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# **Spatial Omics Visualizations:** Lessons Learned from Networks and Maps



# VISUAIZATION design lab

### THE UNIVERSITY OF UTAH





# **EVERYBODY IS TALKING ABOUT SPATIAL OMICS!**

# Method of the Year: spatially resolved transcriptomics

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

Vivien Marx

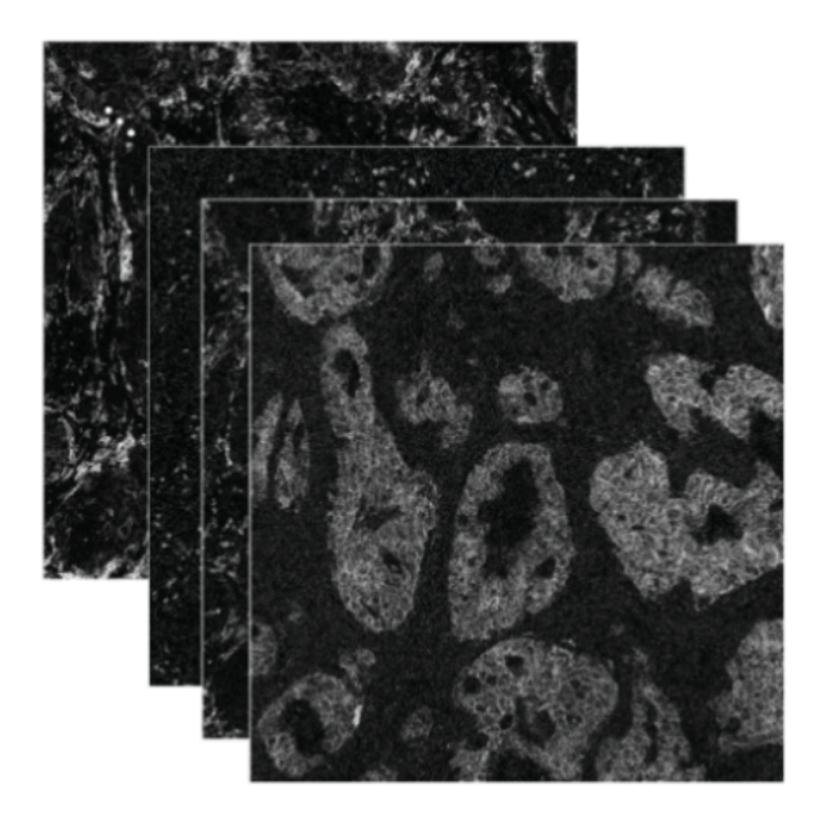
f a researcher is making a smoothie, it might be snack time. Or it could be the moment to prepare a sample for bulk RNA sequencing, in which tissue is homogenized and analyzed to yield

### FOCUS | TECHNOLOGY FEATURE

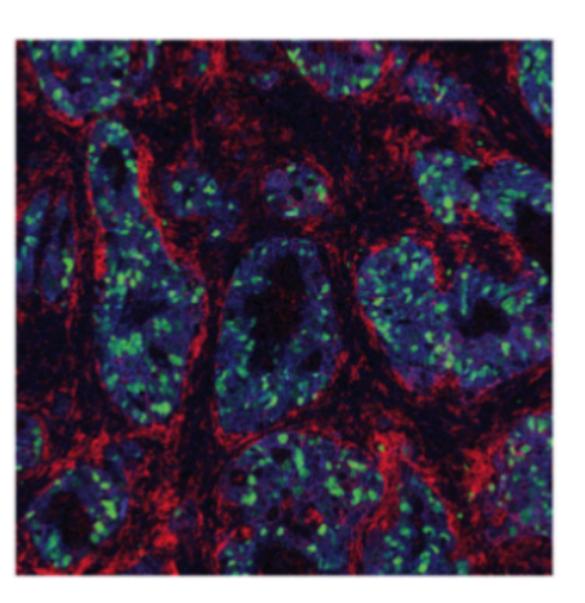


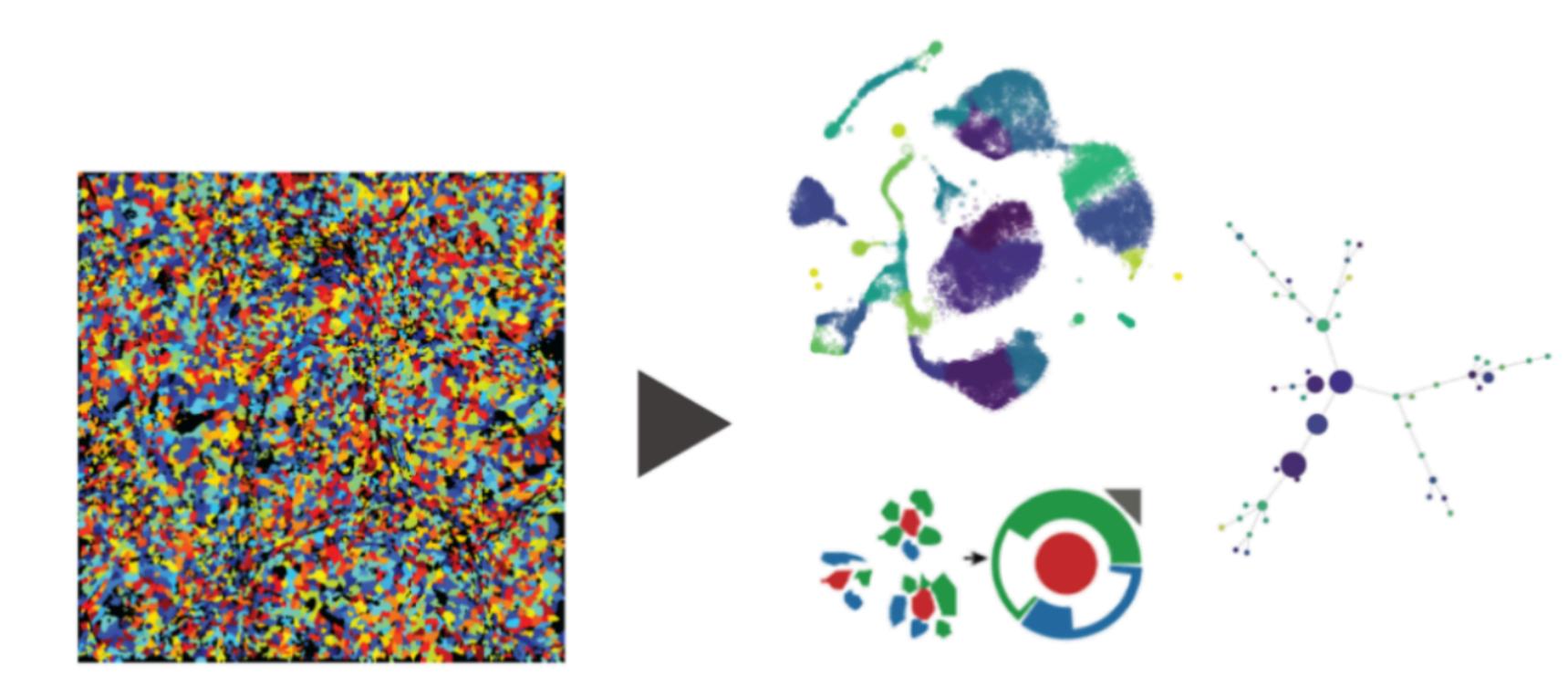


### BIOVIS CHALLENGE: FROM IMAGES TO ANALYSIS



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

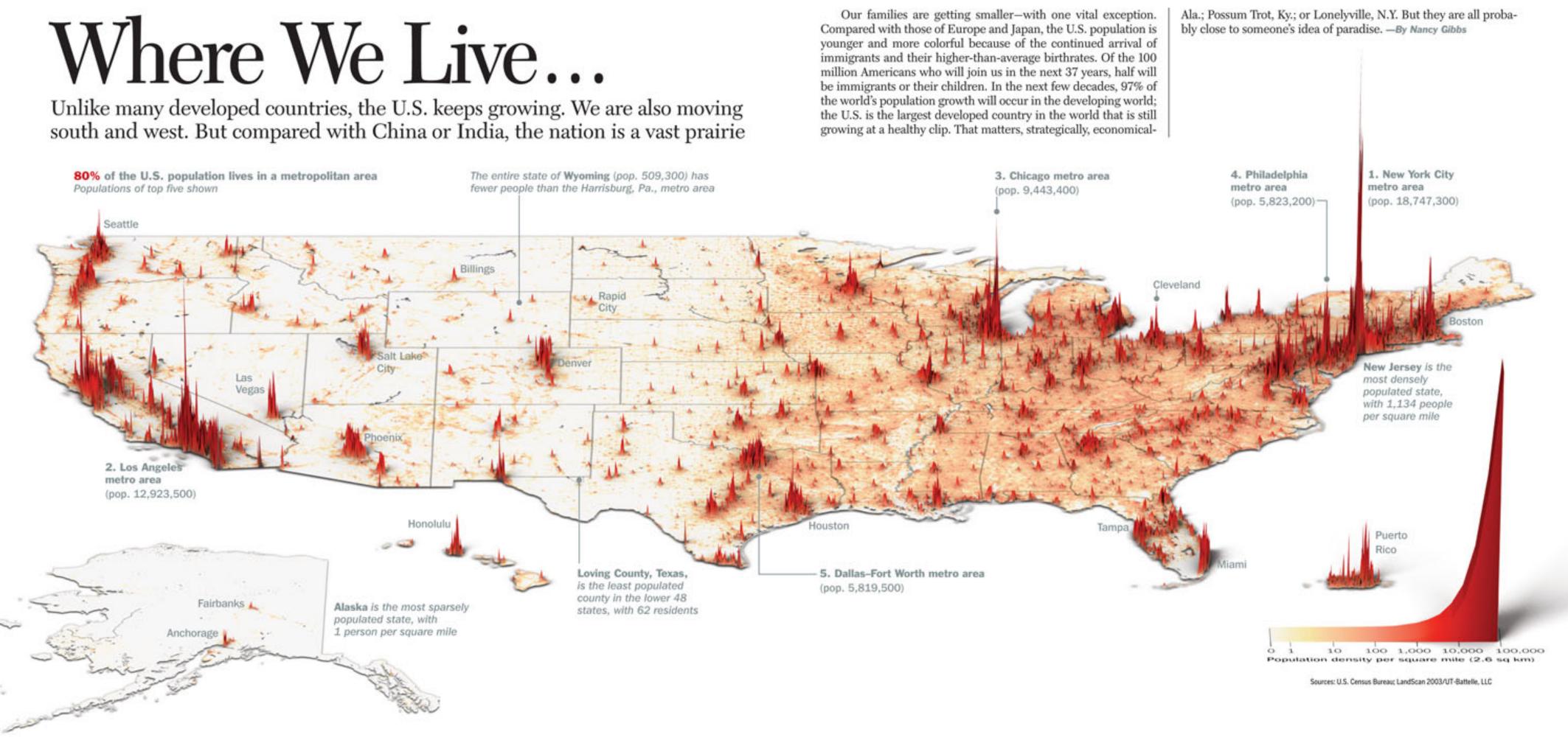
# SPATIAL OMICS VISUALIZATION CHALLENGES

### High dimensional data Similar to "classical" omics data (except for scale) **Spatial location / proximity is important NOT encountered in "classical" omics data.**

