

# HuBMAP CCF User Interfaces

**Katy Börner and the MC-IU HIVE HuBMAP Team**

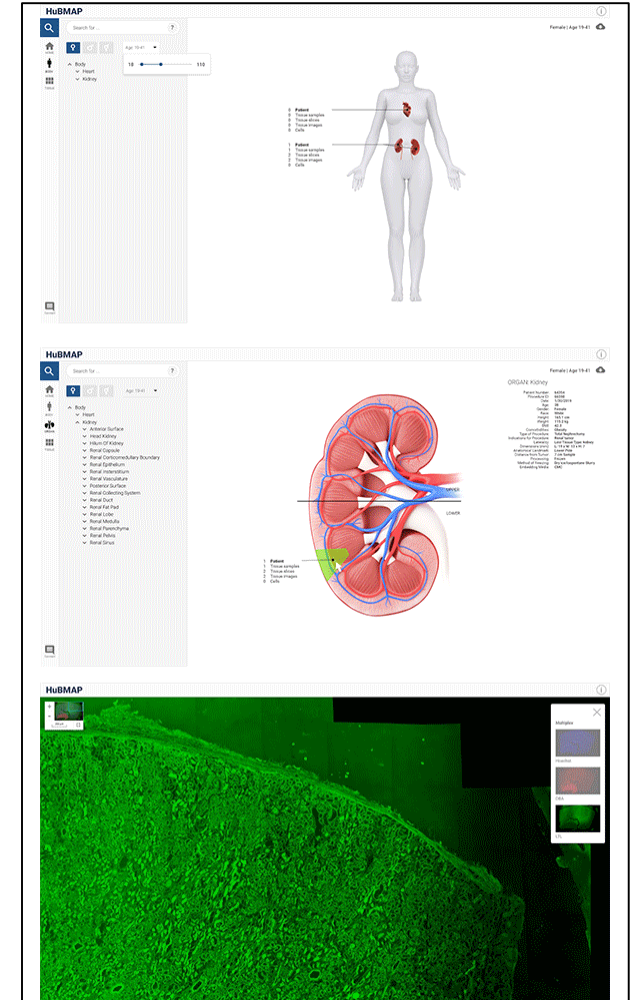
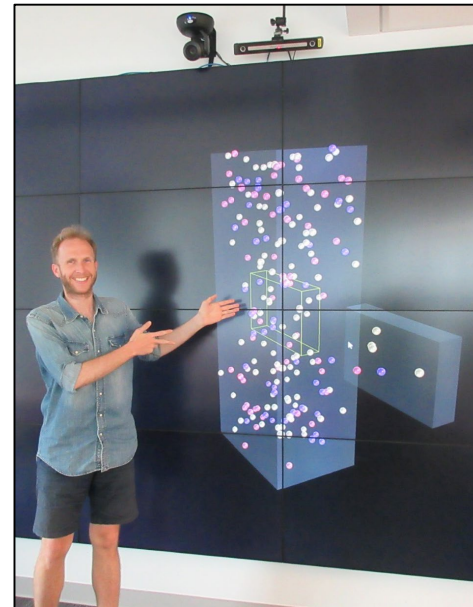
Intelligent Systems Engineering, SICE

