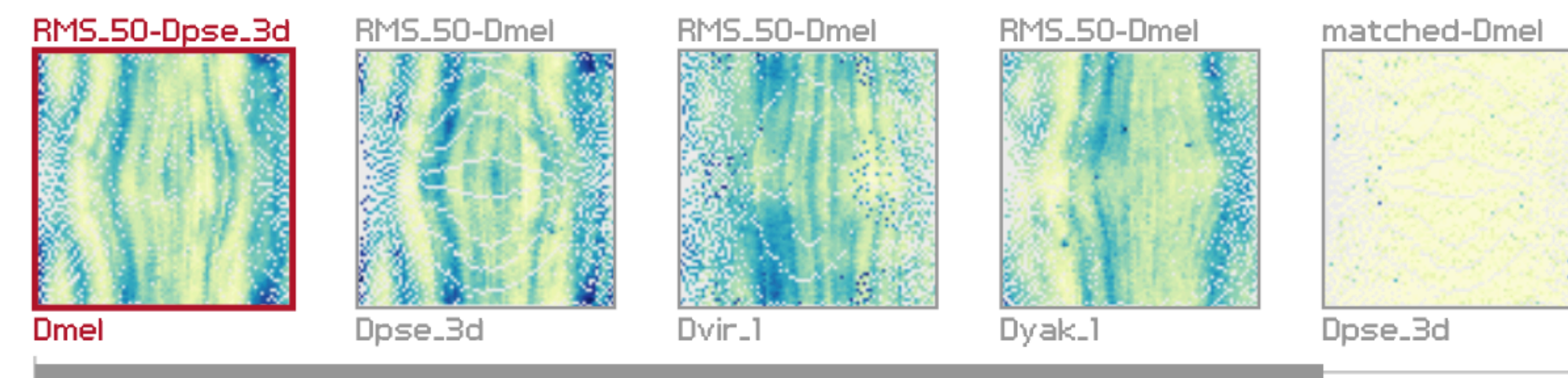
Cell Type	Count	Pct	P-val
Cell Type	91	83.3	2.40e-03
Cell Type	8	7.3	0.0
Cell Type	7	6.4	1.37e-03
Cell Type	3	2.8	0.00
Cell Type	1	0.9	0.00