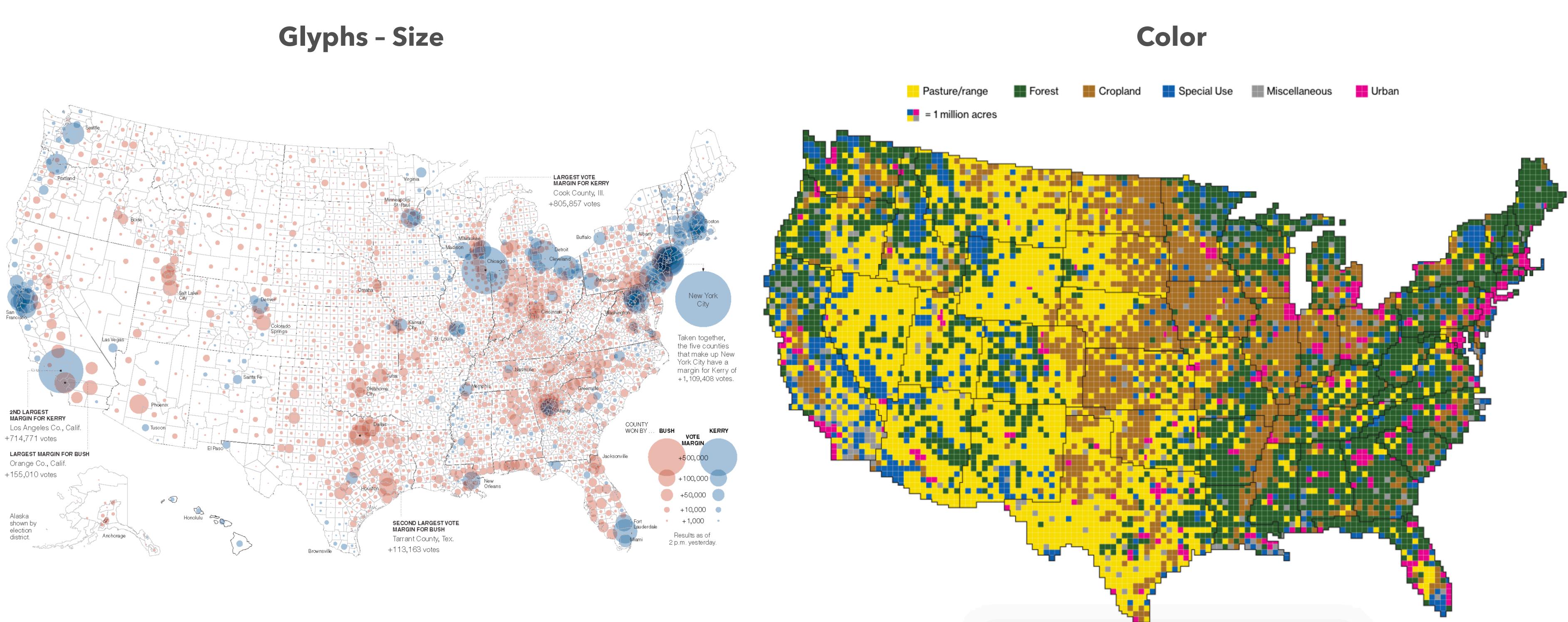


### ANALOGY: MAPS!

### **Fixed location** Potentially high-D vector for each location

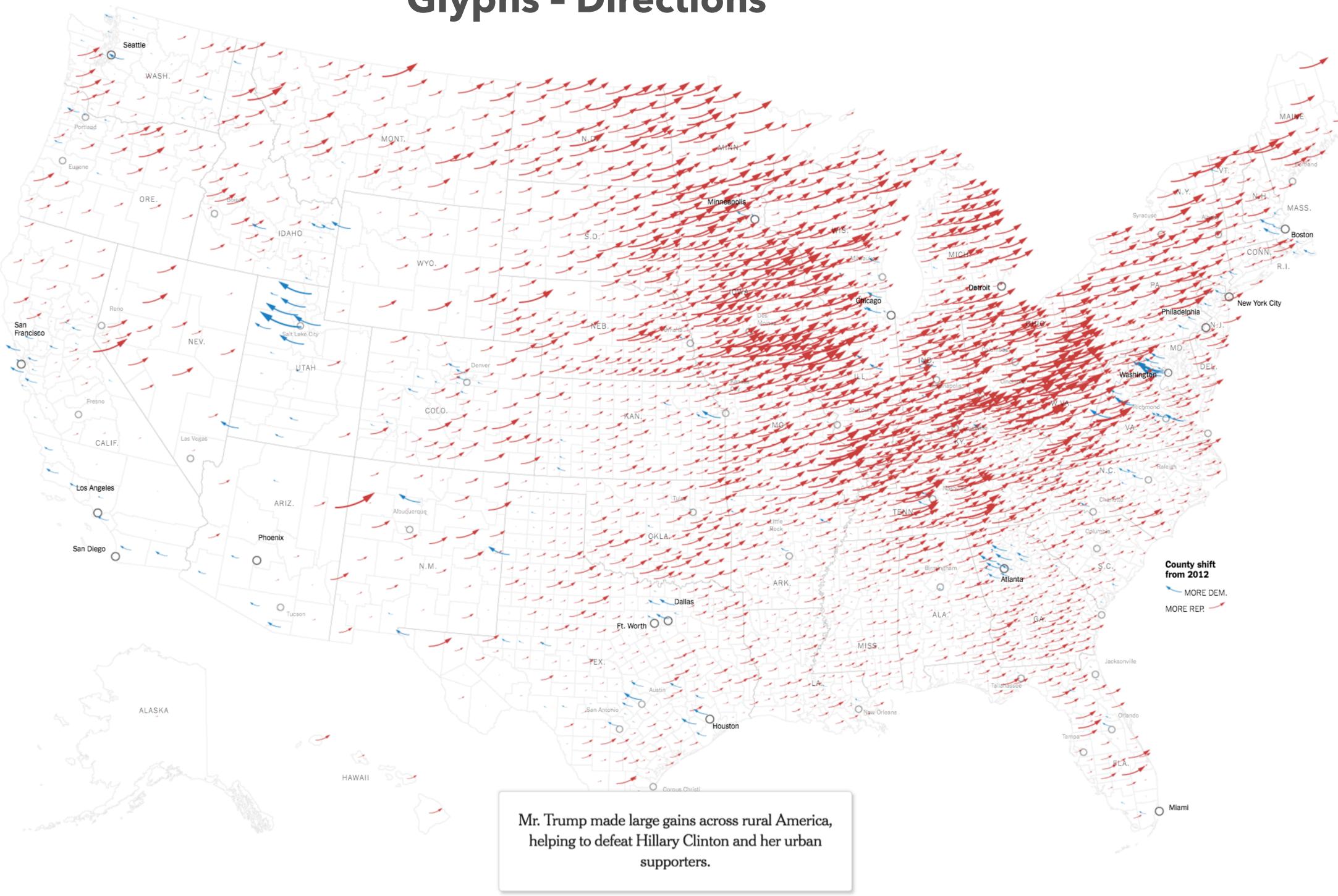


### SINGLE DATA VALUE + SPATIAL LOCATION



Blomberg

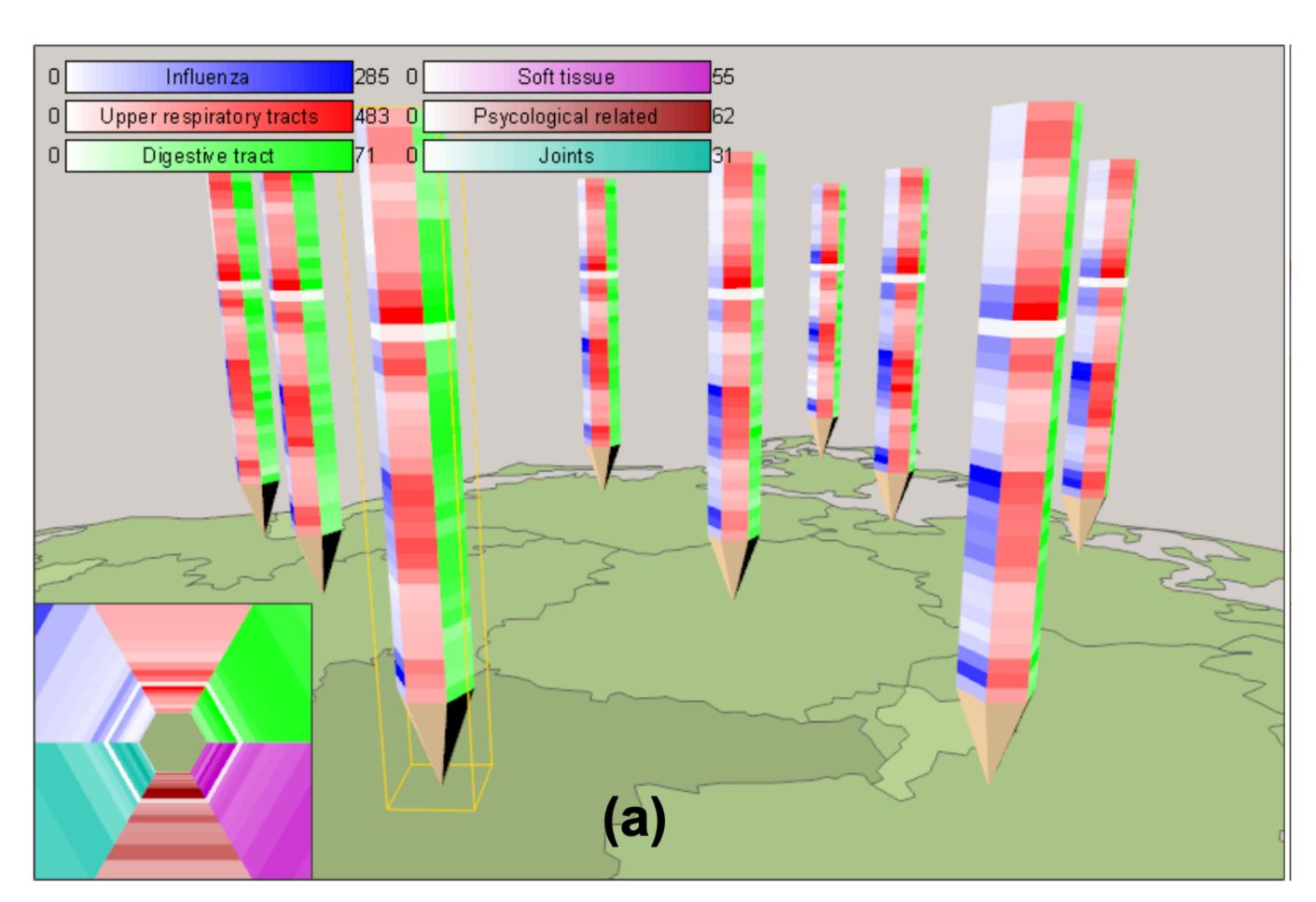
## SINGLE DATA VALUE + SPATIAL LOCATION





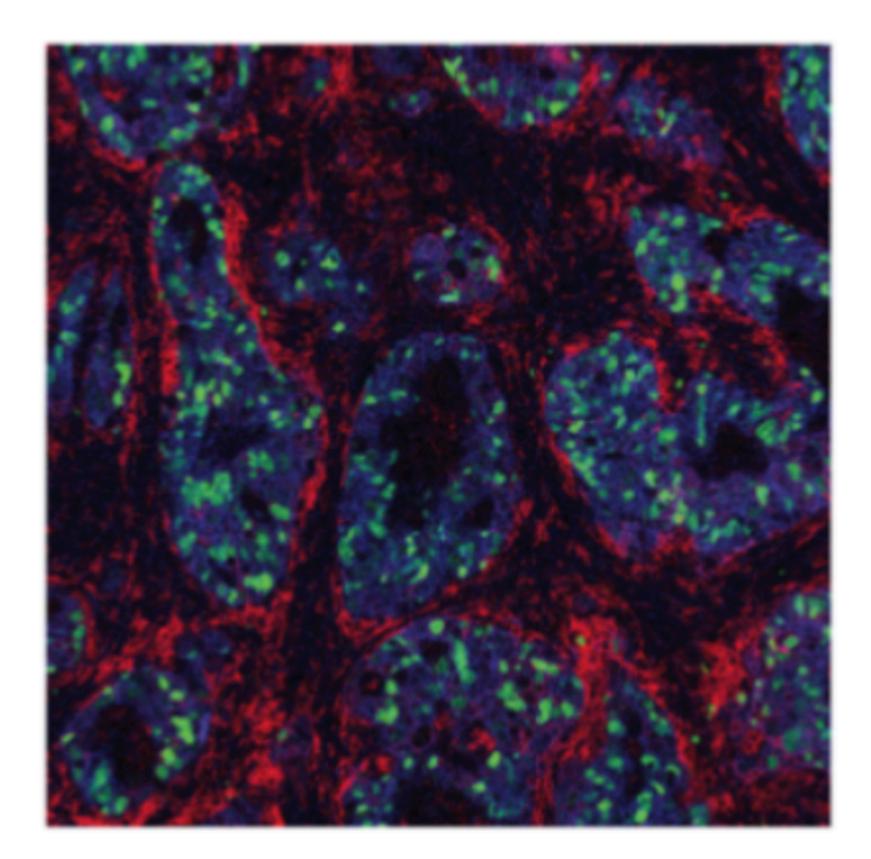
# BUT WHAT IF YOU HAVE MORE THAN ONE DATA VALUE?