Indiana University, Bloomington, IN

HuBMAP CCF WS

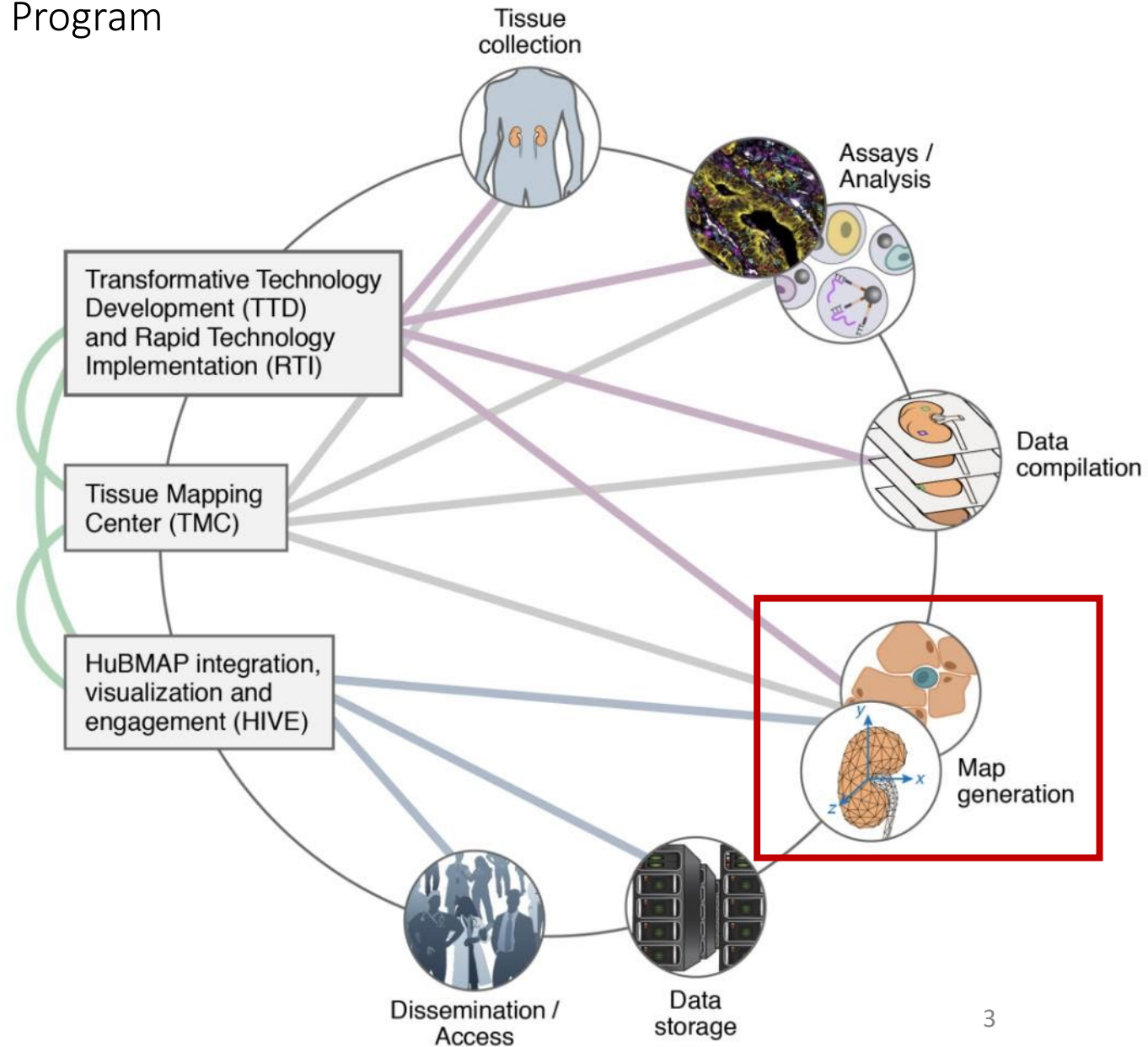
Harvard Medical School, Boston, MA

*June 27, 2019*

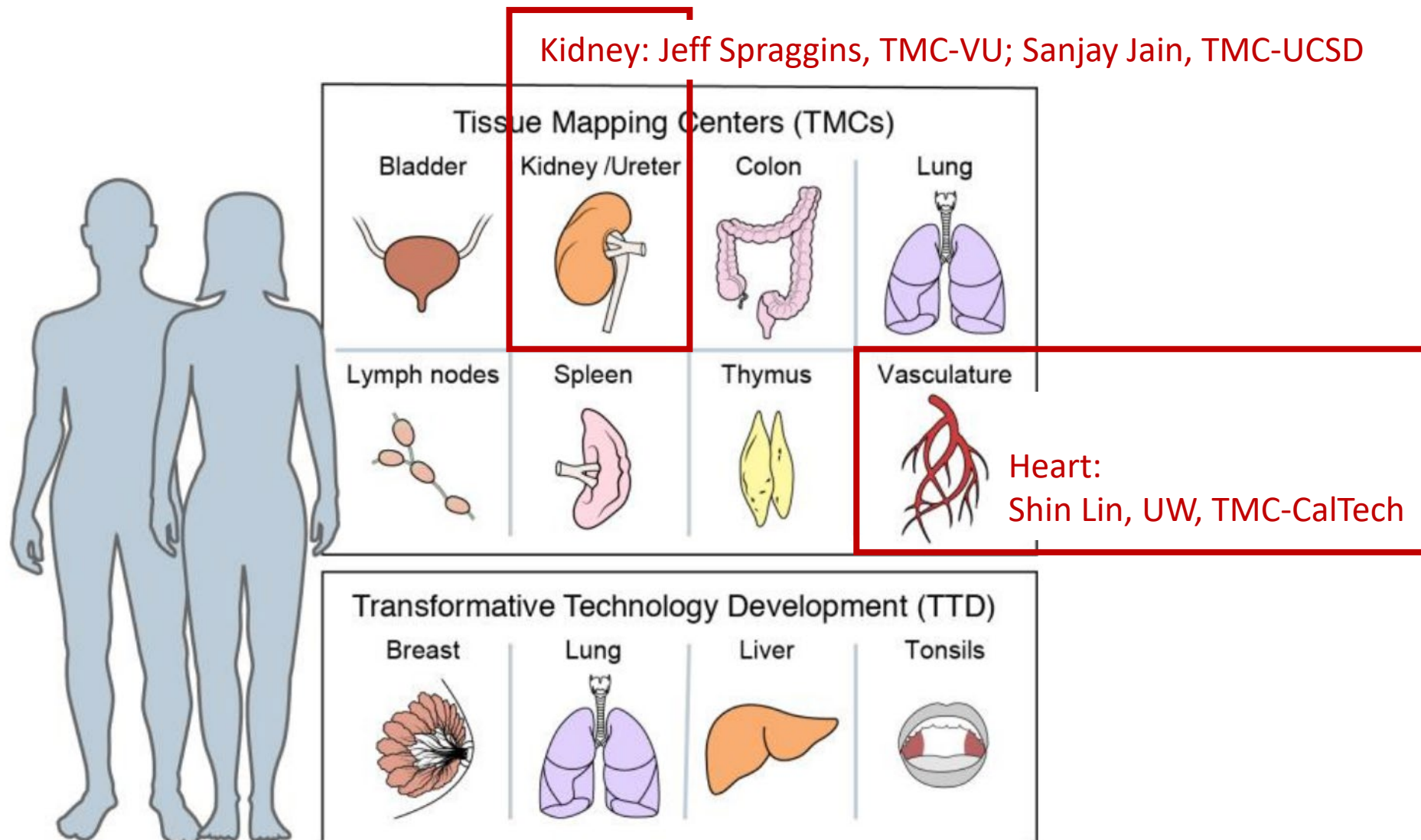


# HuBMAP CCF Data & User Interfaces

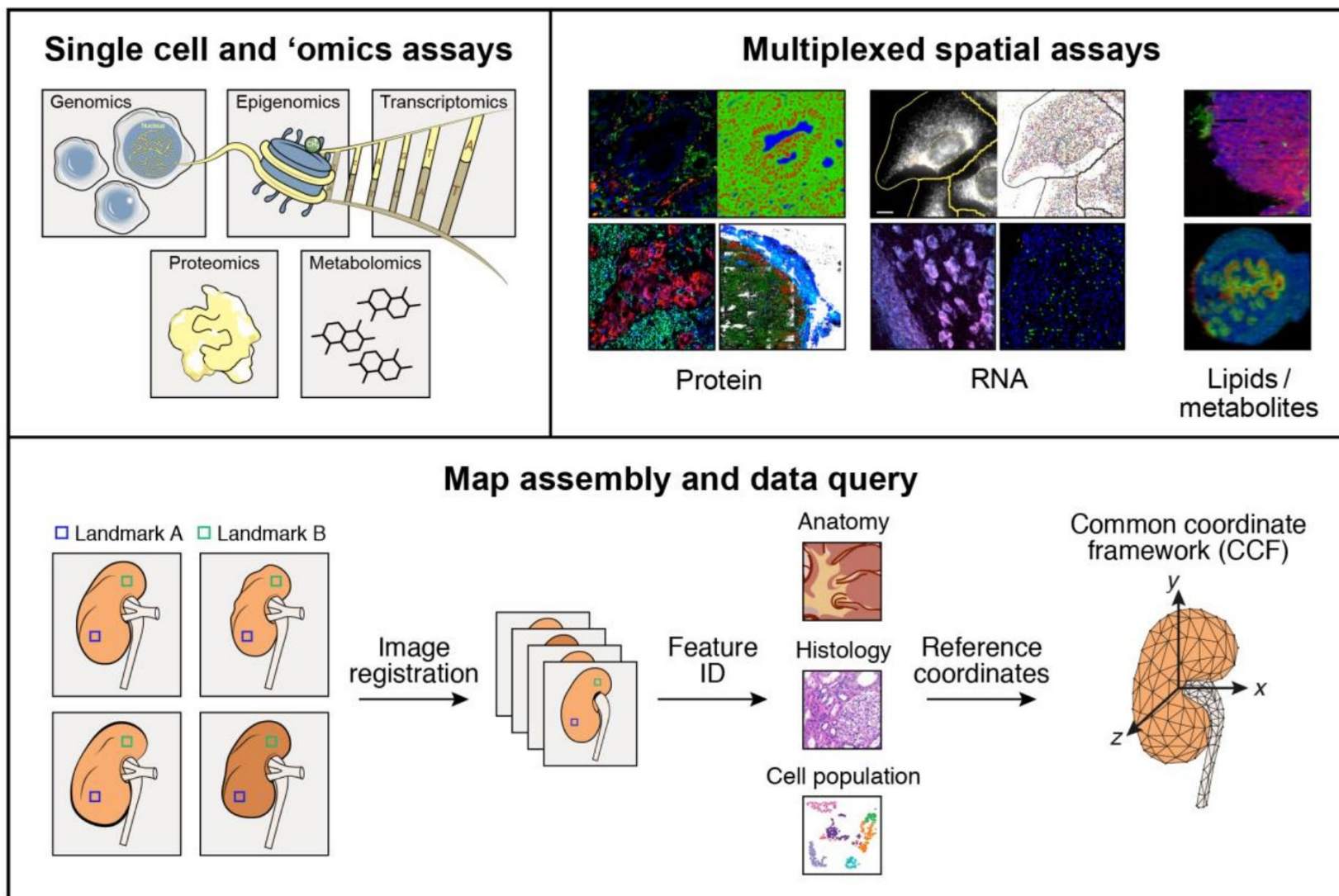
Mapping the Human Body at Cellular Resolution—  
The NIH Common Fund Human BioMolecular Atlas Program  
Snyder et al. <https://arxiv.org/abs/1903.07231>

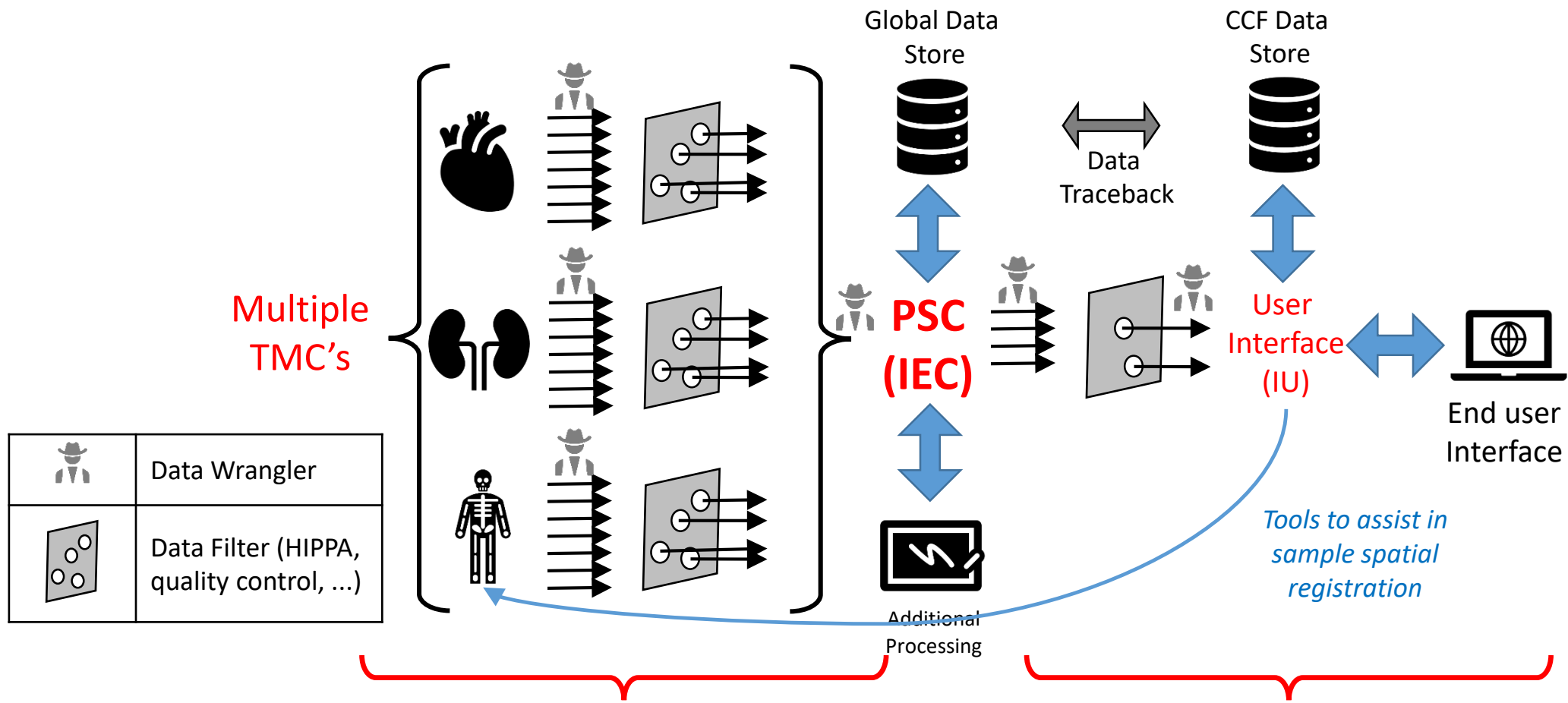


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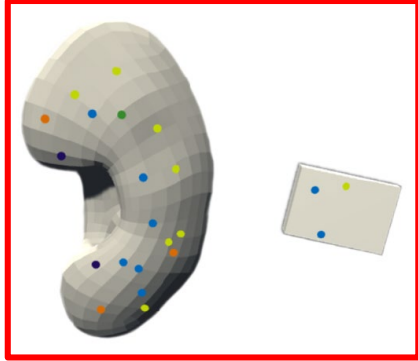


- Provenance
- Patient
- Sample
- Sample Processing
- Technology (MS, IH, ...)
- Analysis
- Etc.

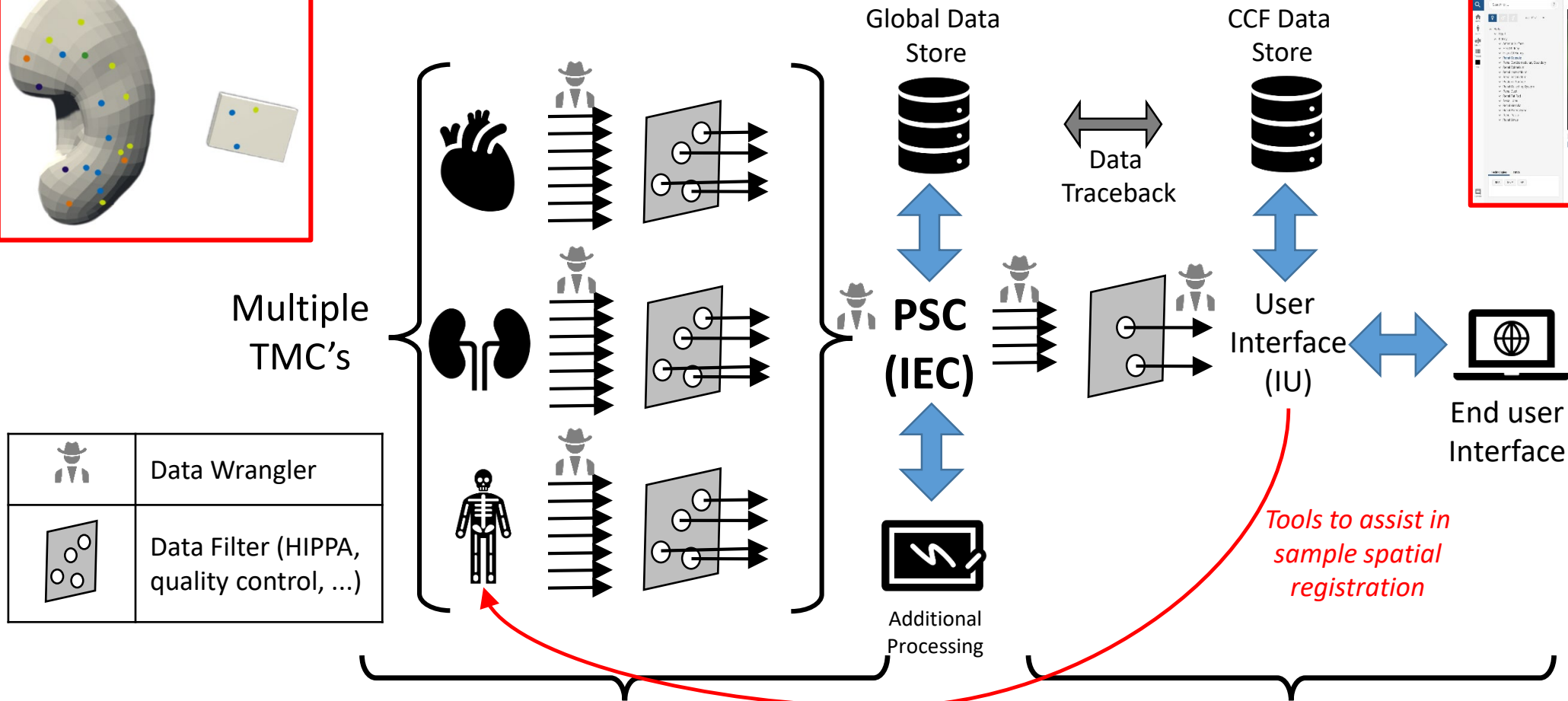
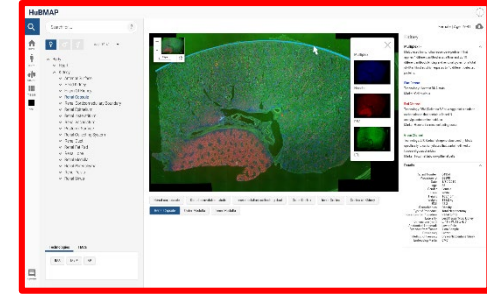
- Only the data needed for the GUI

TMC: Tissue Mapping Center  
PSC: Pittsburgh Supercomputing Center

## Tissue Registration UI



## CCF User Interface (UI)



	Data Wrangler
	Data Filter (HIPPA, quality control, ...)