Selected Columns

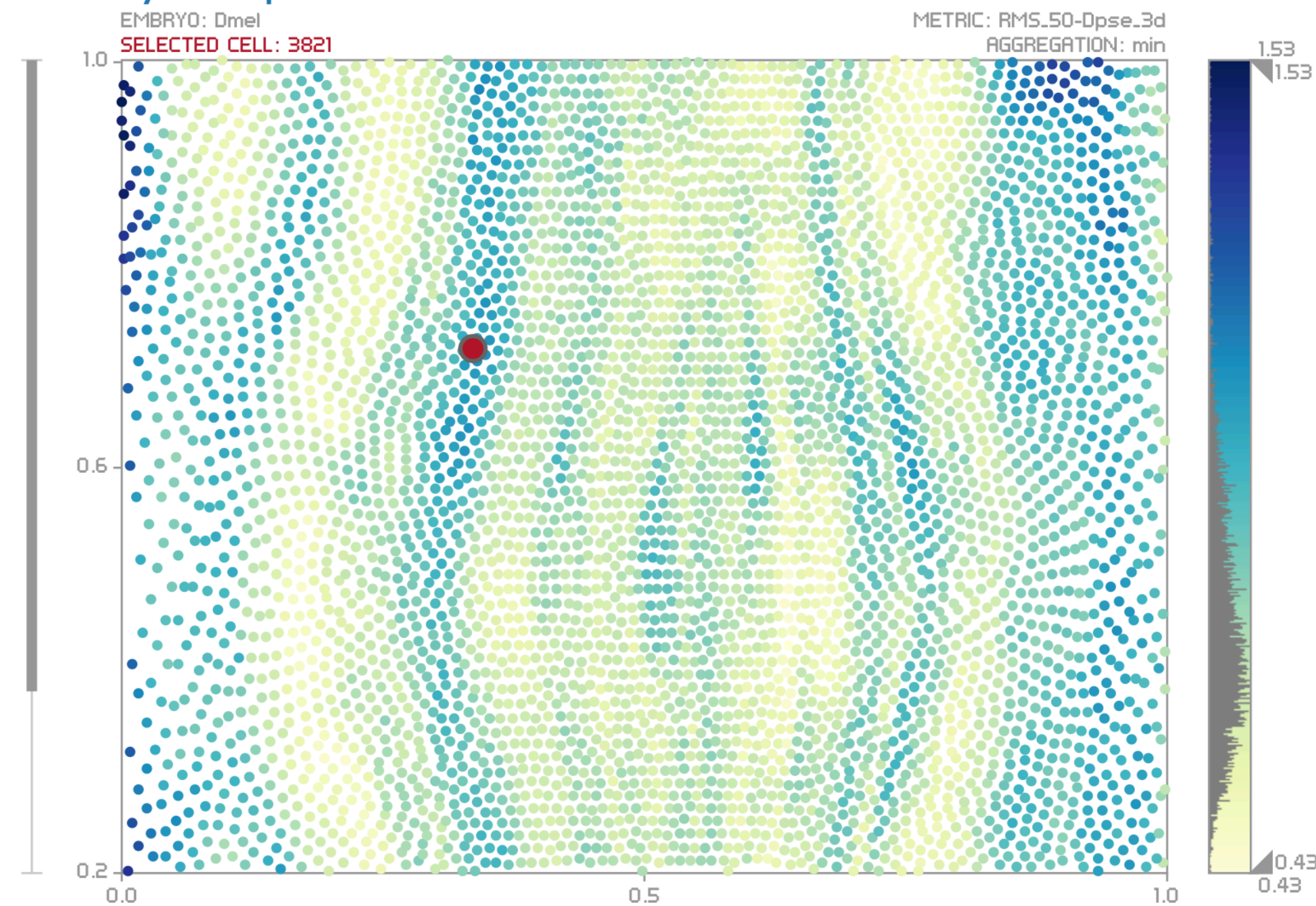
VIS2021

EXAMPLE: DOMAIN SPECIFIC SCORE

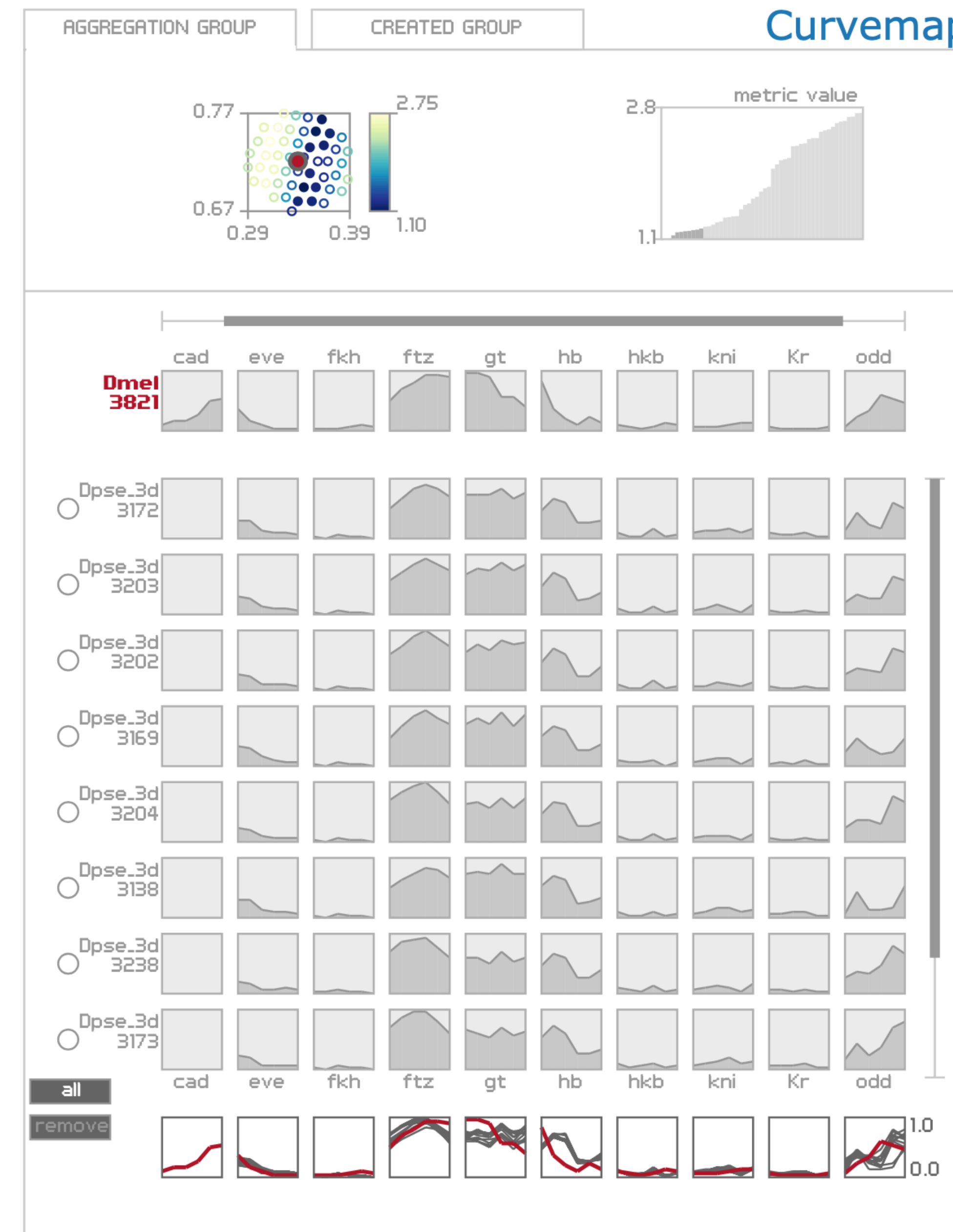
Summaries



Embryo Map



Curvemap

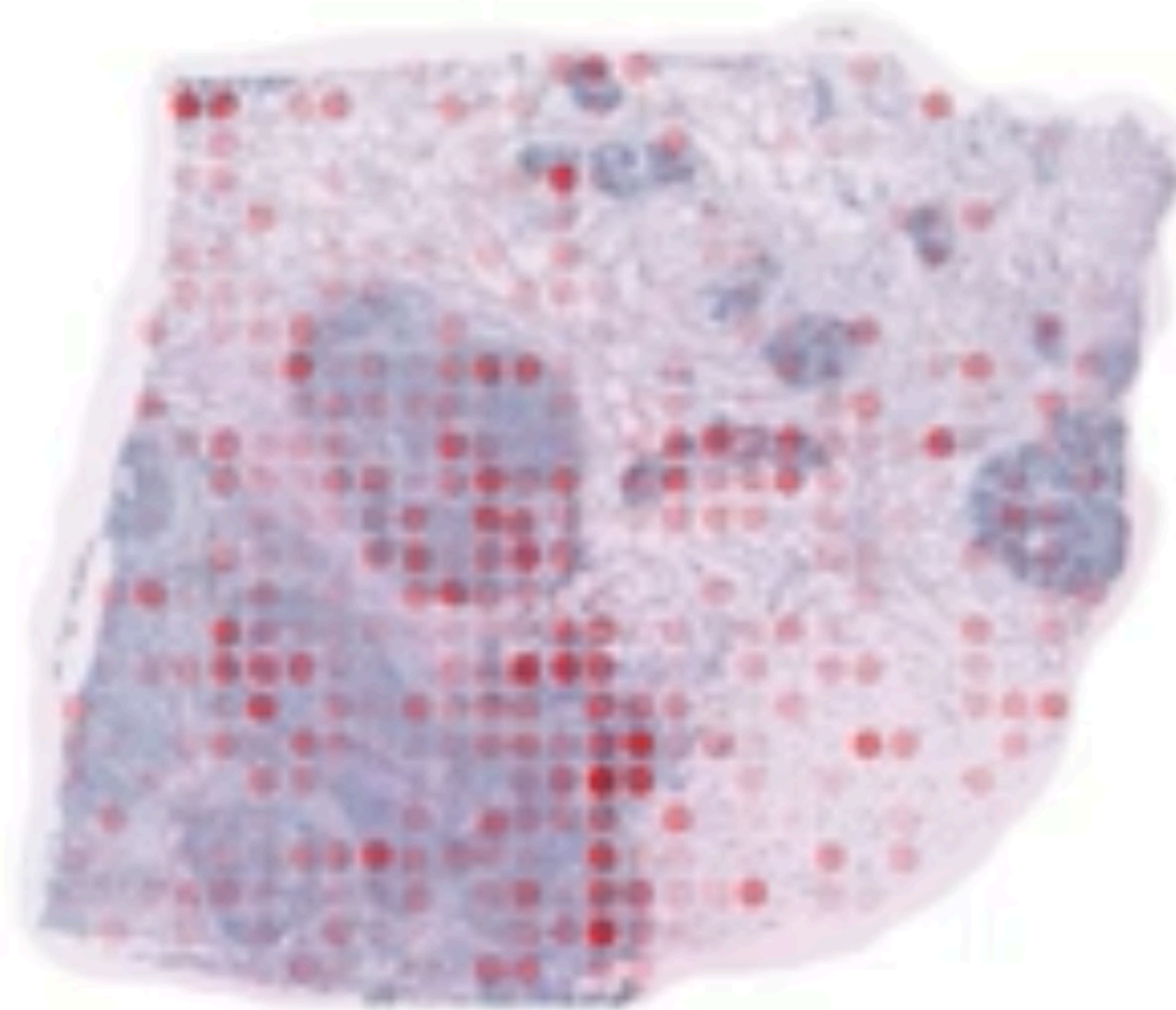


Miriah Meyer, Tamara Munzner, Angela DePace, Hanspeter Pfister
 MulteeSum: A Tool for Comparative Spatial and Temporal Gene Expression Data
 IEEE Transactions on Visualization and Computer Graphics (InfoVis), 16(6): 908--917, doi:10.1109/TVCG.2010.137, 2010.