### Glyphs?





### Don't treat color channels as separate visual channels



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



### DO WE REALLY NEED A MAP?

Chart Size of Lead Chart Electoral Votes

### **Obama Re-elected**

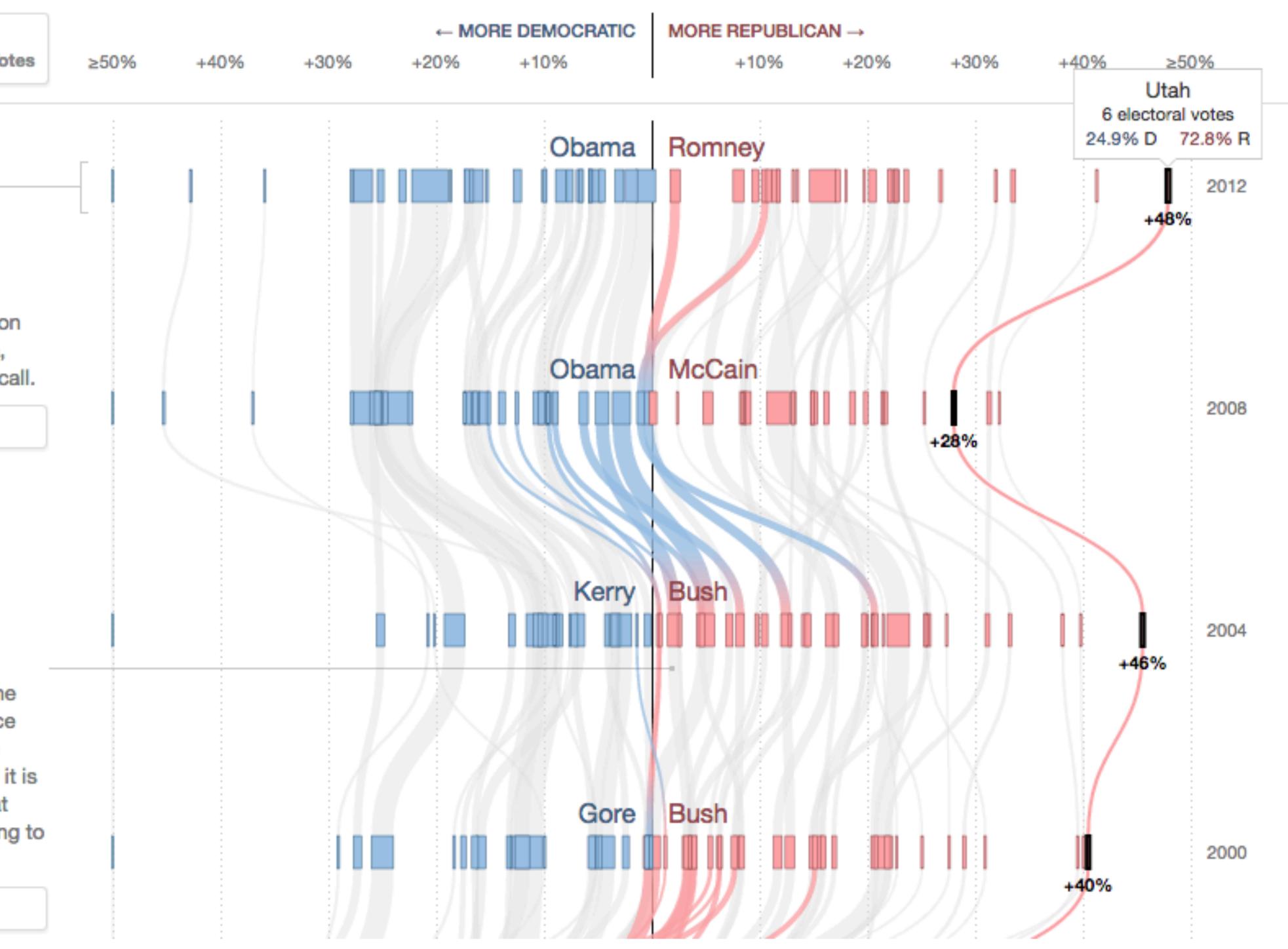
The country voted about 5 percentage points more Republican in 2012 than in 2008. Obama lost North Carolina and Indiana, but won every tossup except Florida, which remains too close to call.

Highlight Tossups

### As Goes Ohio

Ohio, which has voted for the winner in every election since 1964, provided the decisive electoral votes in 2004, and it is the state likeliest to play that role again this year, according to the FiveThirtyEight model.

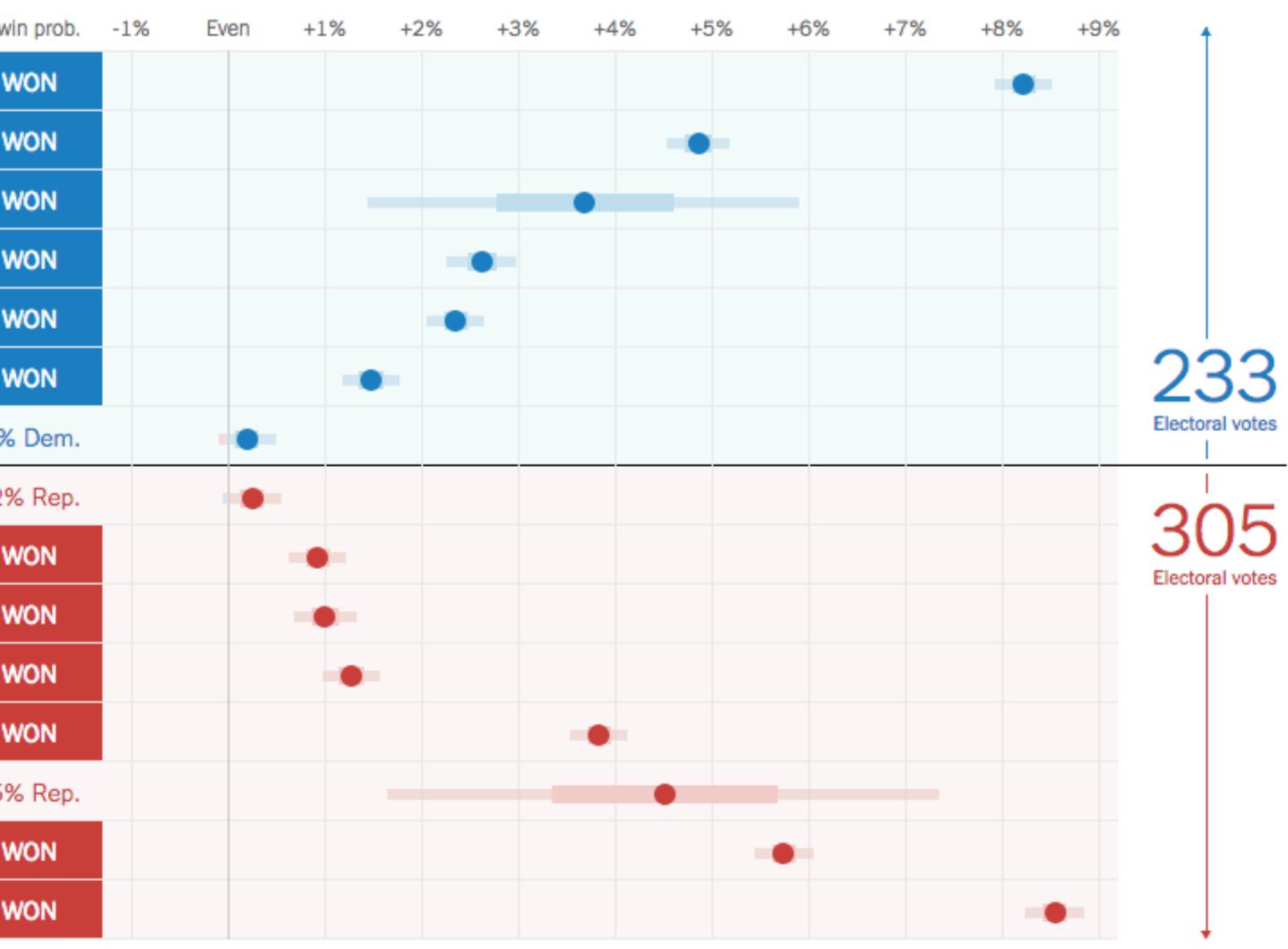
Highlight Ohio



# DO WE REALLY NEED A MAP?

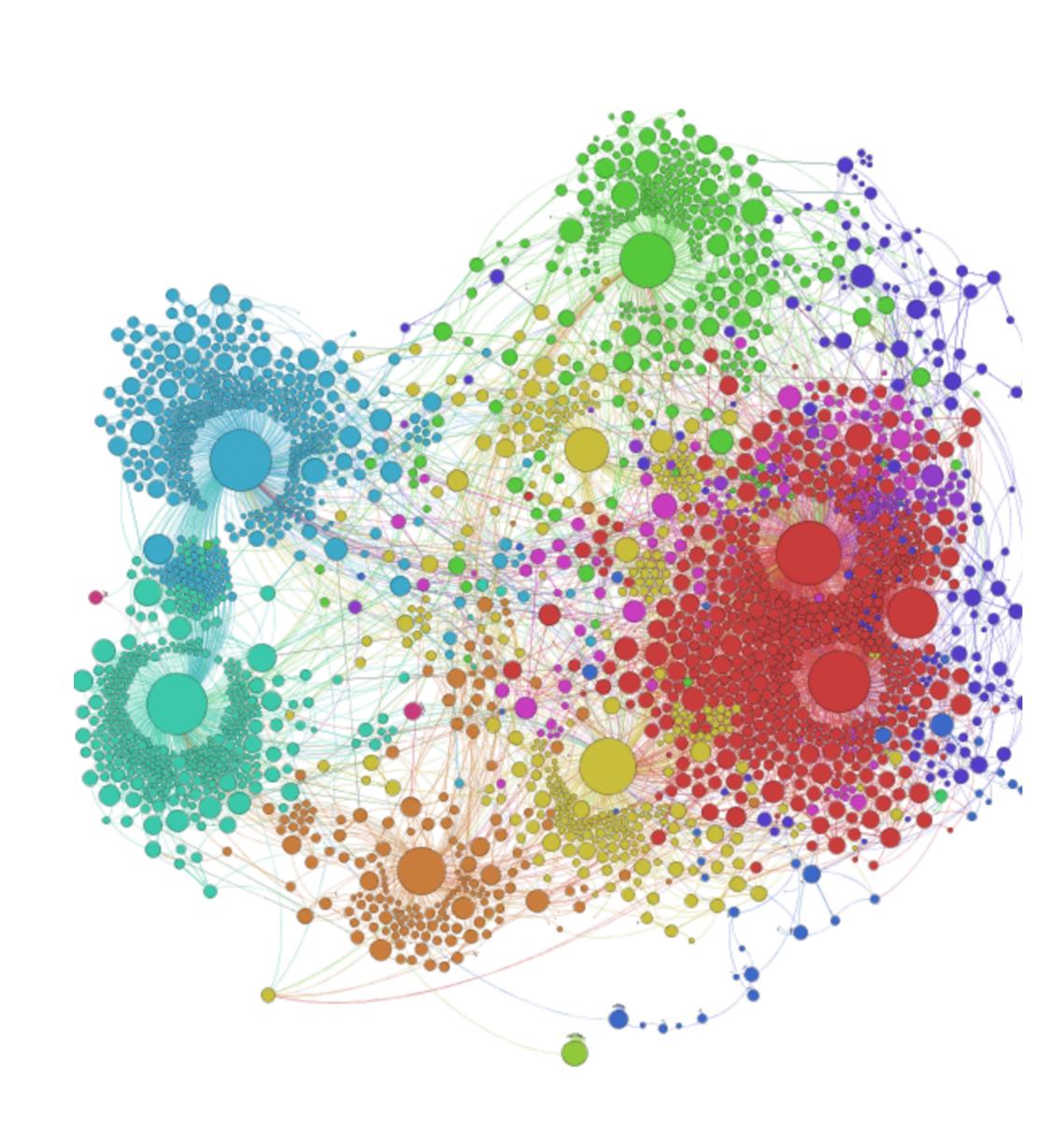
### It's hard to do more complex things with maps Is the spatial context paramoun Is the spatial context a proxy for something?