- Provenance
- Patient
- Sample
- Sample Processing
- Technology (MS, IH, ...)
- Analysis
- Etc.

*Propagate needs back to TMC's*

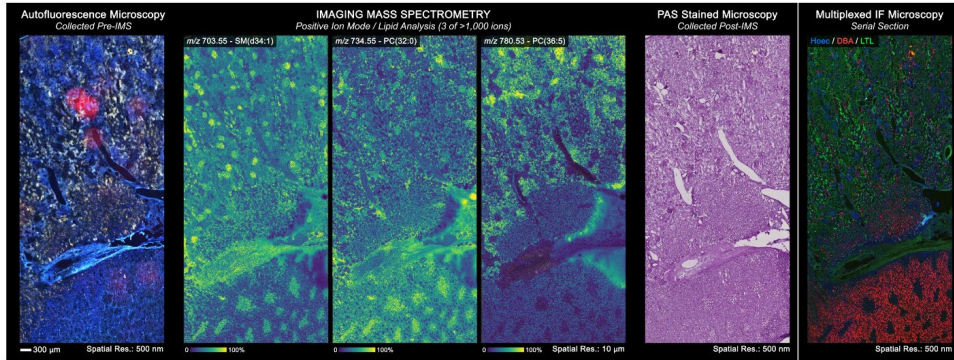
- Only the data needed for the GUI

TMC: Tissue Mapping Center  
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# Data. Is very heterogeneous. Must provide guidance.

**Kidney: Jeff Spraggins et al., VU**

See data on Globus, BIOMIC\_patient-64354



Clinical and Spatial Metadata (21 rows)

Cell type	Subset A	Subset B	Subset C
Tubular Epithelium	Proximal tubular cells	S1	
		S2	
		S3	
Loop on Henle		Thin descending limb	
		Thin ascending limb	
		Thick limb	medullary cortical
		Macula Densa	
Distal convoluted tubule			
Connecting segment			
Collecting duct		Principal cells	
		Intercalated cells	Type A
Glomerulus	Epithelium	Visceral	Type B
		Parietal	
Vasculature	Mesangial cells	Glomerular	
		Peritubular	
		Lymphatic	
	Pericytes		
		Juxta Glomerular Cells	
Interstitial	Fibroblasts	Myofibroblasts	
		EPO producing cells	
		Medullary fibroblasts	
	Mononuclear cells	Resident macrophages	
		Dendritic cells	
	Lymphocytes	T cells	
		B cells	
		NK cells	

Cell Types, on right

Cell States (9 rows)

Cell states	Subset A
Proliferating cells	S-phase
	G2/M
Cell cycle arrest	G0
	G1/S
	G2/M

**Heart: Shin Lin, UW**

Year 1: Tissue data for 1-2cm cubed volumes from 9 sites for 1 heart from 1 individual.

Data Dictionary (115 rows)

Field #	Sort	Field Label	Sort	Field Name	Sort	Field Units	Field Data	Lookup	Tal	Low Value	High Value	Valid value	IsNull	Parent Field	Parent Field	Can Child	Read Only	Sort	
9	Donor	//ABO:	abo				char(3)	lkup_abo					TRUE					FALSE	
10	Donor	//Date of birth:	dob				datetime						TRUE					FALSE	
11	Donor	//Gender:	gender				char(1)	lkup_gender				M,F	TRUE					FALSE	
12	Details	//Age:	age_in_months				smallint			0	1188		TRUE			FALSE		FALSE	
13	Details	//Age Unit:	age_unit				char(1)	lkup_age_unit				M,Y	TRUE	age_in_months				TRUE	
14	Details	//Height:	hgt_cm			cm	decimal(5, 2)			1	241.3		TRUE					FALSE	
15	Donor	hgt_ft //	hgt_ft			ft	int			0	7		TRUE					TRUE	
16	Donor	hgt_in //	hgt_in			in	int			0	11		TRUE					TRUE	
17	Details	//Weight:	wgt_kg			kg	decimal(7, 4)			0.454	294.835		TRUE					FALSE	
18	Donor	wgt_lb //	wgt_lb			lbs	decimal(3, 0)			2	650		TRUE					TRUE	
19	Donor	//Ethnicity/race:	race				bigint	lkup_race_subcat_multi					FALSE					FALSE	
30	Details	//History of diabetes:	hist_diabetes				smallint	lkup_histdiab_dur					TRUE					FALSE	
31	Donor	//History of cancer:	hist_cancer				smallint	lkup_histcancer_site					TRUE			FALSE		FALSE	
32	Donor	History of cancer:	cancer_oth_ostxt				varchar(50)			1	50		TRUE	hist_cancer	999			FALSE	FALSE
33	Details	//History of hypertension:	hist_hypertension				smallint	lkup_histhype_dur					TRUE			FALSE		FALSE	

Cell Types (14)

endothelial cells	
	arterial
	capillary
	venous
	lymphatic
cardiomyocytes	
	atrial
	ventricular
	nodal
fibroblasts	
	fibroblasts
	myofibroblasts
immune cells	
	macrophages



# Data: Clinical

## Kidney: Jeff Spraggins et al., VU

### Clinical and Spatial Metadata (21 rows)

Sample Number:	20
Patient Number:	64354
Procedure ID:	66598
Date:	1/30/2019
Age:	38
Gender:	Female
Race:	White
Height:	165.1 cm
Weight:	115.2 kg
BMI:	42.3
Comorbidities:	Obesity
Type of Procedure:	Total Nephrectomy
Indications for Procedure:	Renal tumor
Laterality:	Left
Tissue Type:	kidney
Dimensions (mm):	L: 19 x W: 13 x H: 7
Anatomical Landmark:	Lower Pole
Distance from Tumor:	7 cm
Sample Processing:	Frozen
Method of Freezing:	Dry Ice/Isopentane Slurry
Embedding Media:	CMC

## Heart: Shin Lin, UW

### Data Dictionary (115 rows)

Field #	Sort	Field Label	Sort	Field Name	Sort	Field Units	Field Data	Lookup Table	Low Value	High Value	Valid value
9	Donor	//ABO:		abo			char(3)	lkup_abo			
10	Donor	//Date of birth:		dob			datetime				
11	Donor	//Gender:		gender			char(1)	lkup_gender			M,F
12	Details	//Age:		age_in_months			smallint		0	1188	
13	Details	//Age Unit:		age_unit			char(1)	lkup_age_unit			M,Y
14	Details	//Height:		hgt_cm		cm	decimal(5, 2)		1	241.3	
15	Donor	hgt_ft //		hgt_ft		ft	int		0	7	
16	Donor	hgt_in //		hgt_in		in	int		0	11	
17	Details	//Weight:		wgt_kg		kg	decimal(7, 4)		0.454	294.835	
18	Donor	wgt_lb //		wgt_lb		lbs	decimal(3, 0)		2	650	
19	Donor	//Ethnicity/race:		race			bigint	lkup_race_subcat_multi			
30	Details	//History of diabe	hist_diabetes				smallint	lkup_histdiab_dur			
31	Donor	//History of cance	hist_cancer				smallint	lkup_histcancer_site			
32	Donor	History of cancer	, cancer_oth_ostxt				varchar(50)		1	50	
33	Details	//History of hyper	hypertension				smallint	lkup_histhype_dur			

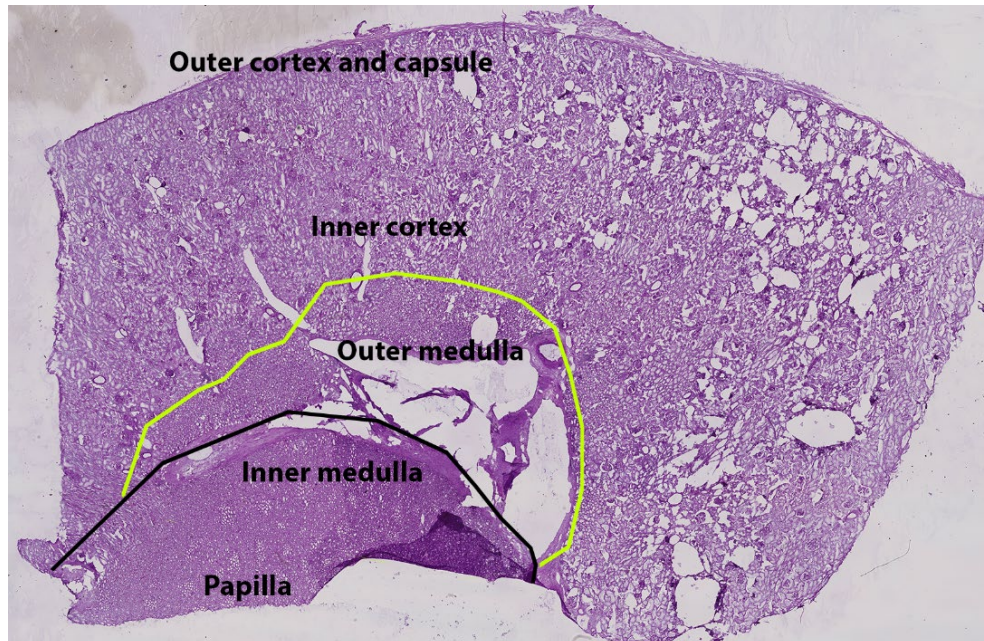
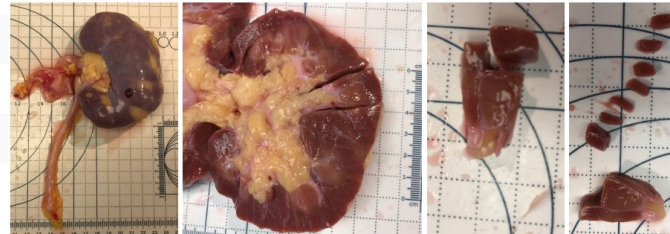
# Data: 3D Tissue

**Kidney: Jeff Spraggins et al., VU**

See data on Globus, BIOMIC\_patient-64354

- 📄 BIOMIC\_patient-64354\_clinical\_and\_spatial\_metadata.xlsx
- 📄 BIOMIC\_patient-64354\_data\_guide.pptx
- 📄 BIOMIC\_patient-64354\_overview.png
- 📄 BIOMIC\_patient-64354\_Sample-20-Histology.tif
- 📁 neg\_ion\_mode\_section
- 📁 pos\_ion\_mode\_section

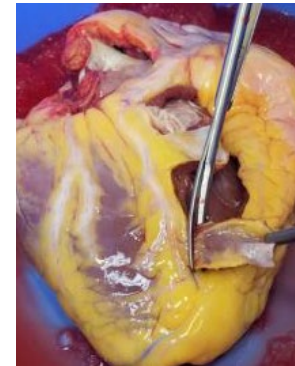
BUKMAP, Zhang Group



**Heart: Shin Lin, UW**

Year 1: Tissue data for 1-2cm cubed volumes from 9 sites for 1 heart from 1 individual.