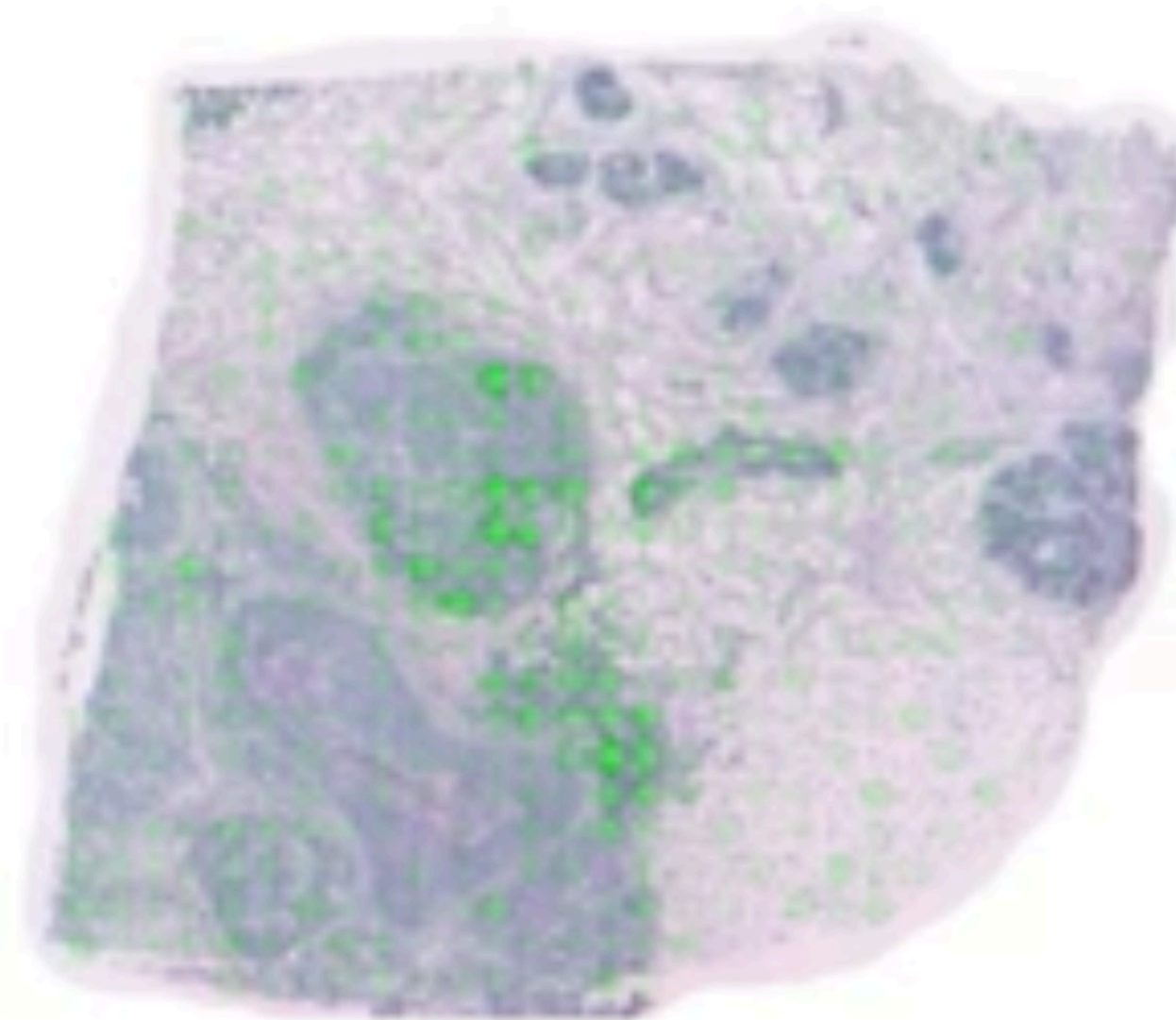
USING SCORES @ BIOVIS BY ALMA ANDERSON

■ ■ ■ Assessing co-localization

B-cells



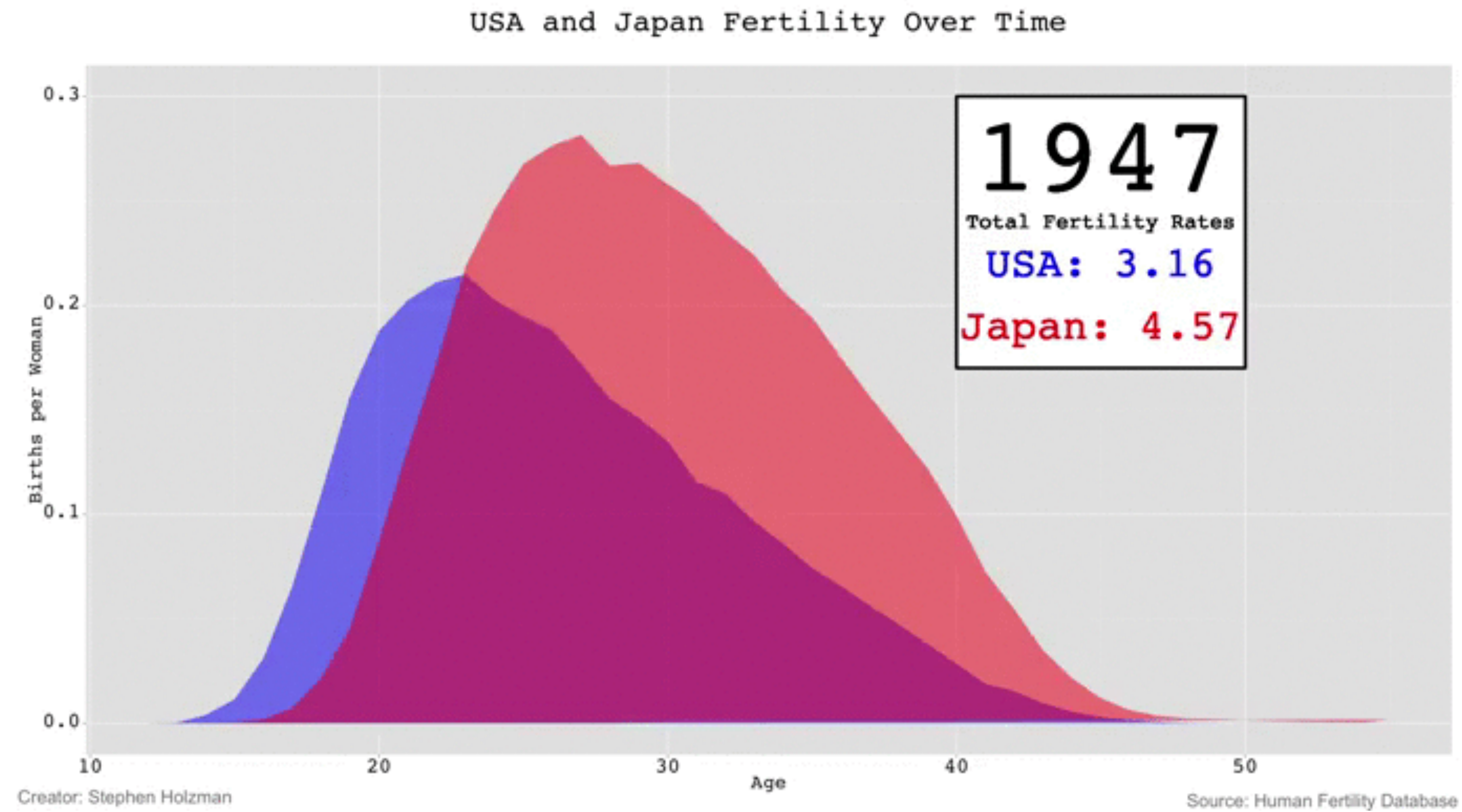
T-cells



Sample from breast cancer patient (HER2-positive)

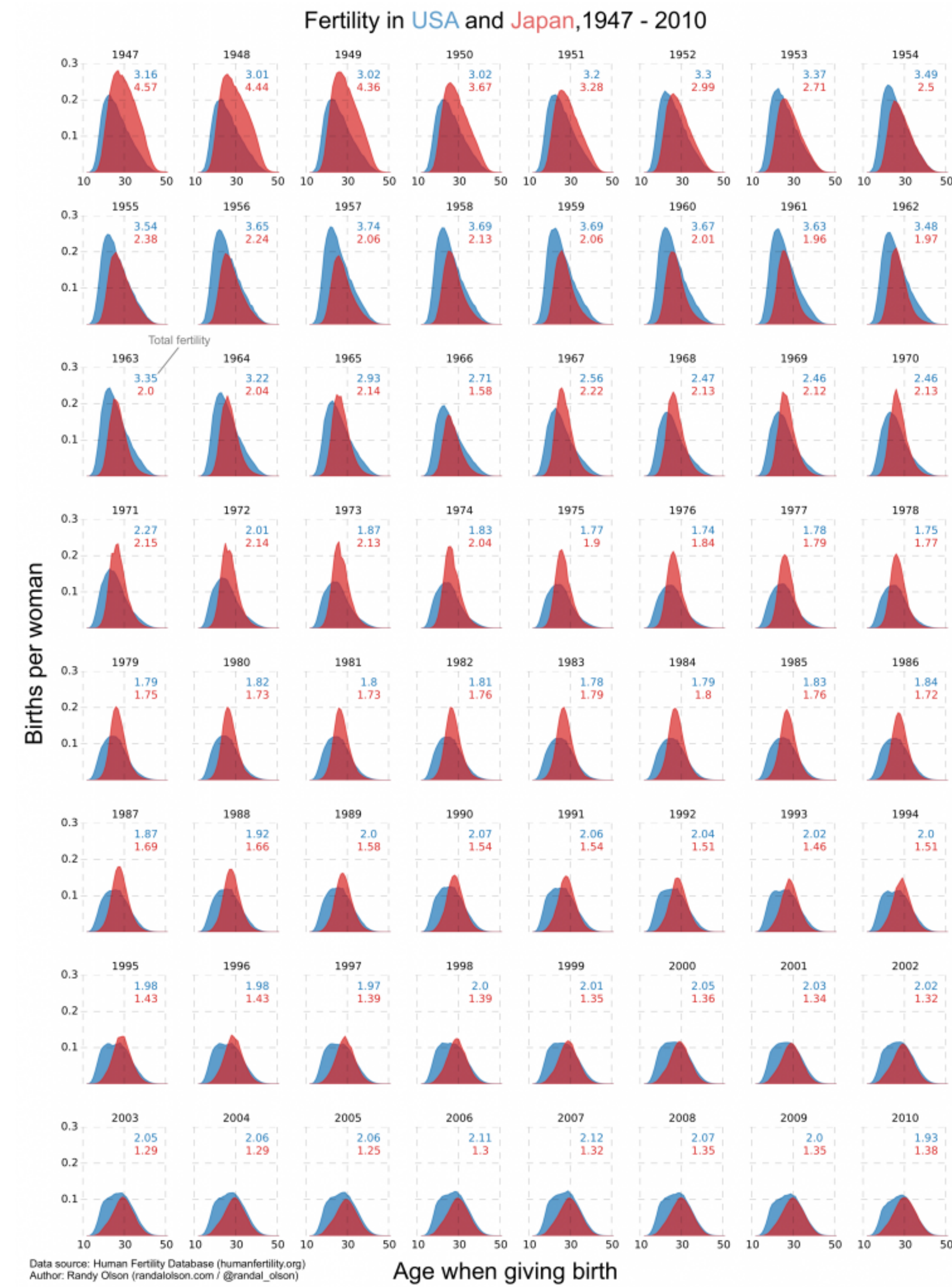
- We were interested in TLS-sites **enriched for both B and T-cells**; these have relevance for prognosis and understanding of the cancer
- Our immunologist wanted to relate this co-localization to the morphology
- **Solution:** transform to single metric (co-localization score) and show