State	Est. pct. of votes	Reported margin	NYT projection	NYT w
New Mexico	100%	Clinton +8.2	Clinton +8.2	<b>√</b> V
Virginia	>95%	Clinton +4.8	Clinton +4.9	<b>√</b> V
Colorado	87%	Clinton +2.1	Clinton +3.7	<b>√</b> V
Maine	>95%	Clinton +2.7	Clinton +2.6	<b>√</b> V
Nevada	100%	Clinton +2.4	Clinton +2.4	<b>√</b> V
Minnesota	100%	Clinton +1.5	Clinton +1.5	<b>√</b> V
New Hampshire	100%	Clinton +0.2	Clinton +0.2	86%
Michigan	100%	Trump +0.2	Trump +0.3	929
Wisconsin	100%	Trump +0.9	Trump +0.9	<b>√</b> V
Pennsylvania	>95%	Trump +1.1	Trump +1.0	<b>√</b> V
Florida	100%	Trump +1.3	Trump +1.3	<b>√</b> V
North Carolina	100%	Trump +3.8	Trump +3.8	<b>√</b> V
Arizona	82%	Trump +4.3	Trump +4.5	>959
Georgia	100%	Trump +5.7	Trump +5.7	<b>√</b> V
Ohio	100%	Trump +8.5	Trump +8.5	<b>√</b> V



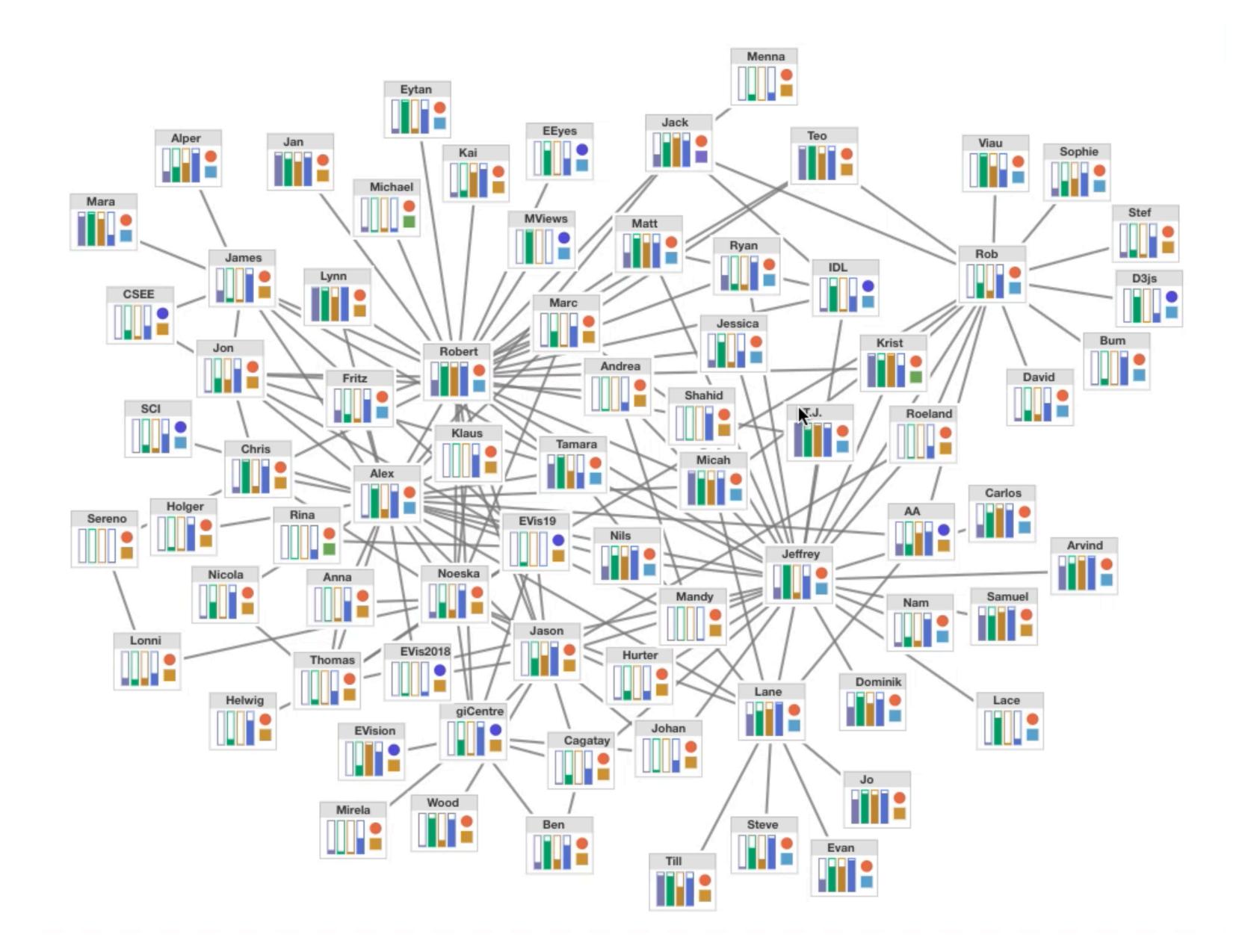
### ANALOGY: MULTIVARIATE NETWORKS

### 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



EUROVIS 2019 (Guest Editors)



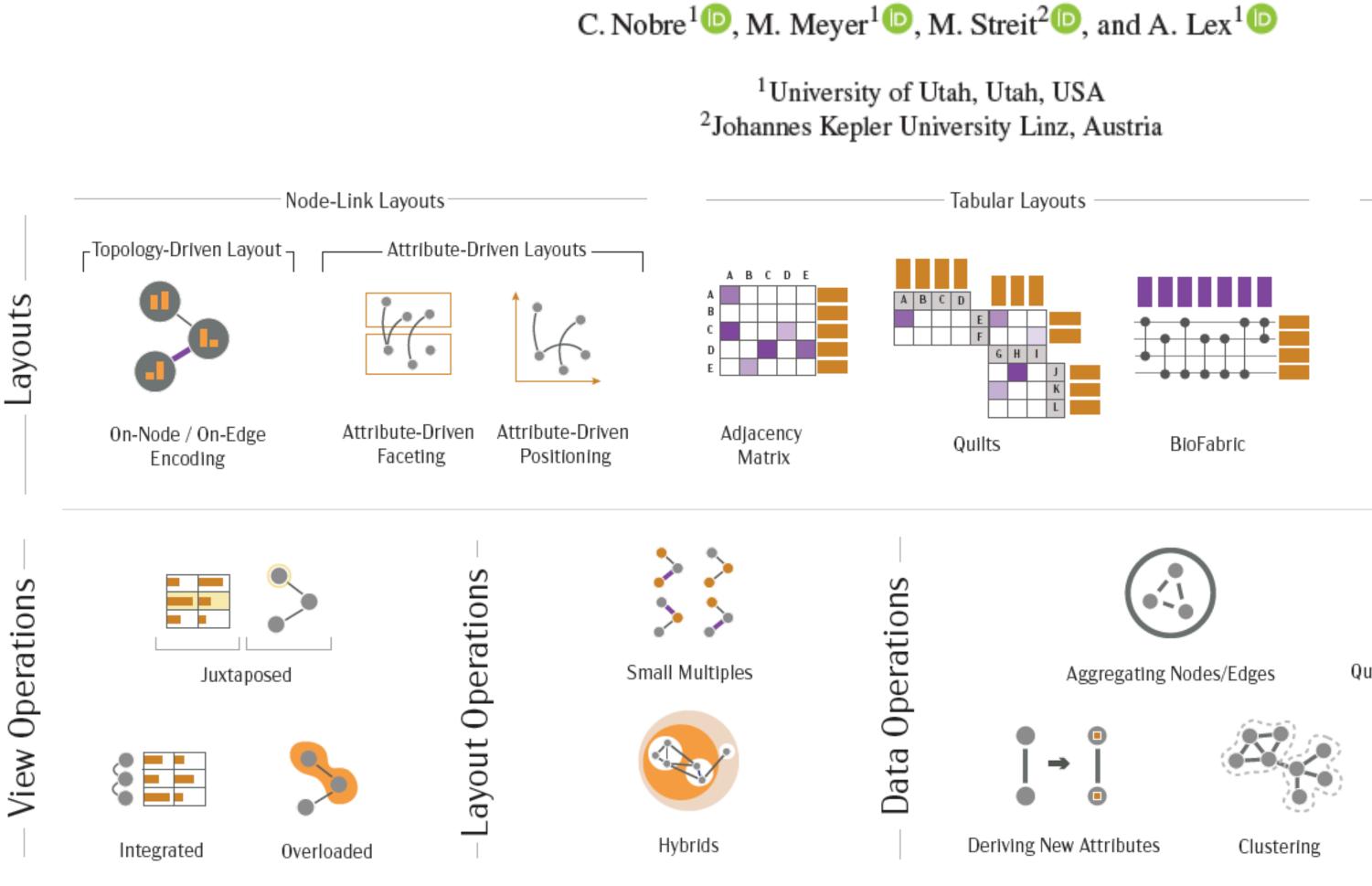


Figure 1: A typology of operations and layouts used in multivariate network visualization. Layouts describe the fundamental choices for encoding multivariate networks. View Operations capture how topology and attribute focused visualizations can be combined. Layout Operations are applied to basic layouts 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).

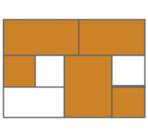
Abstract Multivariate networks are made up of nodes and their relationships (links), but also data about those nodes and links as attributes. Most real-world networks are associated with several attributes, and many analysis tasks depend on analyzing both, relationships and attributes. Visualization 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.