Terminology; Coordinates and photos to spatialize



<u>Sites</u>	<u>Distinctive features</u>
1. LV, apex	
2. LV, free wall 3 cm from apex	
3. septum, 3cm from apex including LAD	major arterial vessel, Purkinje fiber CM
4. RV, free wall 3 cm from apex	
5. RA appendage	
6. RA, SA node to AV node	pacemaker CM
7. LA, appendage	
8. LA, PV inflow	
9. Posterior, adjacent to coronary sinus	major venous vessel

# Data: TMCs x Organs x Data Types x Technologies

BUKMAP, Zhang Group

## Organs (10)

1. Bladder
2. Blood Vessel (Heart)
3. Breast
4. Colon
5. Kidney
6. Liver
7. Lung
8. Spleen
9. Thymus
10. Tonsil

No Bone Marrow  
and Pancreas.

## Data Types (13)

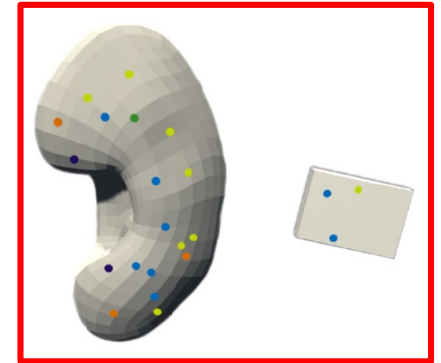
1. Imaging - Proteins
2. Imaging - RNA
3. Imaging - DNA
4. Imaging - Other
5. scRNAseq
6. scDNAseq
7. scProteomics
8. bulk-Proteomics
9. bulk-RNA
10. bulk-DNA
11. Metabolomics
12. Lipids
13. Other

## Technologies (~25)

- CODEX;DART-FISHrp;IF;IHC;LRET-IF;MALDI Imaging MS;PER-DEI  
DART-FISH;LRET-ISH;MERFISH;PER-DEI;seqFISH;smFISH  
PER-DEI  
Lipid and Metabolite MALDI Imaging MS  
snDropseq;scRNAseq  
scATACseq;scTHSseq;SNAREseq  
IMC  
LC-MS/MS  
?  
?  
LC-MS/MS;nano-POTS  
LC-MS/MS;nano-DESI  
Autofluorescence;PAS stained microscopy

# CCF Data Wiki

## Minimum Information Standard



hubmapconsortium / ccf-data-wiki Private

Unwatch 3 Star 0 Fork 0

Code Issues 0 Pull requests 0 Wiki Insights

## Home

Bruce Herr II edited this page 2 days ago · 5 revisions

Edit New Page

## Welcome to the CCF Data Wiki!

Organ	CalTech	UCSD	Stanford	Vanderbilt	Florida
Heart	✓				
Kidney		✓		✓	
Bowel			✓		
Thymus					✓
Spleen					✓
Lymph Nodes					✓
Lung		✓			
Bladder		×			
Colon			×		

### Legend:

- ✓ - Organ proposed and survey submitted
- × - Organ was proposed, but no survey has been submitted

Pages 12

### Links

- Home

### Templates

- TMC-Organ-Template

### Clone this wiki locally

<https://github.com/hubmapcc>

<https://github.com/hubmapconsortium/ccf-data-wiki/wiki>

# CCF User Interfaces (UI)

# CCF User Interface (UI)

The screenshot displays the HuBMAP Semantic Search interface. At the top, the search bar is empty, and the user profile is identified as 'Female | Age 19-41'. The left sidebar contains navigation icons for HOME, BODY, ORGAN, TISSUE, and CELL. A 'Semantic Browsing' section lists anatomical regions under 'Body' and 'Kidney', with 'Renal Capsule' highlighted. Below this is a 'Semantic Filters' section with tabs for 'Technologies' and 'TMCs', showing filters for 'IMS', 'MxIF', and 'AF'. The main area features a large kidney tissue image with a 25 µm scale bar and a 'Multiplex' panel showing three channels: 'Hoechst' (blue), 'DBA' (red), and 'LTL' (green). The right panel provides details for the 'Kidney' sample, including patient information, procedure details, and a list of markers.

**HuBMAP Semantic Search**

Female | Age 19-41

**Body**

- Heart
- Kidney**
  - Anterior Surface
  - Head Kidney
  - Hilum Of Kidney
  - Renal Capsule**
  - Renal Corticomedullary Boundary
  - Renal Epithelium
  - Renal Interstitium
  - Renal Vasculature
  - Posterior Surface
  - Renal Collecting System
  - Renal Duct
  - Renal Fat Pad
  - Renal Lobe
  - Renal Medulla
  - Renal Parenchyma
  - Renal Pelvis
  - Renal Sinus

**Semantic Browsing**

**Semantic Filters**

Technologies TMCs

IMS MxIF AF

**Kidney**

**Multiplex IF**

Multiplex

**Blue Channel**

Technology: Hoechst DNA stain  
Marker: Cell nucleus

**Red Channel**

Technology: DBA (Dolichos Biflorus Agglutinin) binds to carbohydrates that contain a-linked N-acetylgalactosamine residues  
Marker: General for renal collecting ducts.

**Green Channel**

Technology: LTO (Lotus tetragonolobus Lectin) binds specifically to carbohydrates that contain a-linked L-fucose oligosaccharides  
Marker: Proximal tubule epithelial cells

**Details**

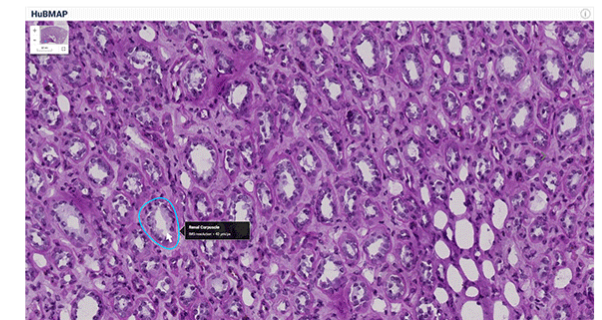
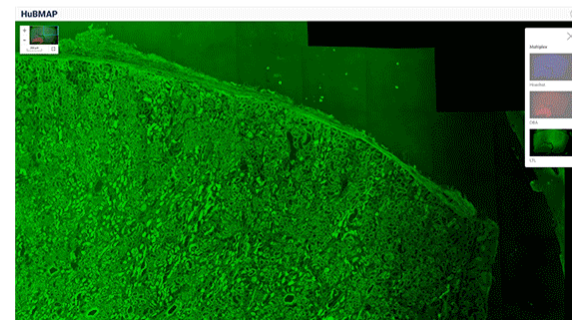
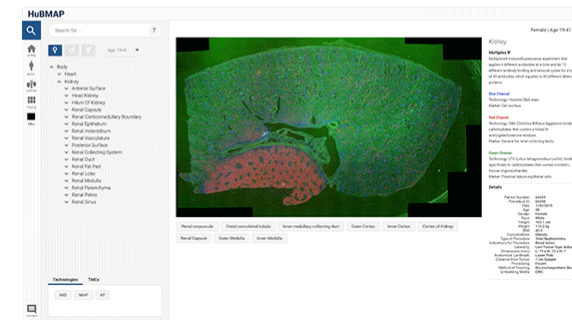
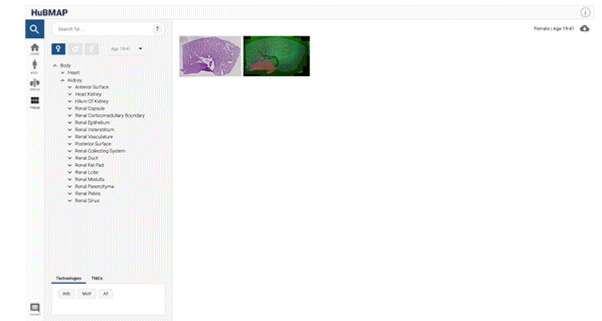
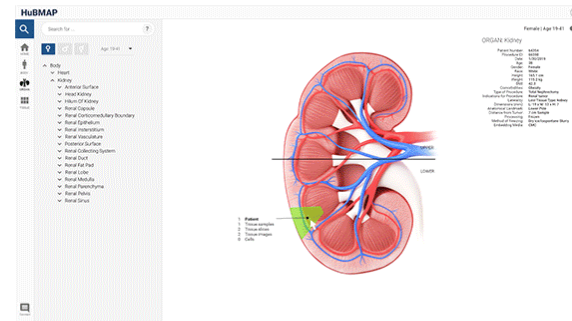
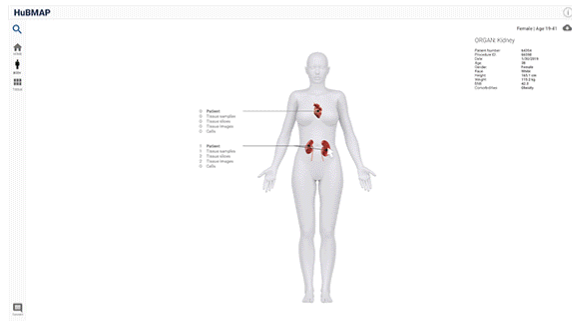
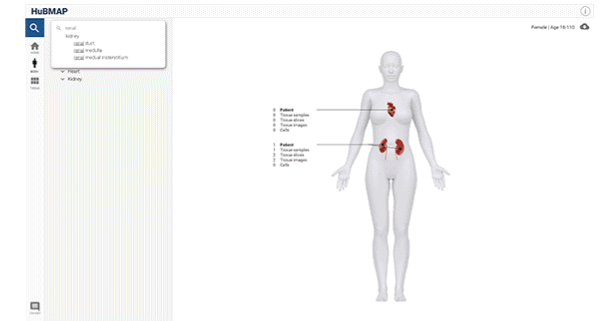
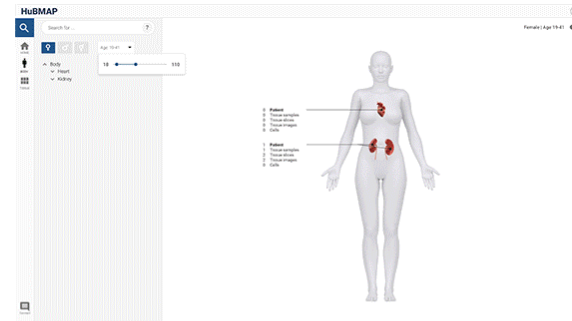
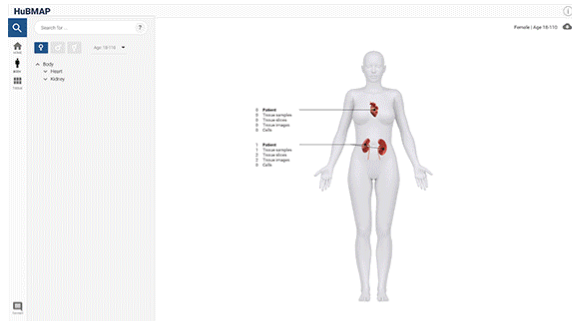
Patient Number: 64354  
Procedure ID: 66598  
Date: 1/30/2019  
Age: 38  
Gender: Female  
Race: White  
Height: 165.1 cm  
Weight: 115.2 kg  
BMI: 42.3  
Comorbidities: Obesity  
Type of Procedure: Total Nephrectomy  
Indications for Procedure: Renal tumor  
Lateralality: Left Tissue Type: kidney  
Dimensions (mm): L: 19 x W: 13 x H: 7  
Anatomical Landmark: Lower Pole  
Distance from Tumor: 7 cm Sample  
Processing: Frozen  
Method of Freezing: Dry ice/Isopentane Slurry  
Embedding Media: CMC