REDESIGN EXAMPLE

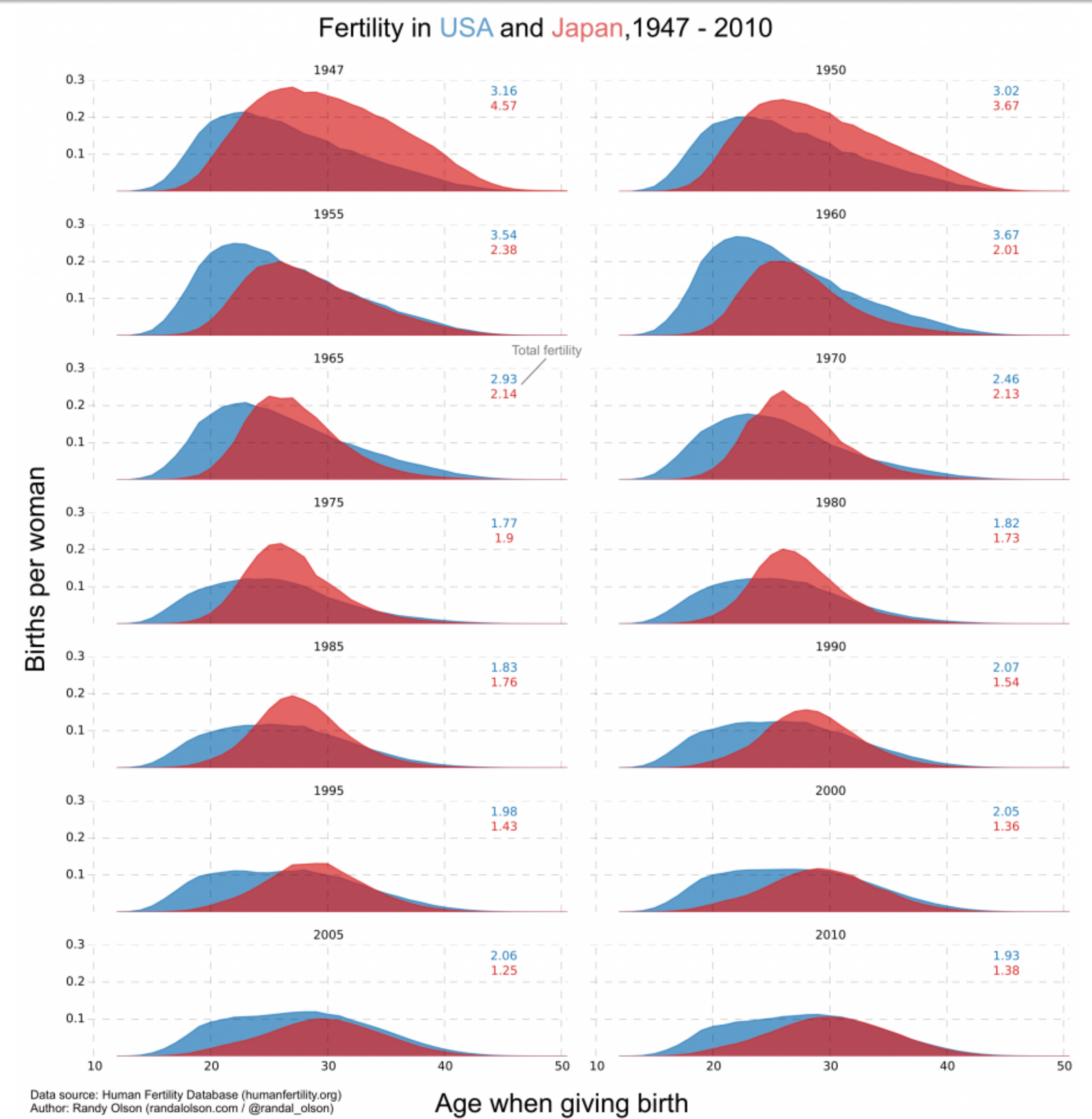


Randal S. Olson

EYES BEAT MEMORY: SMALL MULTIPLES

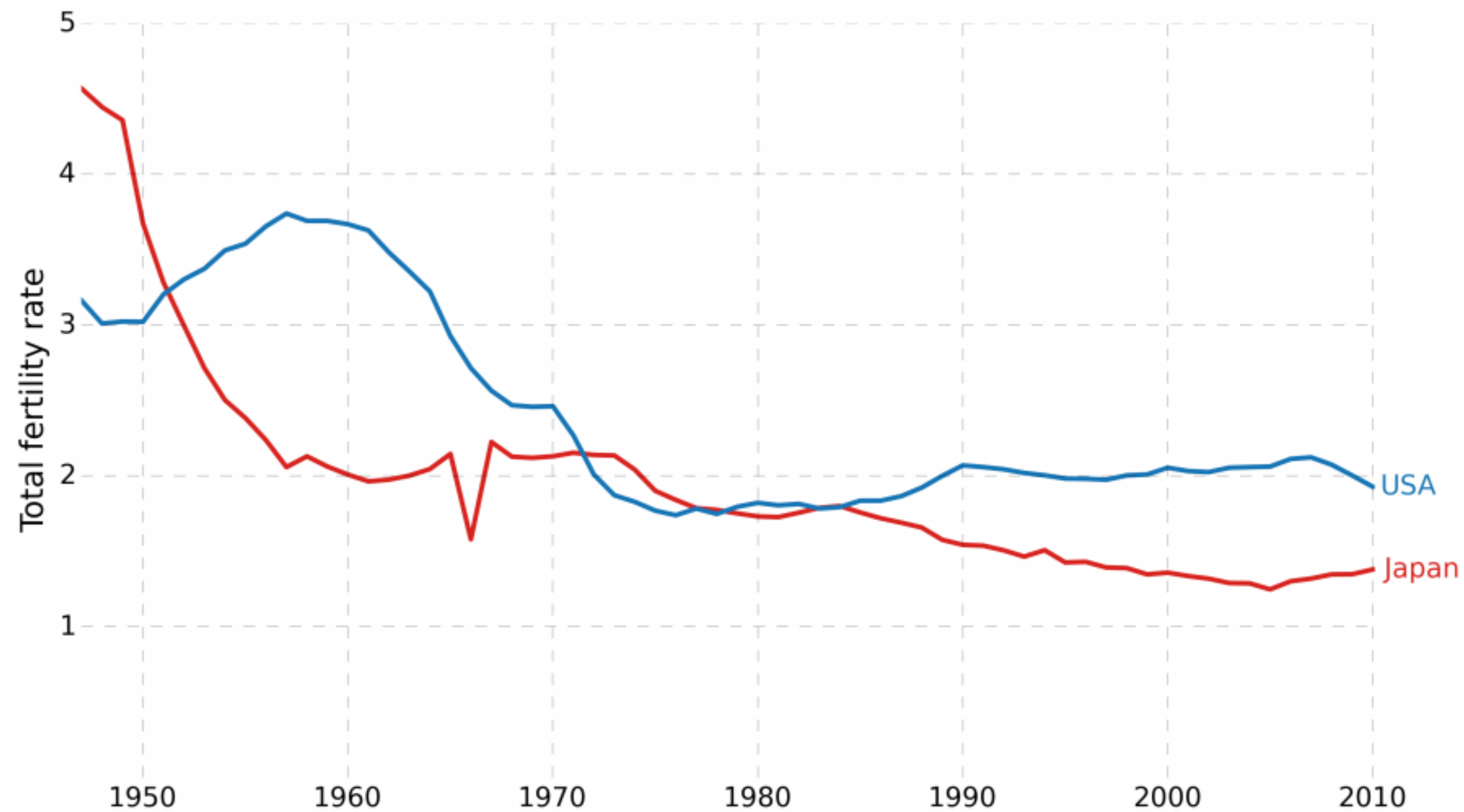


EYES BEAT MEMORY: SMALL MULTIPLES



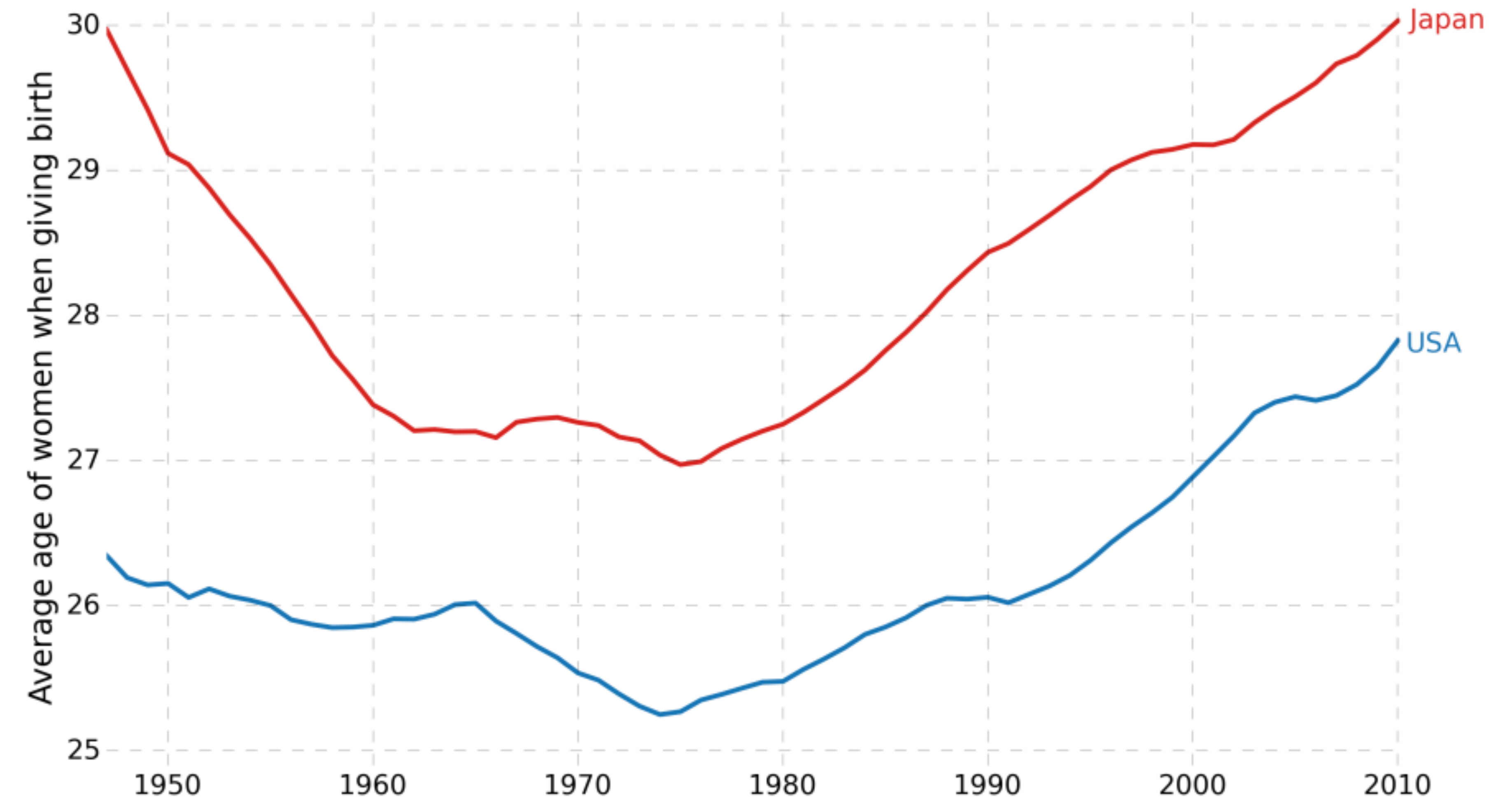
SIMPLIFY!

Total fertility rate in USA and Japan, 1947 - 2010



Data source: Human Fertility Database (humanfertility.org)
Author: Randy Olson (randalolson.com / @randal_olson)

Average age when giving birth in USA and Japan, 1947 - 2010



Data source: Human Fertility Database (humanfertility.org)
Author: Randy Olson (randalolson.com / @randal_olson)

CONCLUSION

You can't show all the data!

But you can show what's important about the data.

Build tools that give analysts the ability to show that!

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Thanks to: Carolina Nobre, Kiran Gadhave, Jen Rogers, Haihan Lin, Dylan Wootton, Jochen Görtler, Oliver Deussen, Miriah Meyer, Jeff Phillips, Samuel Gratzl, Holger Stitz, Marc Streit, Nils Gehlenborg, Hilary Coon, Lane Harrison, Hendrik Strobelt, Romain Vuillemot, Hanspeter Pfister, and many Others!



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