### The State of the Art in Visualizing Multivariate Networks

Volume 38 (2019), Number 3 STAR – State of The Art Report

Implicit Tree Layouts



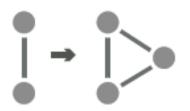


Inner Nodes + Leaves

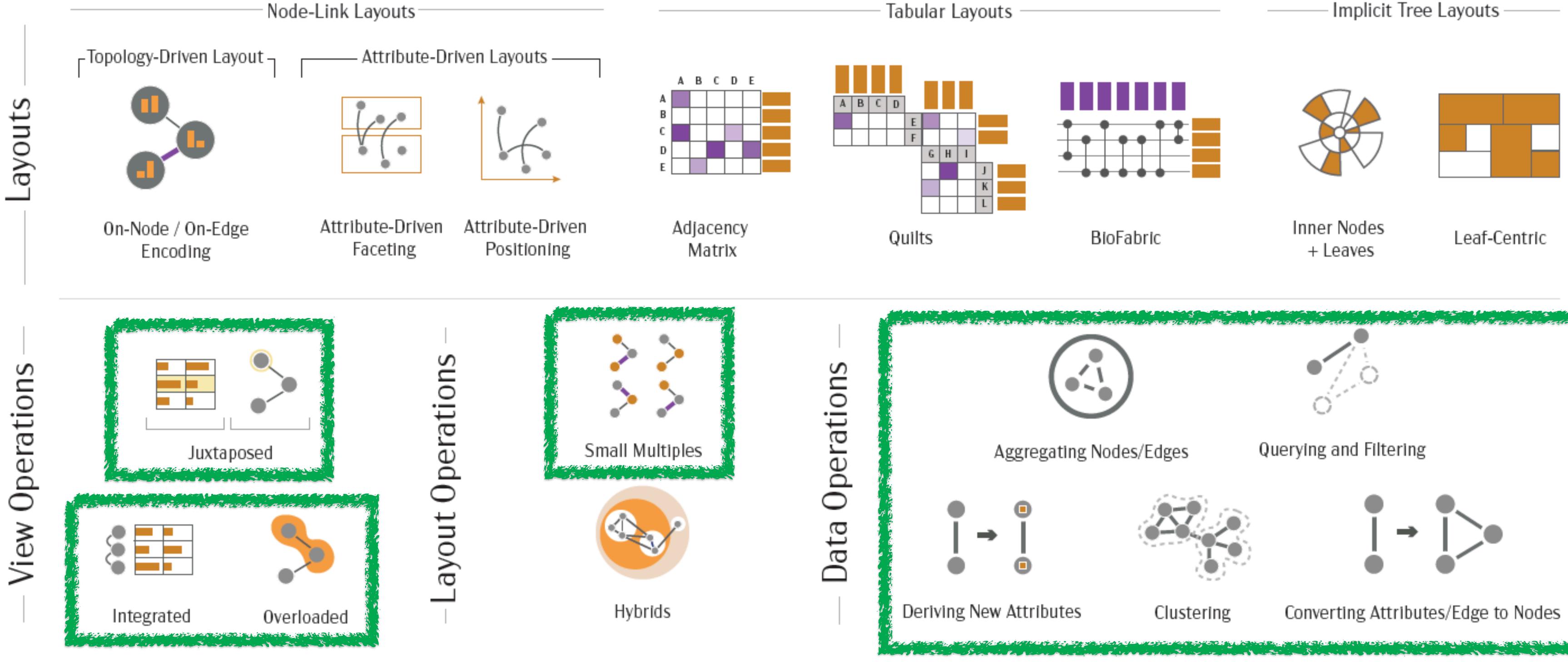
Leaf-Centric



Querying and Filtering



Converting Attributes/Edge to Nodes



Implicit Tree Layouts



### SO WHAT ELSE CAN You do?

### INTERACTION!

### Two Paths: 1. Select regions to show 2. Select / derive data to show

# SELECTING REGIONS TO SHOW

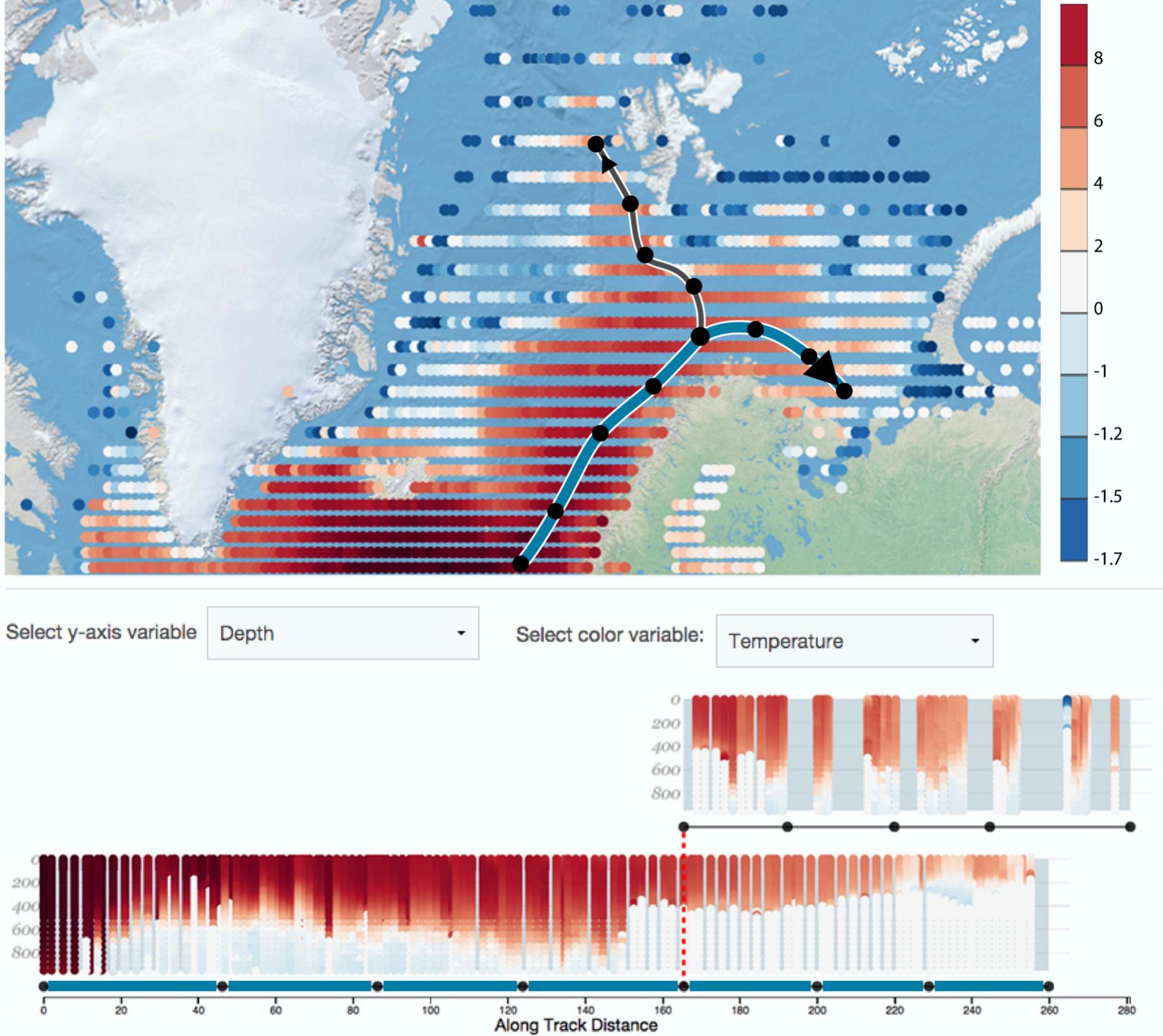
### PRINCIPLE

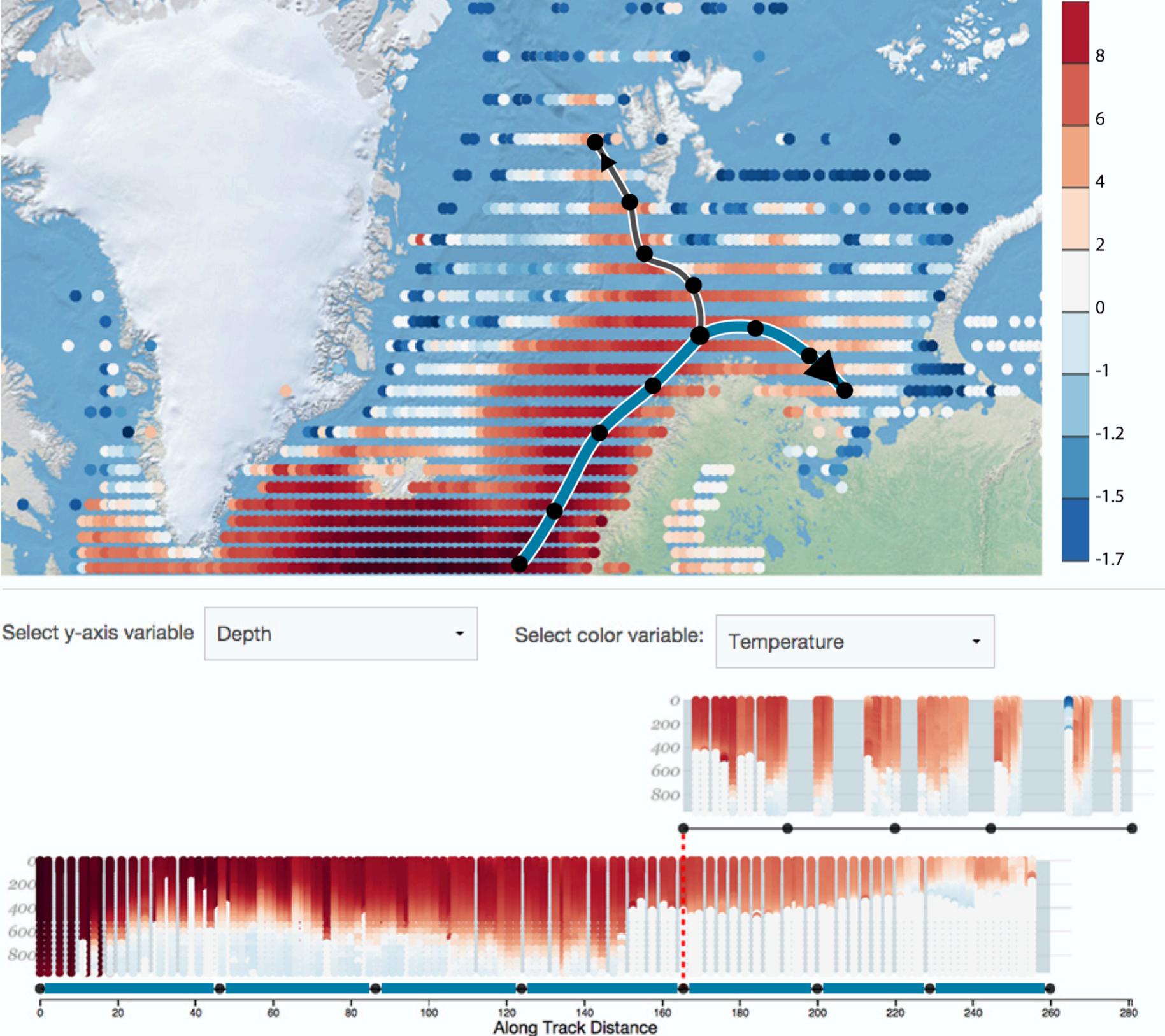
### Select one or multiple items/regions Show rich data about them in separate view



# SELECT A PATH IN A





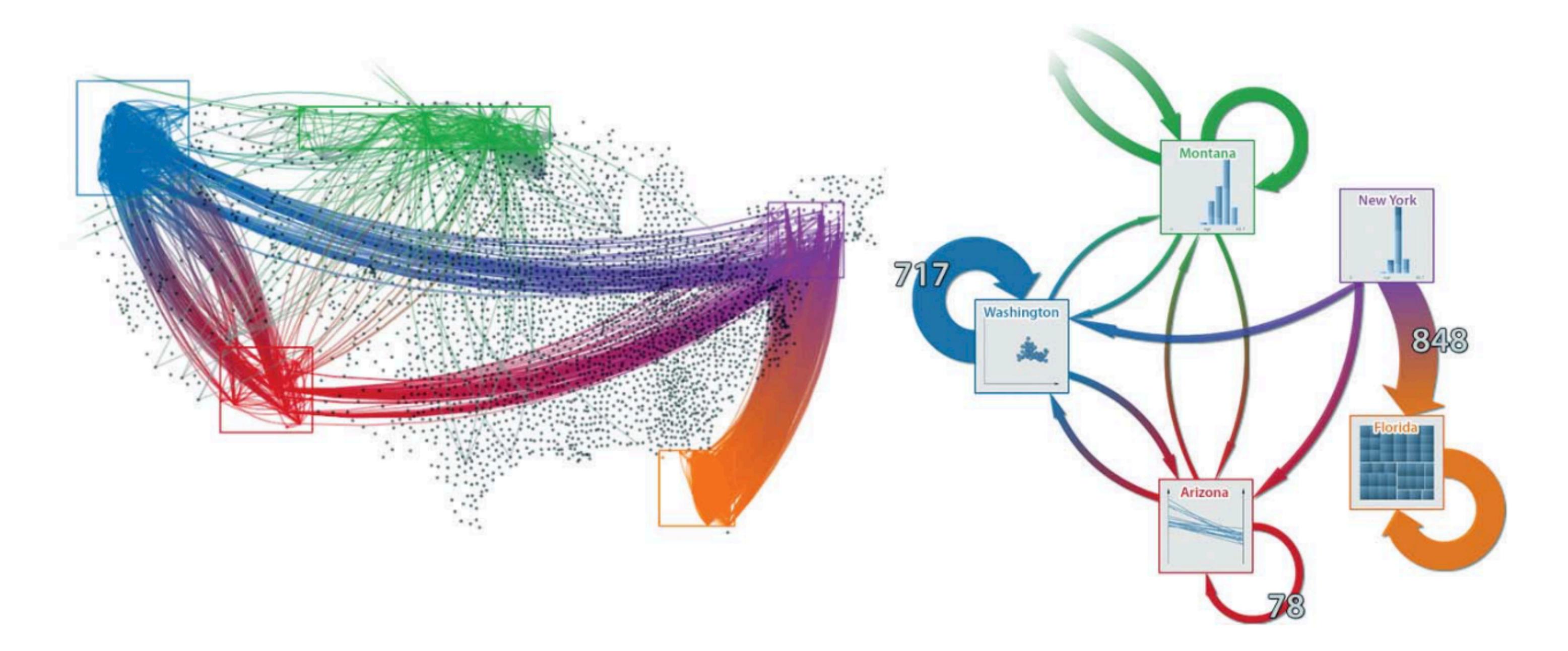


Carolina Nobre, Alexander Lex OceanPaths: Visualizing Multivariate Oceanography Data