Renal corpuscule Distal convoluted tubule Inner medullary collecting duct Outer Cortex Inner Cortex Cortex of Kidney

Renal Capsule Outer Medulla Inner Medulla

GitHub demo site: <https://hubmapconsortium.github.io/ccf-ui/>

# CCF User Interface (UI)



# CCF User Interface (UI)

MC-IU has released CCF user interface v0.5.0, a proof-of-concept version of the CCF UI.

The user interface supports:

- Visual browsing of tissue samples and metadata at the whole body, organ, tissue, and cell level.
- Filtering by metadata (age, gender, TMC, and technology), results are presented at all views.
- Submission of questions and comments on the CCF UI.
- Semantic search by ontology, results are presented at all levels.
- Data download at the whole body, organ, tissue, and cell level, i.e., link to <https://sampledata.hubmapconsortium.org>.

See also:

Recorded demo: [https://www.youtube.com/watch?v=rWMqKQc\\_00w&feature=youtu.be](https://www.youtube.com/watch?v=rWMqKQc_00w&feature=youtu.be)

GitHub link to code: <https://github.com/hubmapconsortium/ccf-ui>

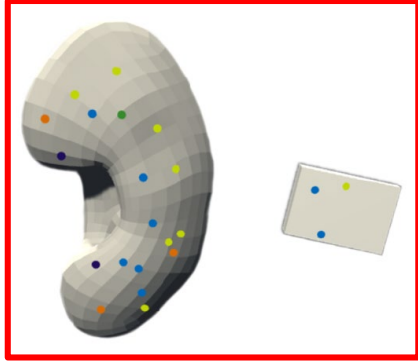
GitHub demo site: <https://hubmapconsortium.github.io/ccf-ui/>

Original specs: <https://drive.google.com/open?id=1tqUzmVLxwqcGprtRlevfY86YvHHPEsDR>

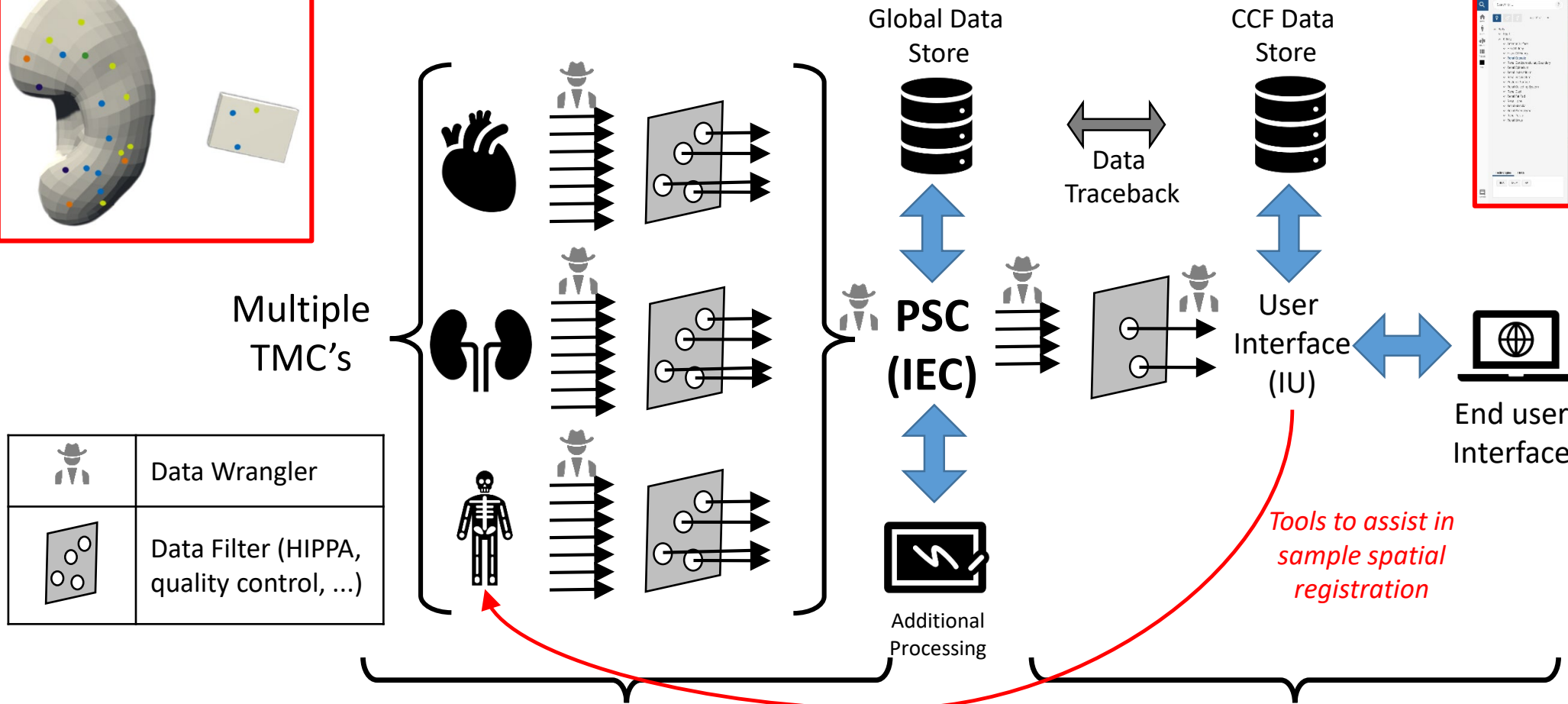
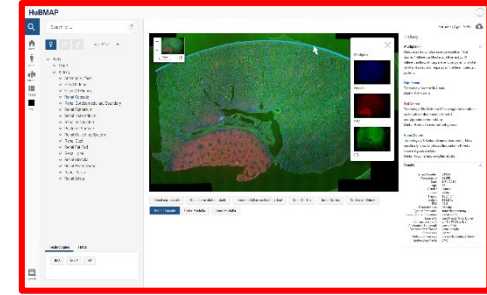
**Live Demo!**



## Tissue Registration UI



## CCF User Interface (UI)



	Data Wrangler
	Data Filter (HIPPA, quality control, ...)

- Provenance
- Patient
- Sample
- Sample Processing
- Technology (MS, IH, ...)
- Analysis
- Etc.

*Propagate needs back to TMC's*

- Only the data needed for the GUI

TMC: Tissue Mapping Center  
PSC: Pittsburgh Supercomputing Center

# Tissue Registration UI: Heart (depending data availability)

Align 9 tissue samples in 3D heart using a combi of

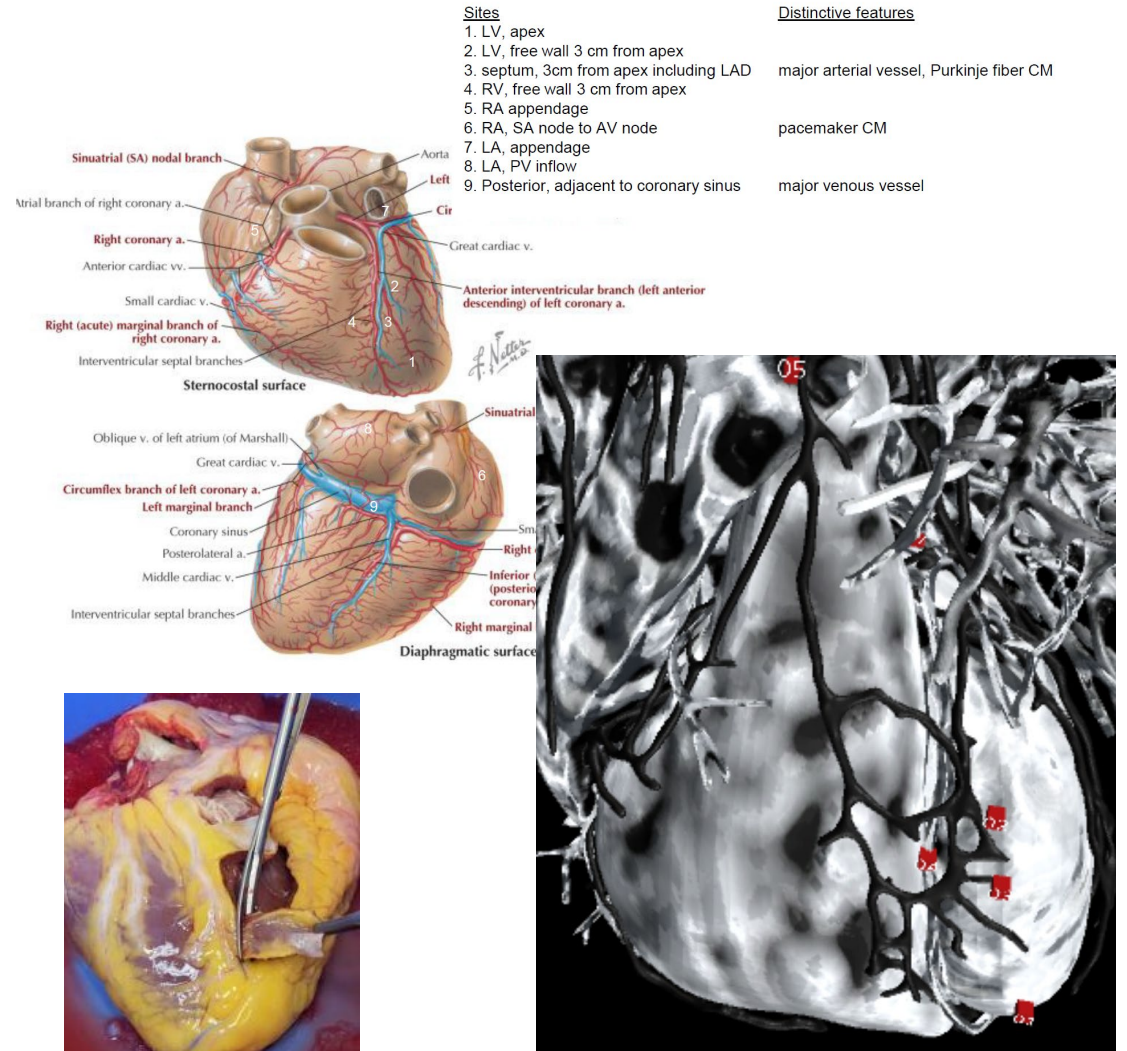
- Rough placement using human expertise/3D pattern matching and
- Fine adjustments using machine learning

Virtual tissue samples will be sized 1-2cm cubed, numbered (1 ... 9), and oriented (left-right, top and bottom tissue slice of z-stack).

Measure error from

- precision of tissue sample procurement and
- placement in the 3D browser

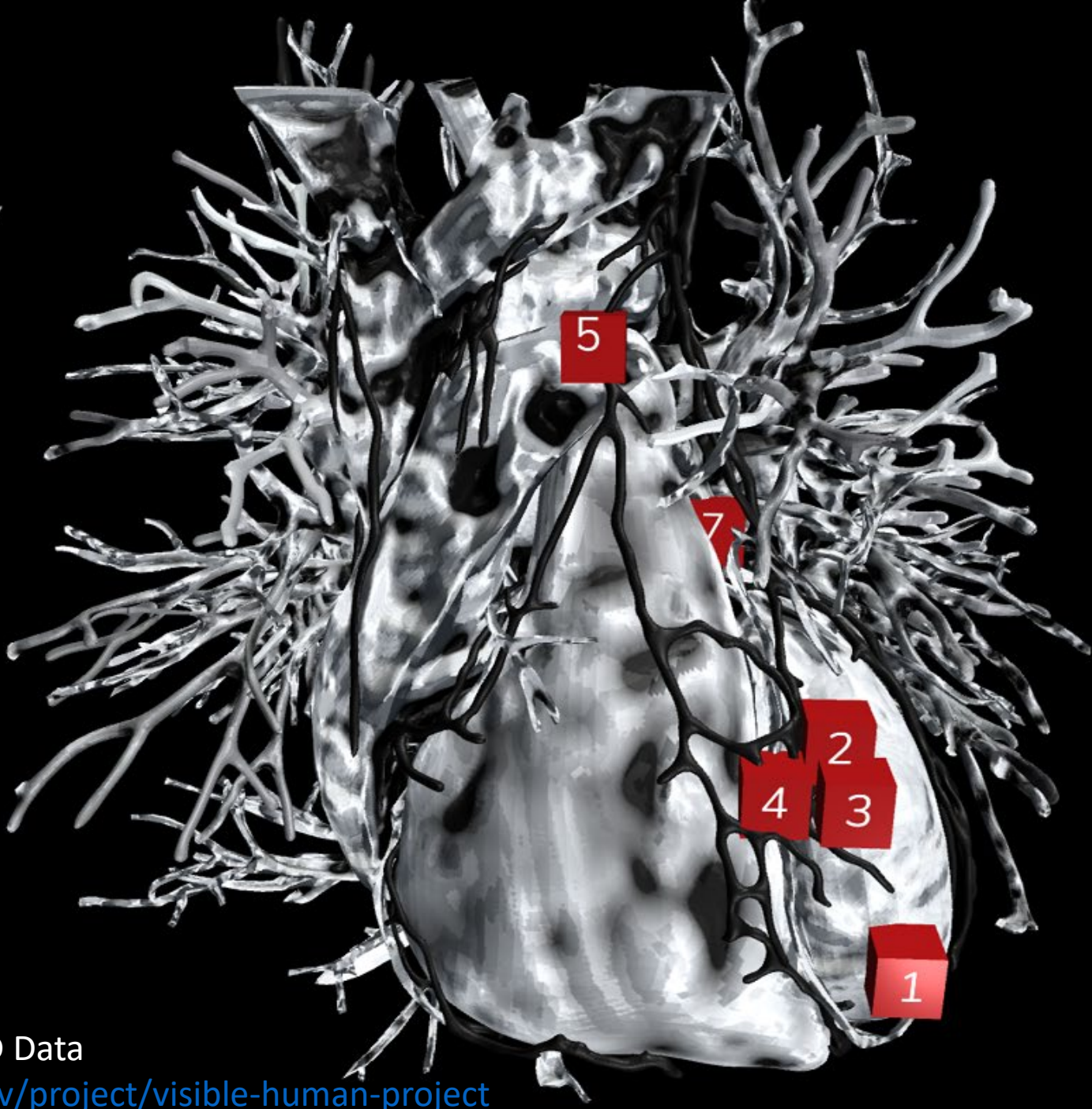
We hypothesize that placement accuracy will improve when additional information (e.g., landmarks, major scaffolds, MR/CT scan of heart after 9 samples were extracted) is being visible in virtual organ.



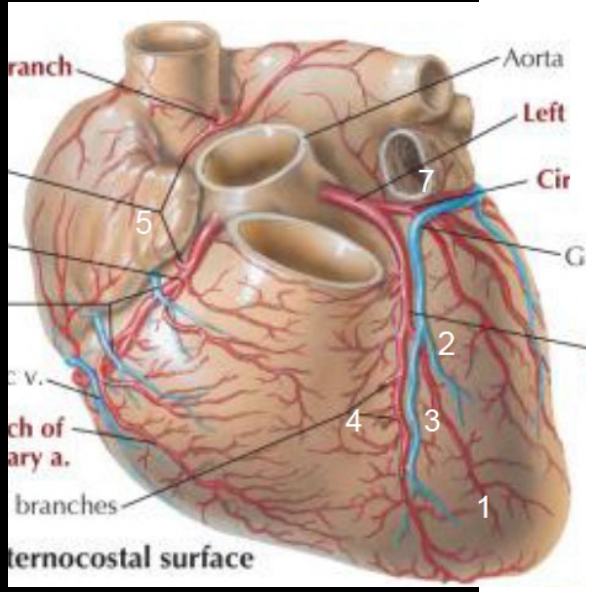
Human heart with data overlay  
Developer: Andreas Bueckle

- Show/hide
- Coronary arteries
  - Coronary veins
  - Left atrium
  - Left ventricle
  - Right atrium
  - Right ventricle
  - Markers

Adjust camera speed



Currently Selected  
Please click any of the red markers!



Heart model from NLM3D Data  
<https://lhncbc.nlm.nih.gov/project/visible-human-project>

# Tissue Registration UI: Kidney

(depending data availability)

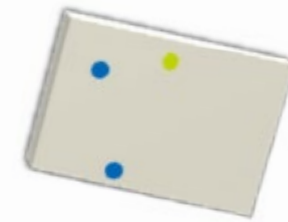
- Exploit human pattern recognition and fine motor skills (by surgeons) to register tissue in organs.
- Add info on anatomical landmarks (fiducial marks), cell types, molecular data to support alignment.
- LATER: Use human alignment data as training data for machine learning algorithms, to better support manual alignment OR to possibly fully automatize alignment.



VIVE™ | VIVE Virtual Reality System  
vive.com

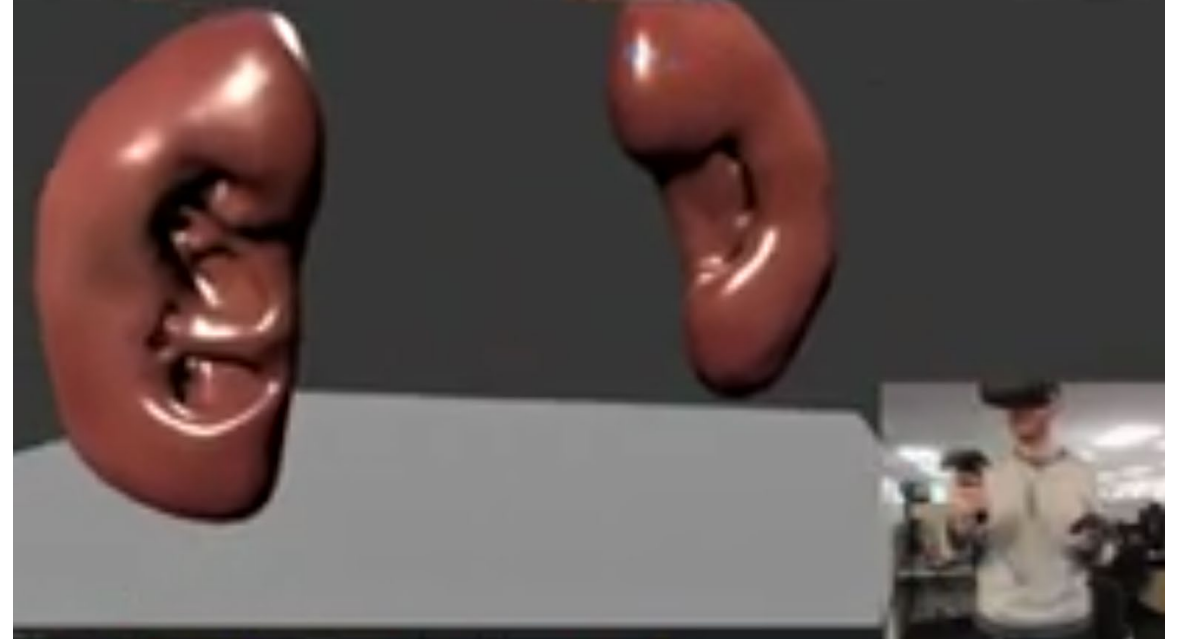
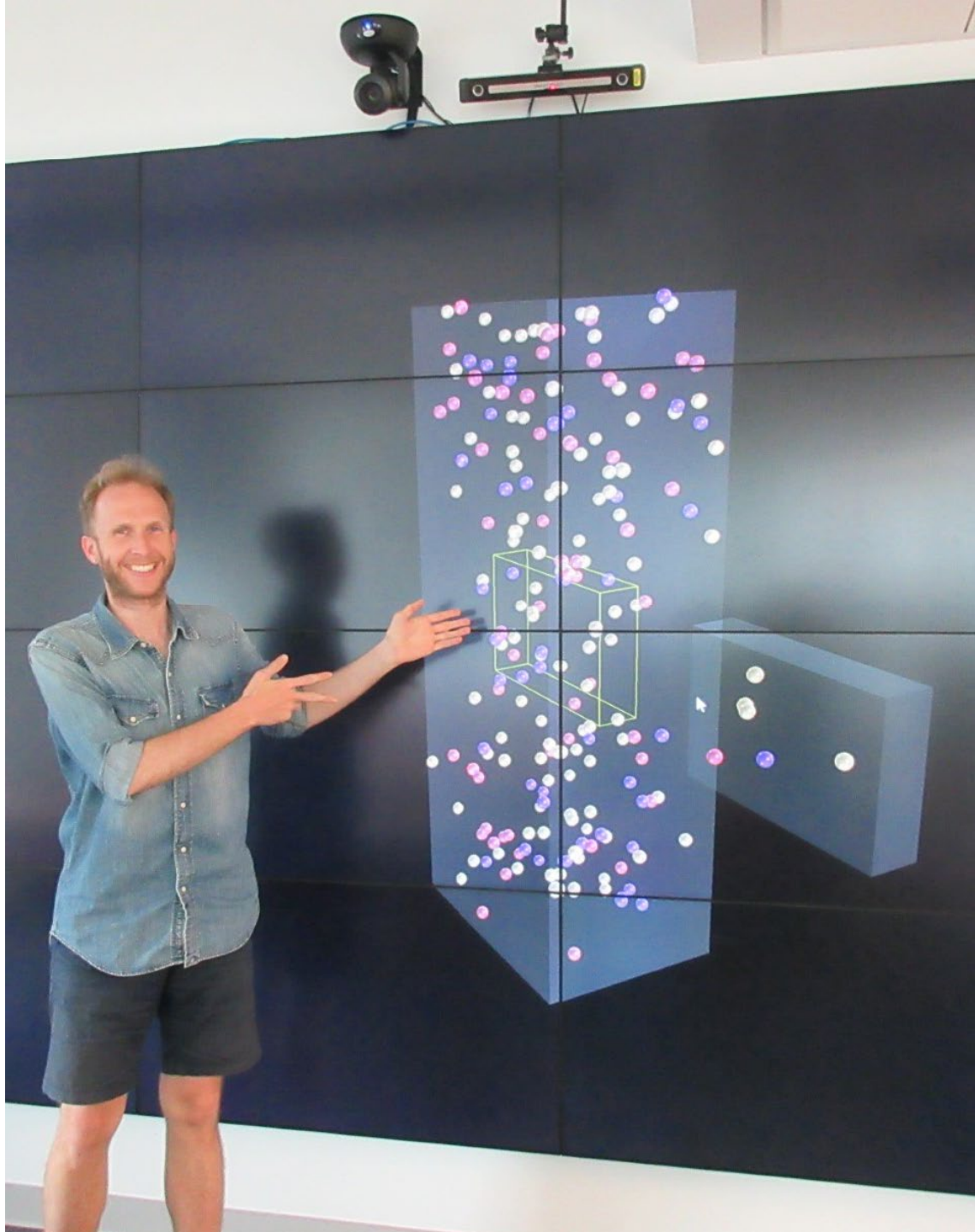