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

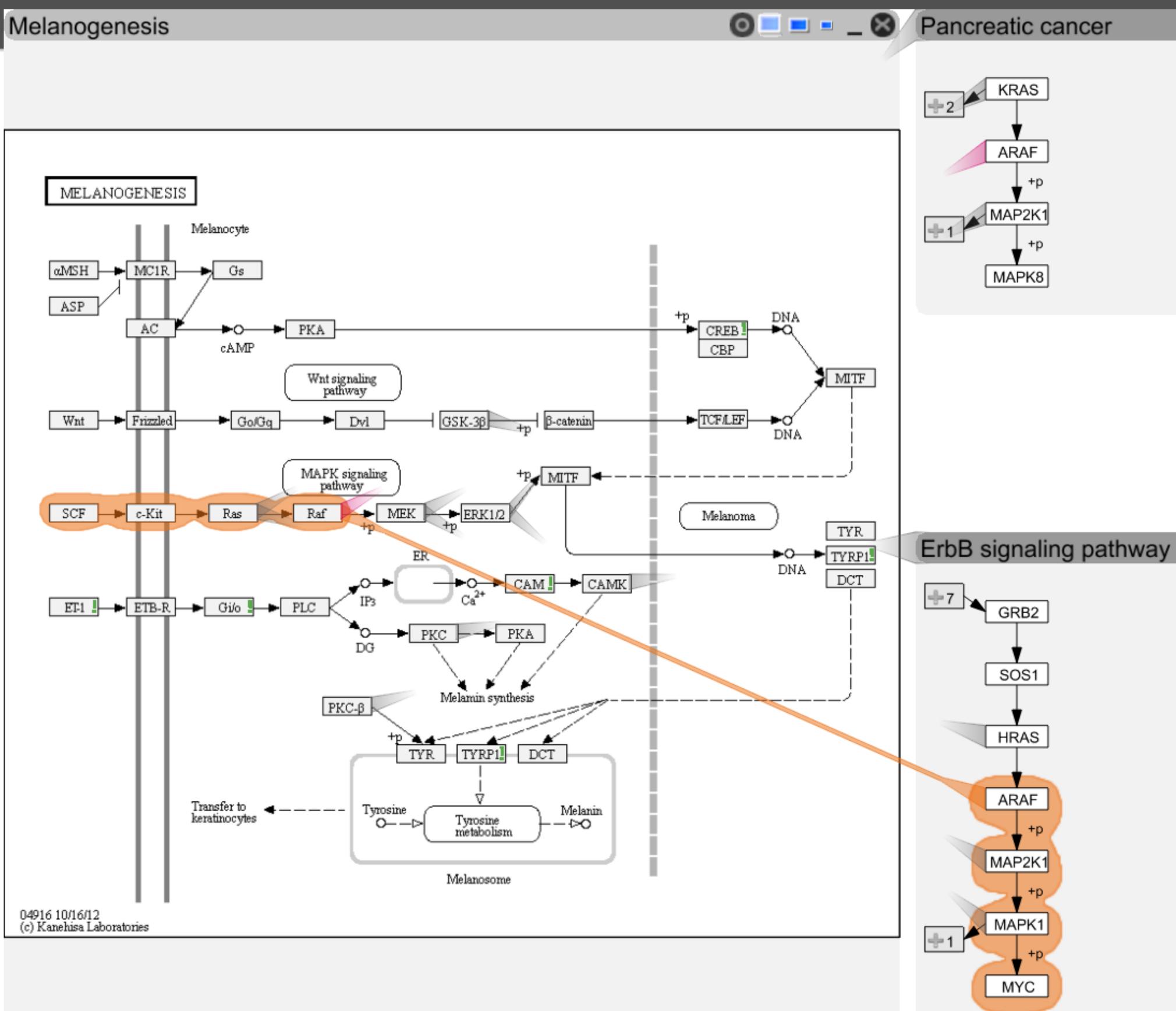


### SELECT A REGION AND AGGREGATE!

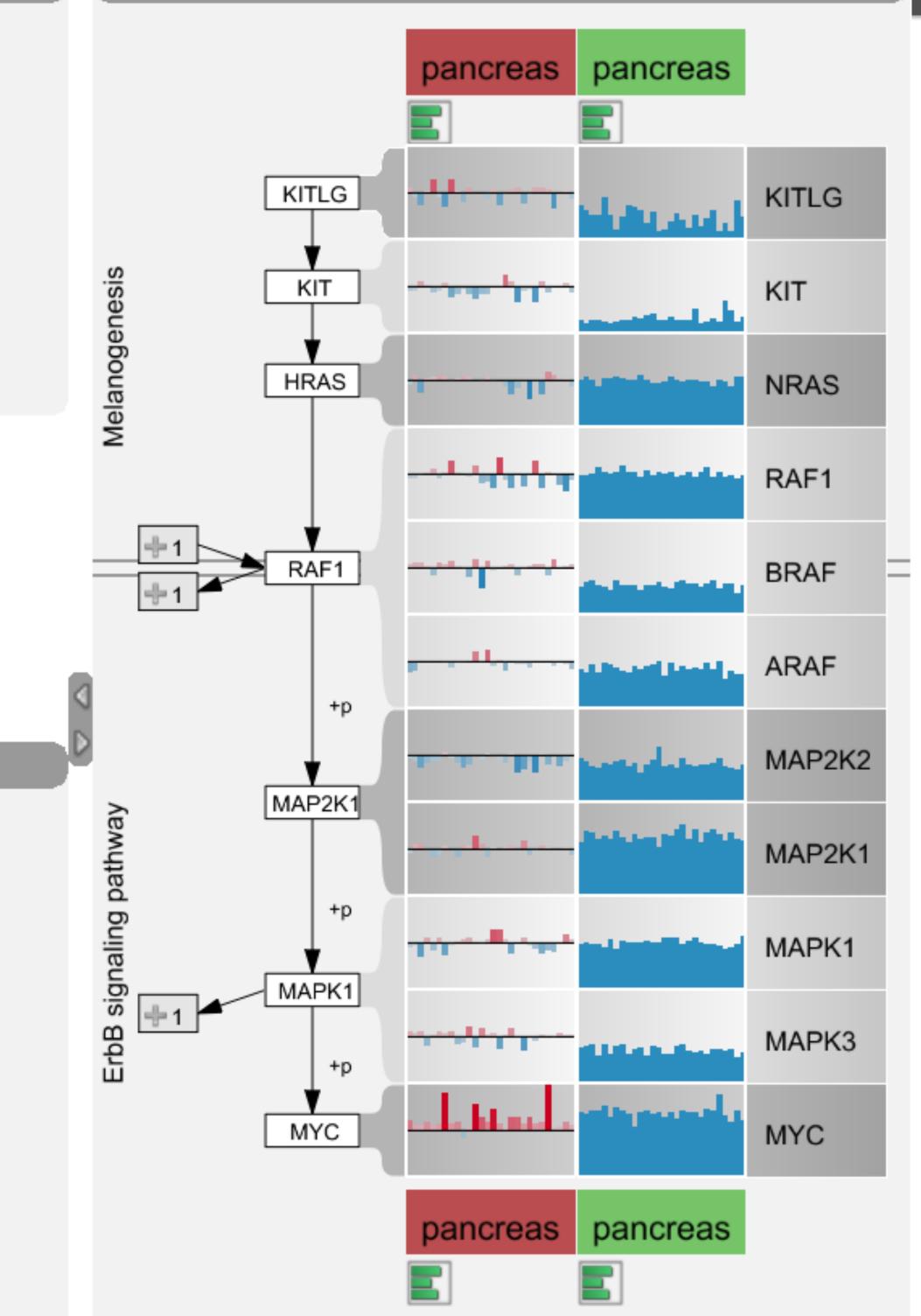


S. van den Elzen and J. J. van Wijk, "Multivariate Network Exploration and Presentation: From Detail to Overview via Selections and Aggregations," IEEE Transactions on Visualization and Computer Graphics (InfoVis '14), vol. 20, no. 12, pp. 2310-2319, 2014, doi: 10.1109/TVCG.2014.2346441.

### ENROUTE – PATH SELECTION



Selected Path



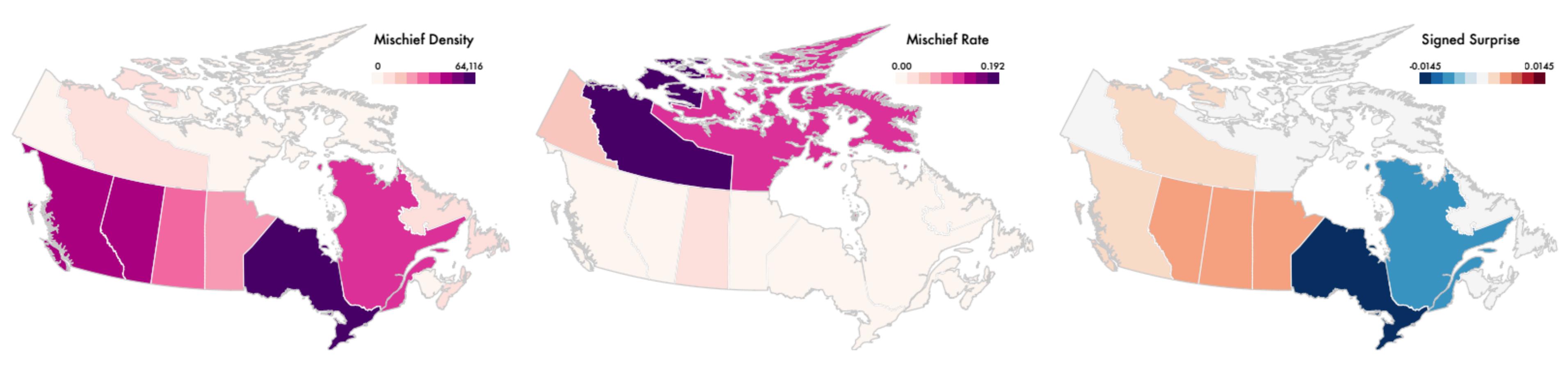
# SELECT / DERIVE DATA TOSHOW

### 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**

 $\bullet \bullet \bullet$ 

### APPROACH: USE A PRIOR, SHOW DIFFERENCE.



(a) The Event Density of "mischief" in Canada.

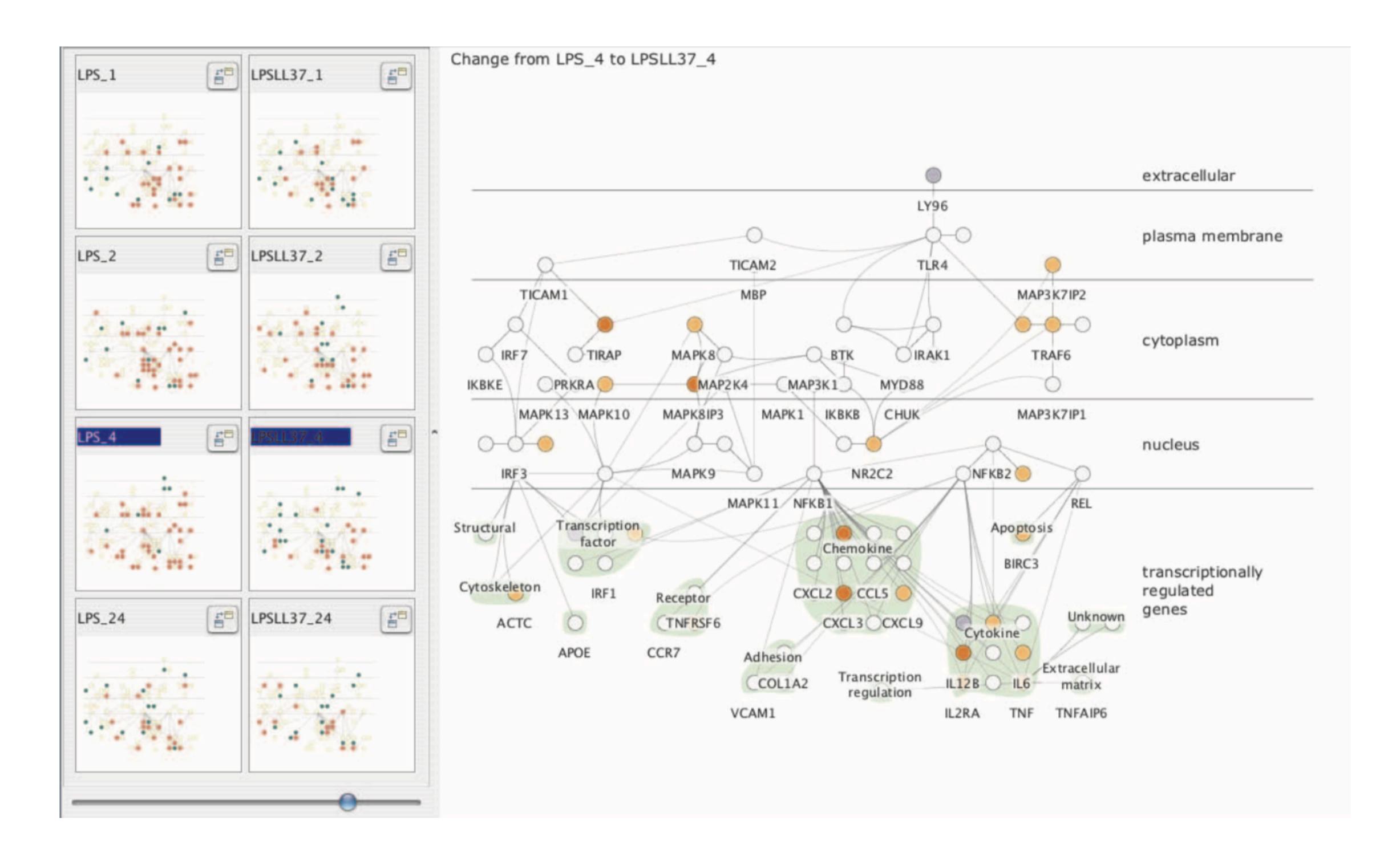
(b) The per-capita Event Rate of mischief.