**How many of you have used  
a VIVE or space mouse?**



Kidney model from NLM3D Data

<https://lhncbc.nlm.nih.gov/project/visible-human-project>

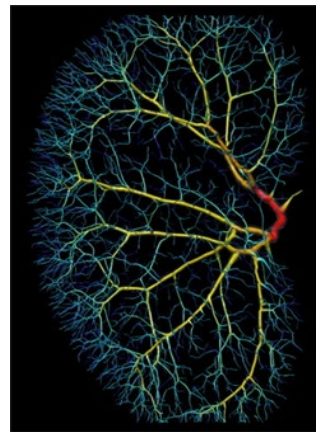
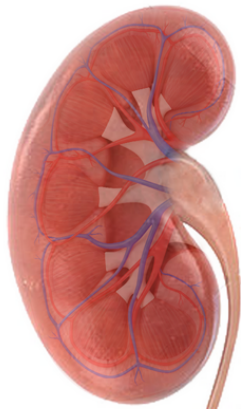
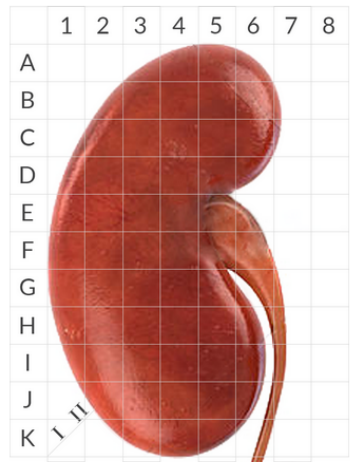


Kidney model from NLM3D Data

<https://lhncbc.nlm.nih.gov/project/visible-human-project>



For the kidney, there exist no predefined tissue extraction sites. The current kidney Registration UI uses a grid system and a picture of a kidney slice to guide placement, see **Fig. 3 left and middle**. Funding of this GLUE grant will make it possible to use 3D image volumes collected from intact kidney tissue within KPMP that feature anatomically meaningful structures of different cell types, see **Fig. 3 right**.

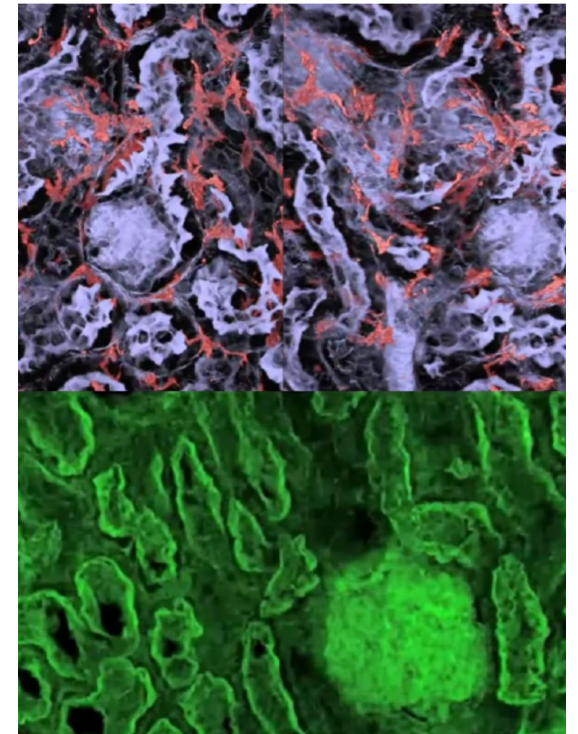


Structural morphology of renal vasculature. Nordsletten et al. 2006. <https://doi.org/10.1152/ajpheart.00814.2005>

**Figure 3:** 3D grid system (left), 2D image (middle), 3D tissue reconstructions from KPMP (right)

Using the 3D grid system, a user can use 3D coordinates to refer to a particular area in the kidney, e.g., A-4-II would correspond to the lower-middle part of the kidney, on the inside (occluded in this view). Using the 2D image, sample placement within a semi-transparent reference kidney object is guided by anatomical structures. Using KPMP data, it will be possible to show complex volumetric structures inside the 3D reference kidney to allow for more granular alignment at the molecular level.

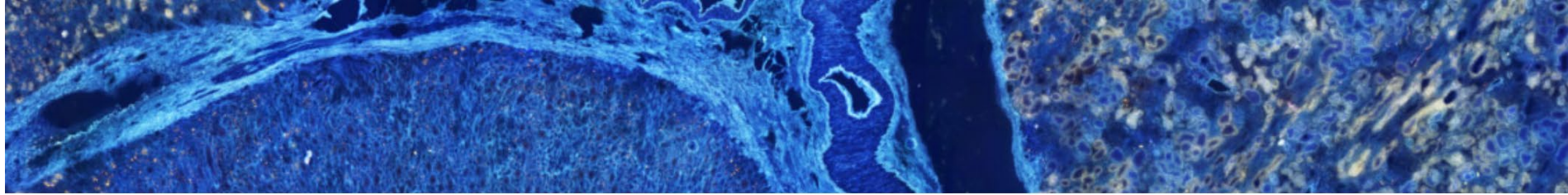
KPMP GLUE grant proposal in progress.



**Fig. 4.** KPMP volumetric data.

See presentation by Seth Winfree for details <https://ccfws.cns.iu.edu>

# CCF Workshop (MC-IU, Jeff Spraggins, TMC-Vanderbilt, Sanjay Jain, TMC-UCSD)



**HuBMAP**  
Human BioMolecular Atlas Program

COMMON COORDINATE  
FRAMEWORK WORKSHOP  
CCFWS-01

## Time & Date

9:00am–5:00pm EDT on May 9, 2019

All slides, video recordings are at <https://ccfws.cns.iu.edu>

## Goals

HuBMAP will develop a common reference map or coordinate system called the Common Coordinate Framework (CCF). As stated in the Common Coordinate Framework Meeting (CCFM) document, a CCF makes it possible to uniquely and effectively define and name any location in the human body. A set of robust origin points (serving as landmarks) make it possible to reference organs, tissues, cells over different anatomical scales, tolerate human variability and function across lifespan and disease, and help integrate heterogeneous data layers and a wide range of reference maps such as whole body spatial maps, genetic variant maps, and coordinate systems that align with vascular pathways.

This CCF workshop will focus on a kidney-specific CCF and atlas but also discuss other relevant CCF/atlas efforts. It will feature presentations and discussions on:

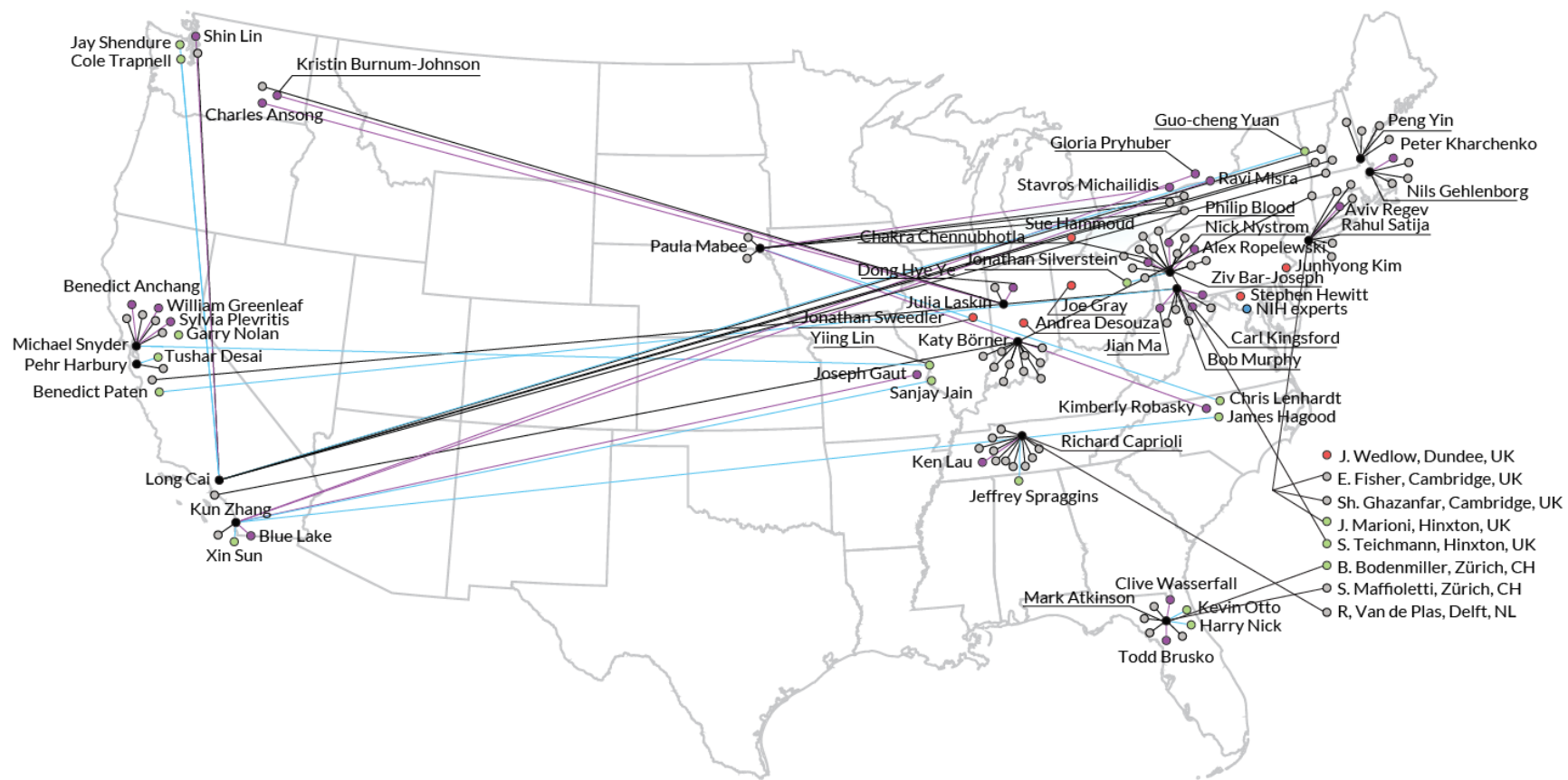
- CCF metadata—what data are currently captured, how can they be unified across tissue mapping centers (TMCs), what additional data are needed to meet stakeholder (research) needs.
- CCF ontologies—what ontologies exist and are used in what part of the data pipeline; what ‘desirable properties’ and ‘success criteria’ exist?
- CCF mapping and numerical construction—including dealing with human variation and using CCF user interfaces as a means to properly register data and review data completeness.
- General principles and processes that can inform CCF design for other organs and continuous adaptation of CCF to emerging technologies and ever changing user requirements.