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

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



### HAVE A HANDFUL OF SCORES? VIS FEASIBLE!





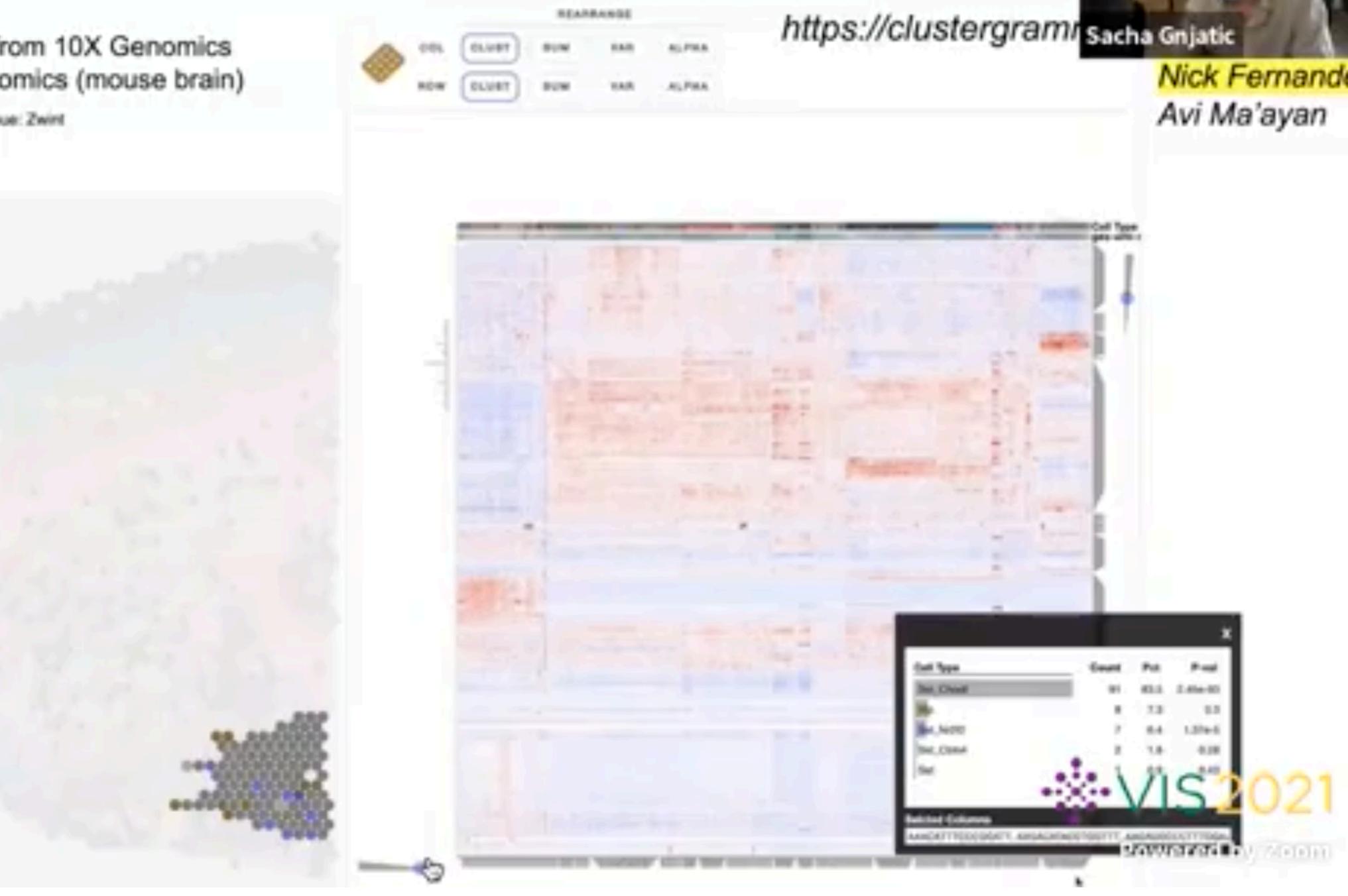


### **CLUSTERGRAMMER JUST EARLIER AT BIOVIS:** DYNAMIC FILTERS

### Clustergrammer: an interactive tool to visua.

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

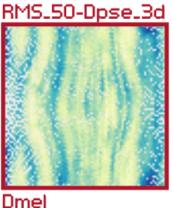
Visium Tissue: Zwint

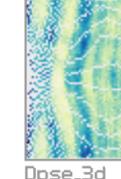


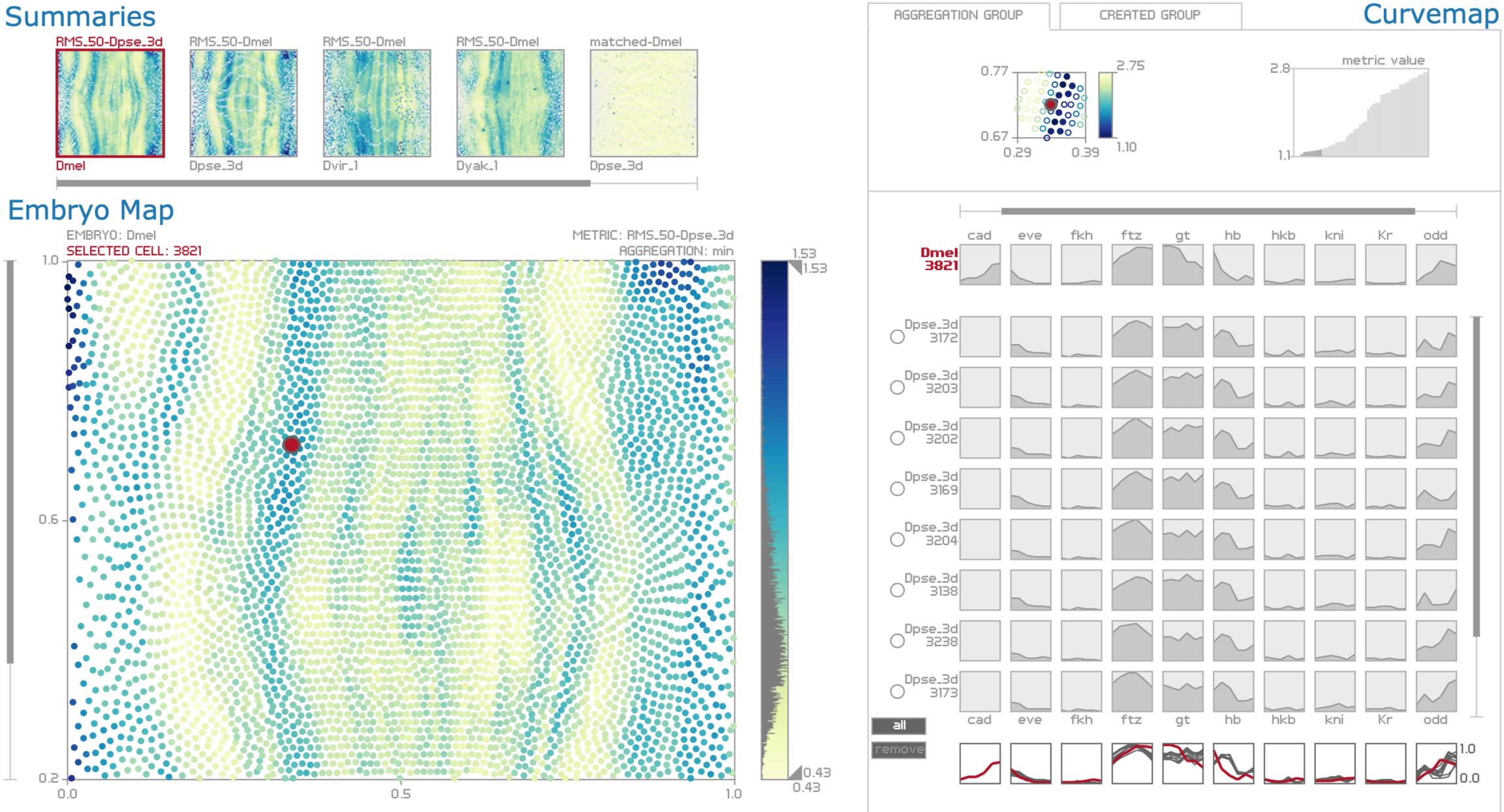


Nick Fernandez

### **EXAMPLE: DOMAIN SPECIFIC SCORE**



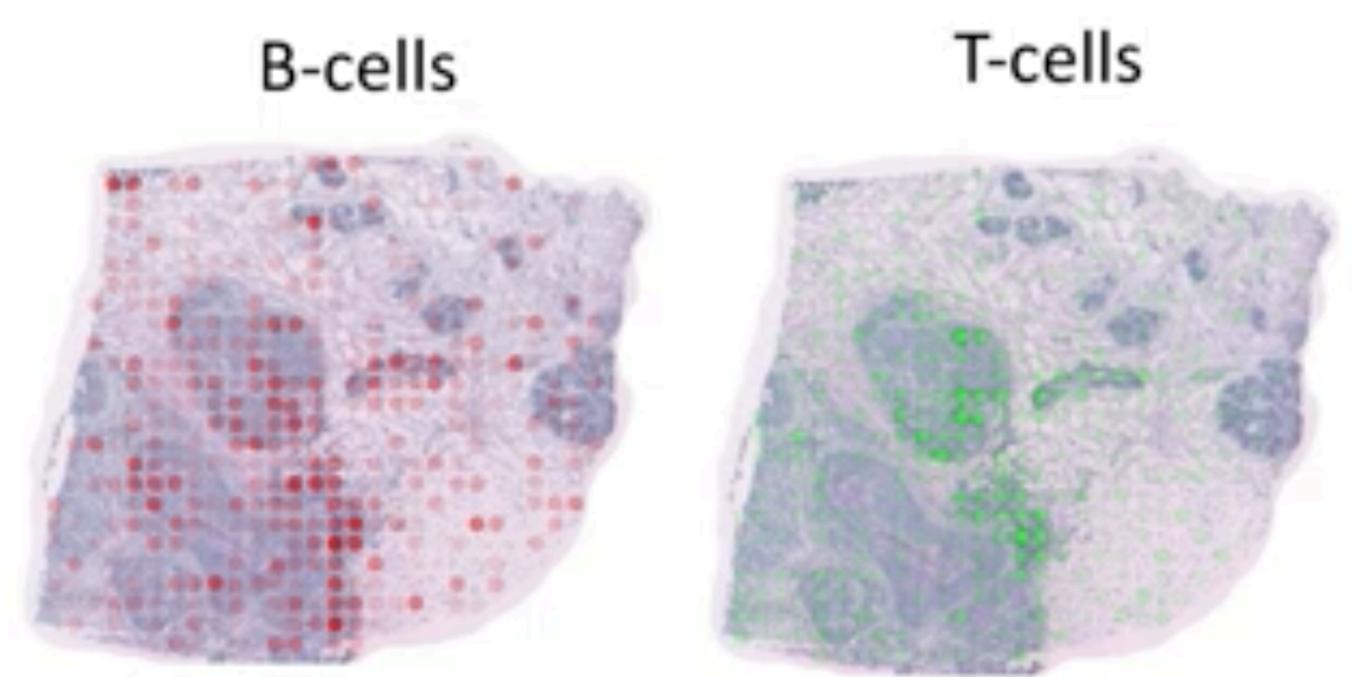




Miriah Meyer, Tamara Munzner, Angela DePace, Hanspeter Pfister MulteeSum: A Tool for Comparative Spatial and Temporal Gene Expression Data IEEE Transactions on Visualization and Computer Graphics (InfoVis), 16(6): 908--917, doi:10.1109/TVCG.2010.137, 2010.

# USING SCORES @ BIOVIS BY ALMA ANDERSON

### Assessing co-localization



Sample from breast cancer patient (HER2-positive)

- .

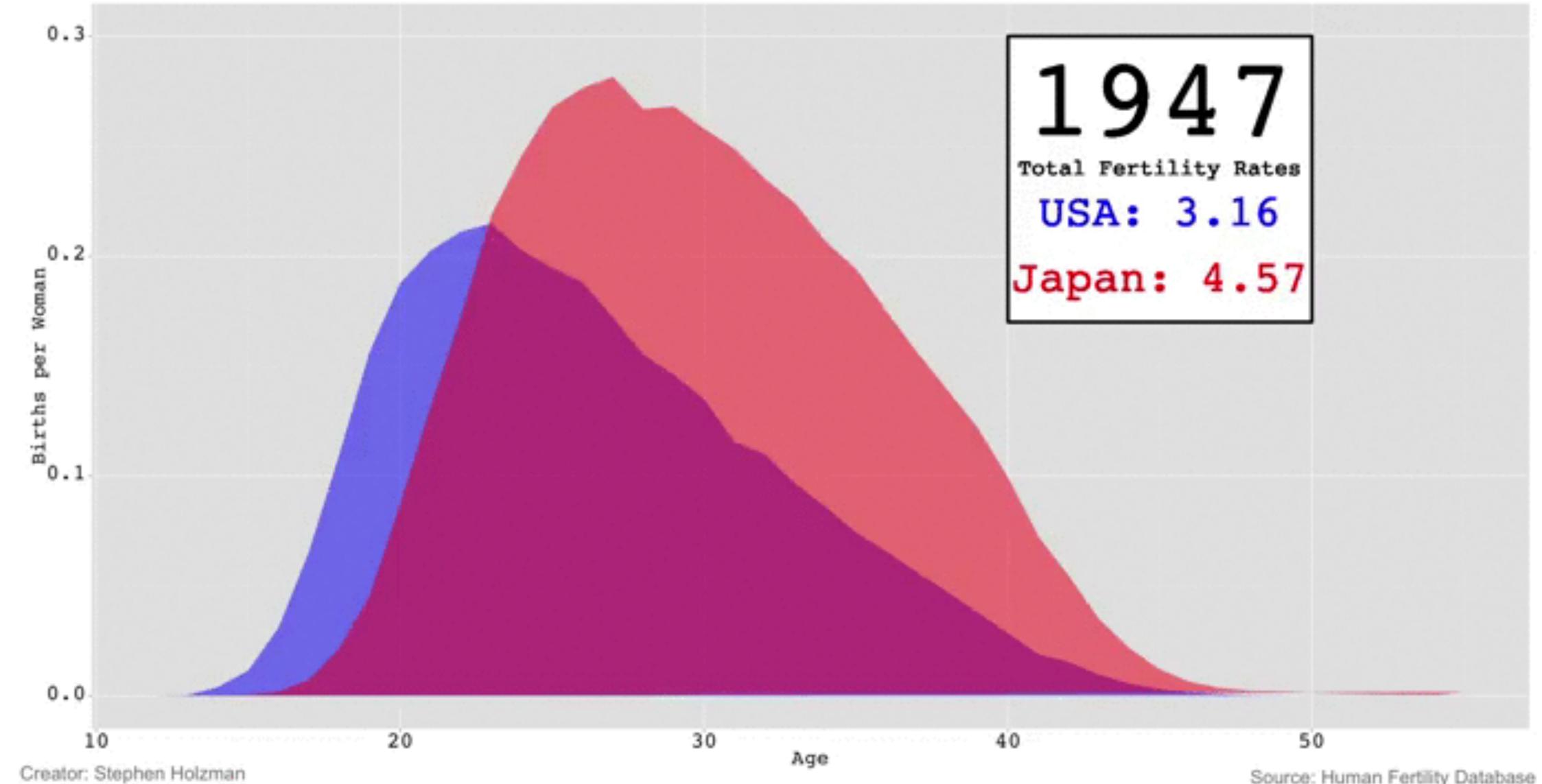
We were interested in TLS-sites enriched for both B and T-cells; these have relevance for prognosis and understanding of the cancer