The ultimate goal is a set of draft guidelines for TMCs detailing what data to provide in which formats to maximize CCF mapping accuracy and data utility.

# Geospatial Layout of HuBMAP Teams

MC-IU within HuBMAP (<http://hubmapconsortium.org>)

04/16/2019



## Legend

Label: Experts  
Color: Role

Nodes and Edges

- PI (Contact)
- PI
- Co-Investigator

- NIH
- EPC

## How To Read This Visualization

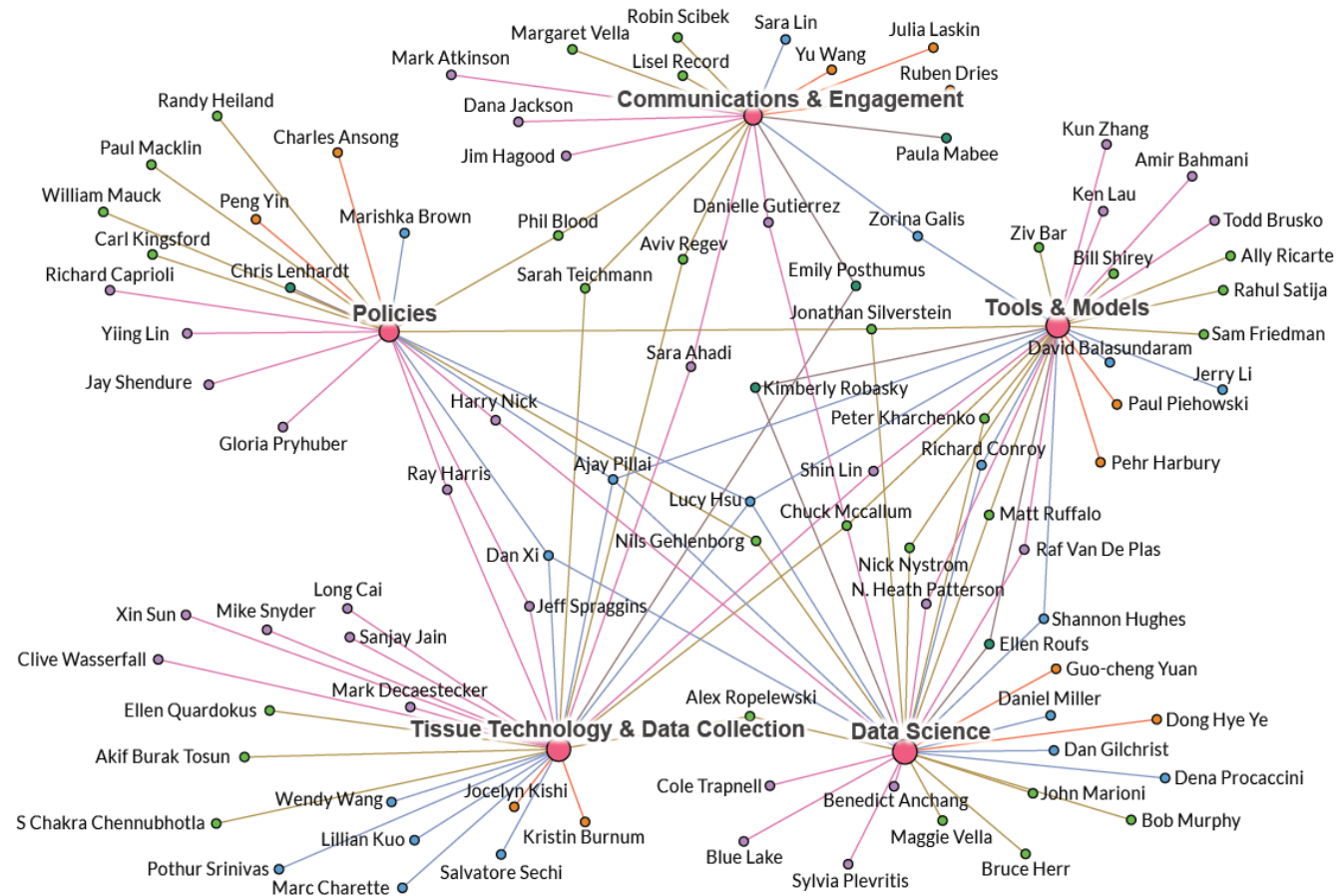
This geospatial map shows the collaboration network of 134 experts. Principal investigators (PIs) are placed at their exact geolocation. Team members are placed nearby and are linked to PIs. Nodes and edges are colored according to their roles.



# Bimodal Network of Experts and Working Groups

MC-IU within HuBMAP (<https://hubmapconsortium.org>)

04/09/2019



## Legend

Label: Experts  
Working Groups  
Color: Team Type

Nodes and Edges  
■ WGs    ■ TMC  
■ HIVE    ■ NIH  
■ TTD    ■ CC

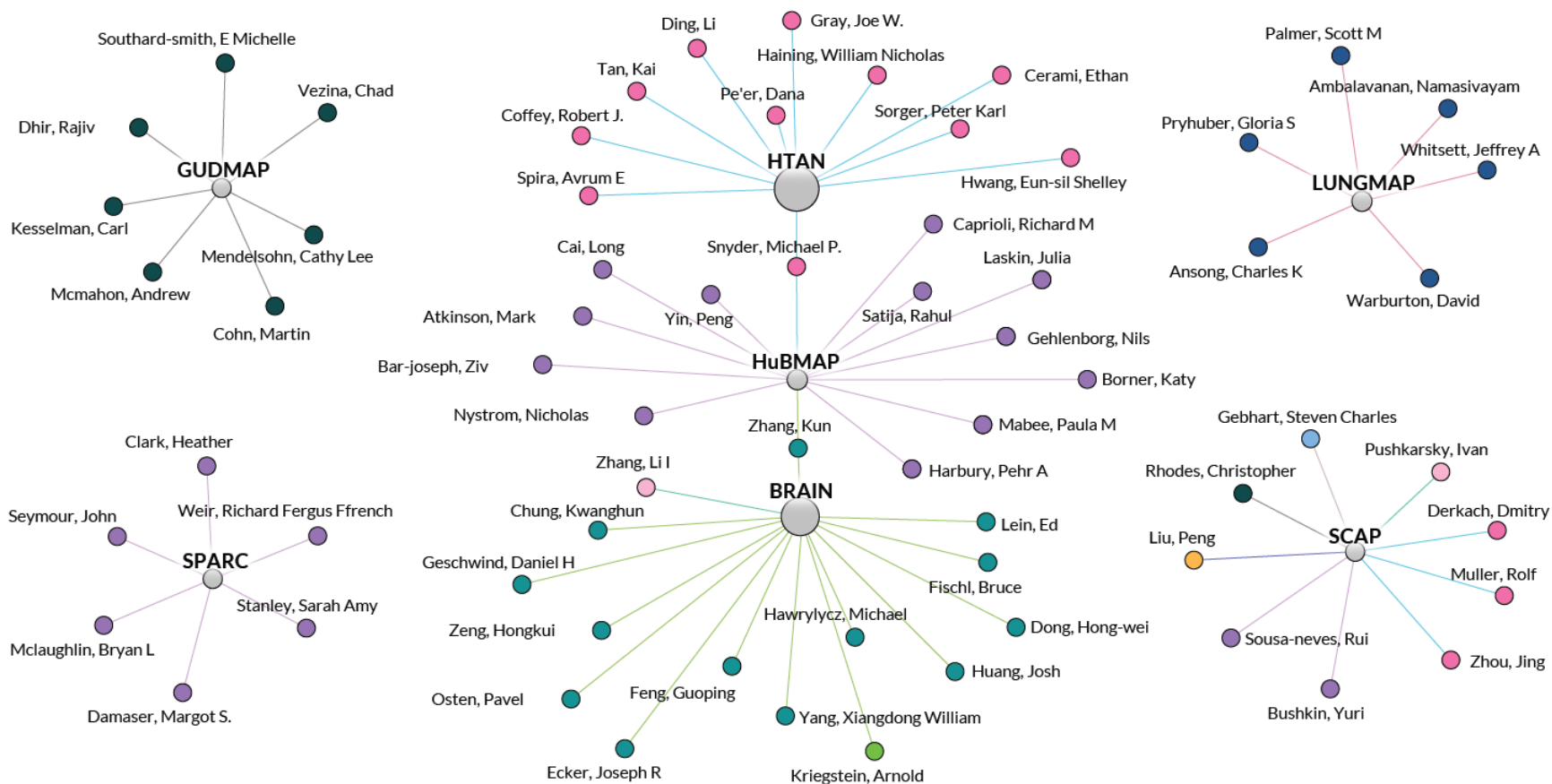
## How To Read This Visualization

This bimodal network represents a network between five working groups and 126 experts of those groups. Nodes and edges are colored according to the team type that expert belongs to. Working group nodes are sized by the total number of experts in the group.

# Bimodal Network of PI (Contacts) and NIH Projects

MC-IU within HuBMAP (<https://hubmapconsortium.org>)

04/22/2019



## Legend

Label: PI (Contacts)  
 NIH Projects  
 Maximum = 71,882,084  
 Minimum = 3,065,740

- OD
- NIMH
- NCI
- NIDDK
- NHLBI
- NCCIH
- NHGRI
- NIGMS

## How To Read This Visualization

This bimodal network represents a network between six NIH projects and 73 PI (Contacts). Nodes and edges are colored according to the Funding Institute & Center that project of Pi (Contact) is financed from. NIH project's node sizes based on award amount.