Our immunologist wanted to relate this co-localization to the morphology

Solution: transform to single metric (co-localization score) and show



### REDESIGN EXAMPLE

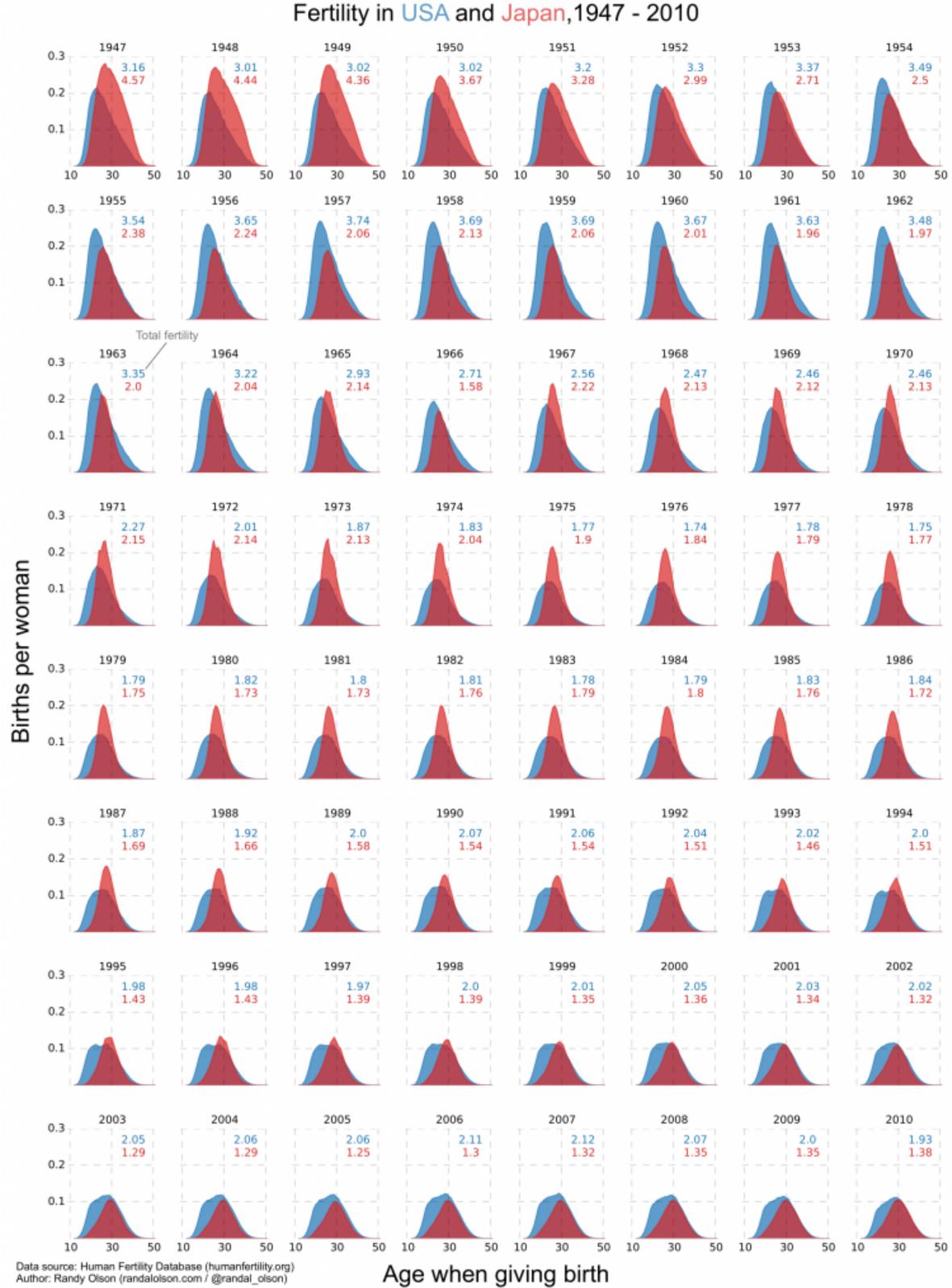


### USA and Japan Fertility Over Time

Source: Human Fertility Database

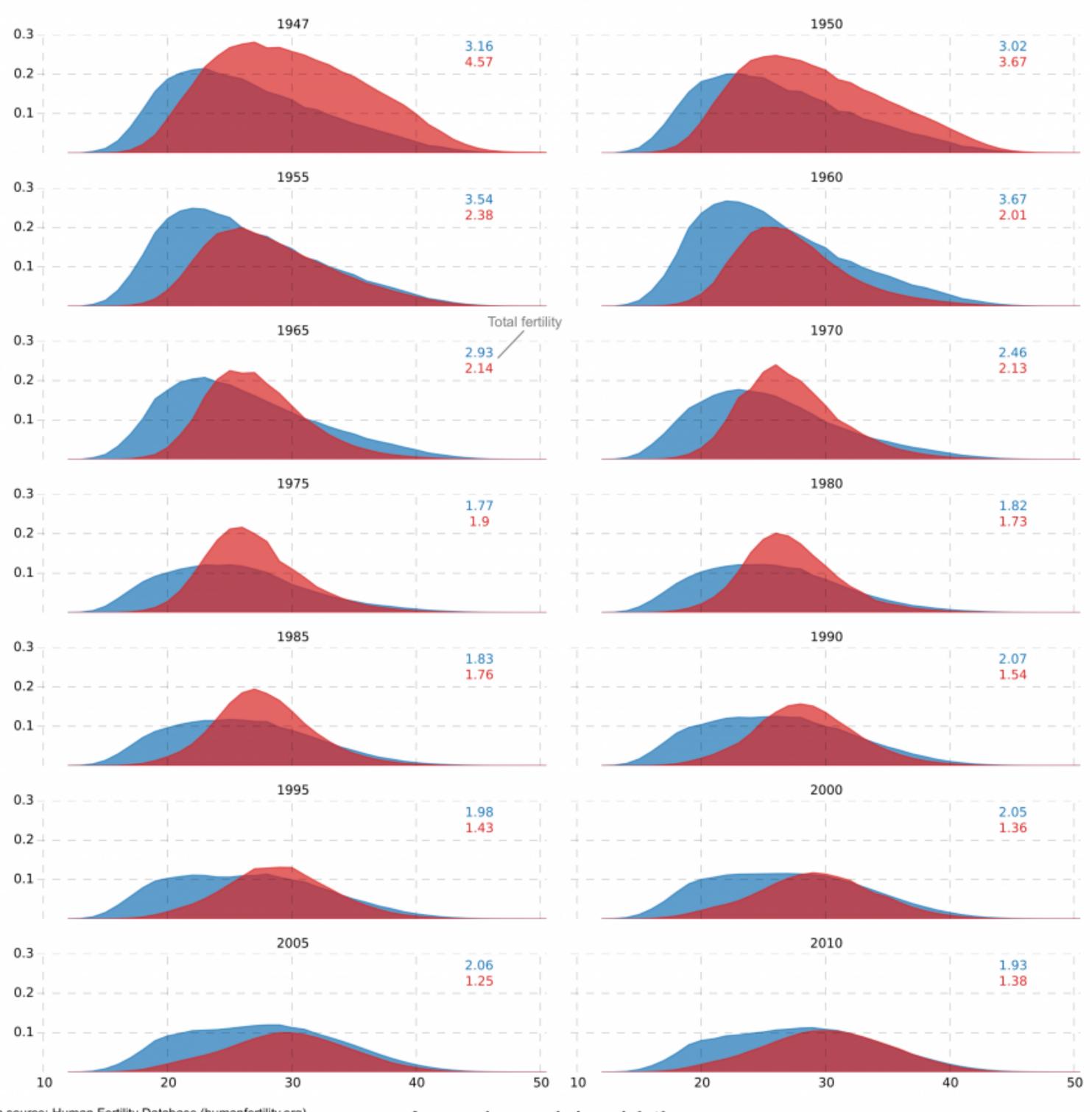


# EYES BEAT MEMORY: SMALL MULTIPLES



### EYES BEAT MEMORY: SMALL MULTIPLES

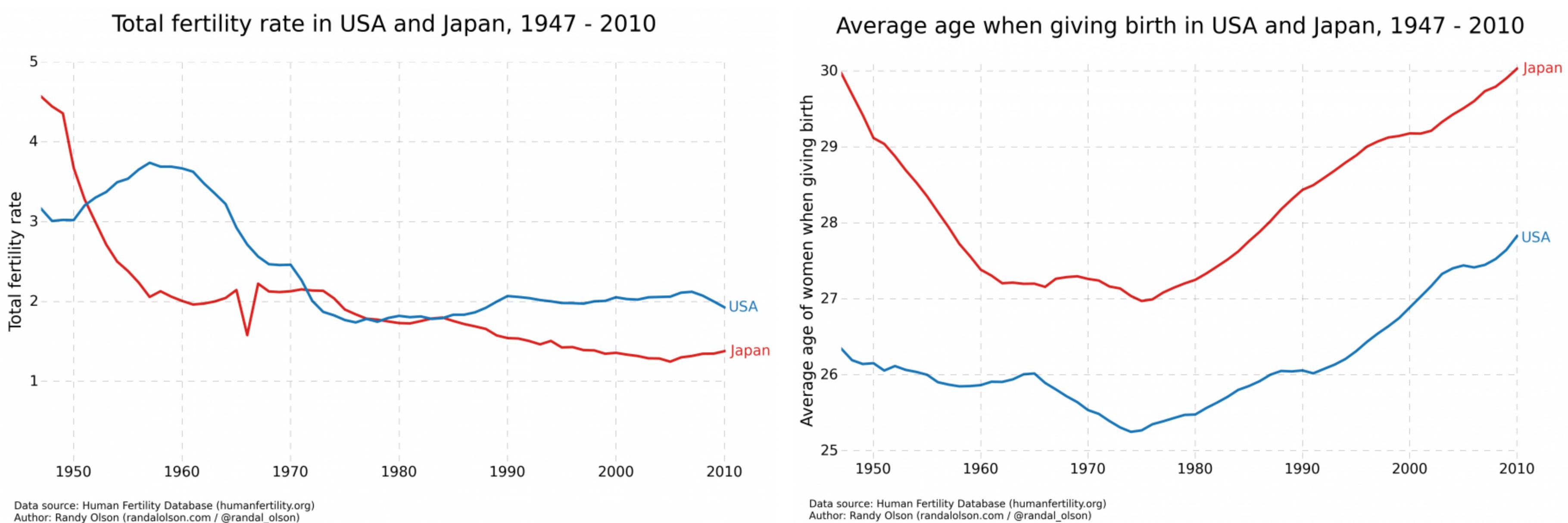
Fertility in USA and Japan, 1947 - 2010



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

Age when giving birth

### SIMPLIFY



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!



# VISUAIZATION design lab

### THE UNIVERSITY OF UTAH



