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# Whole-brain microscopy image analysis

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# Fluorescence

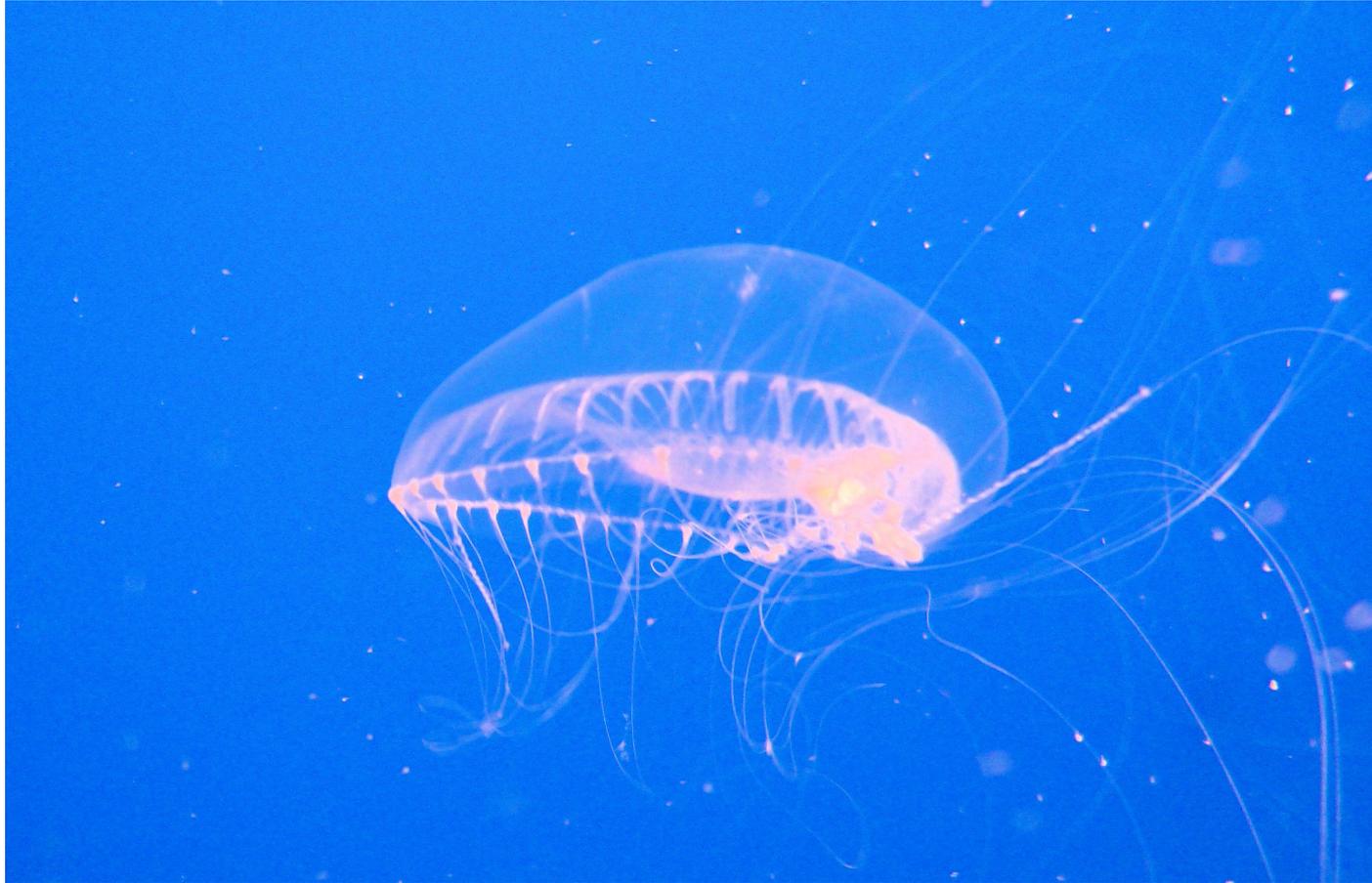
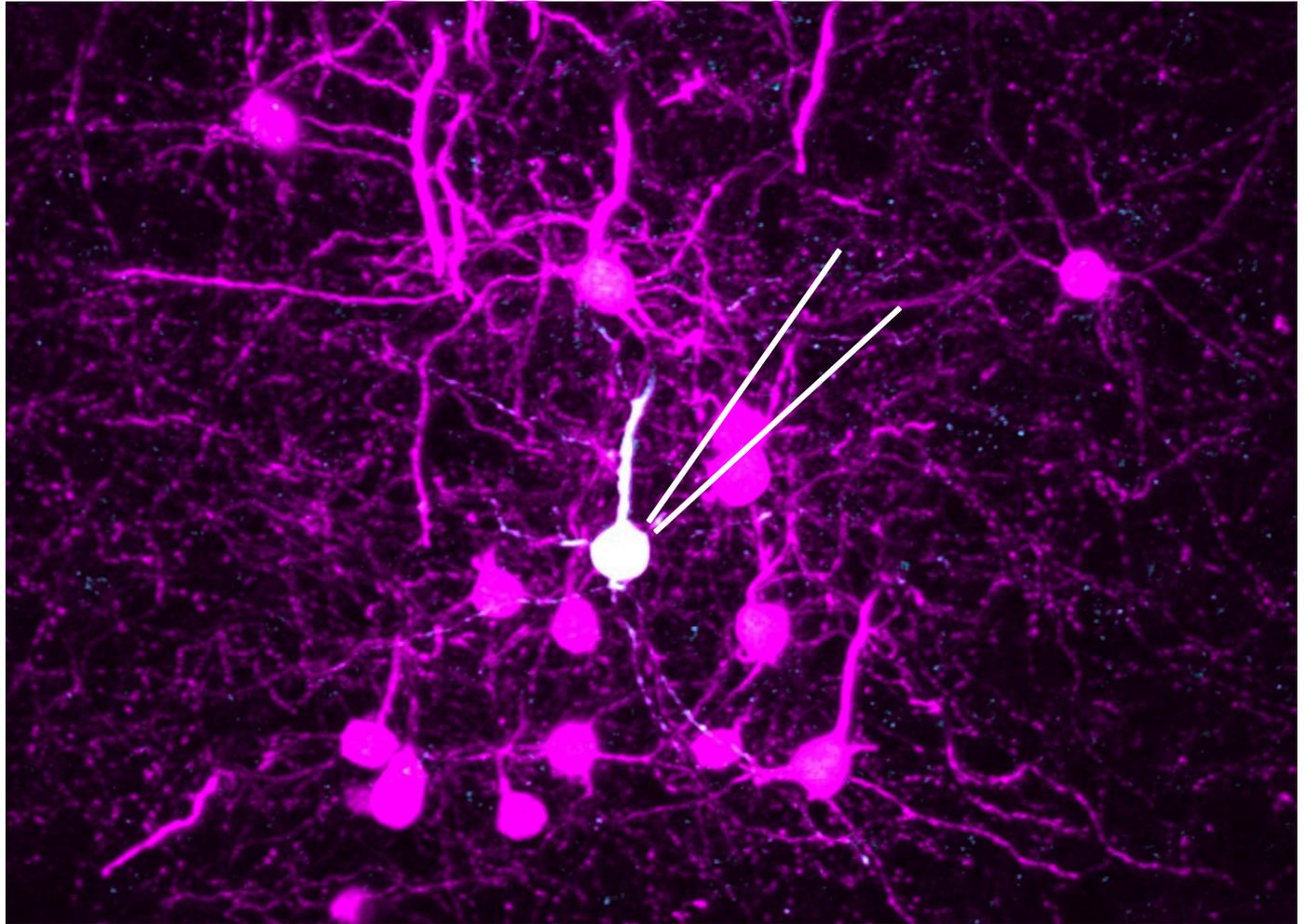
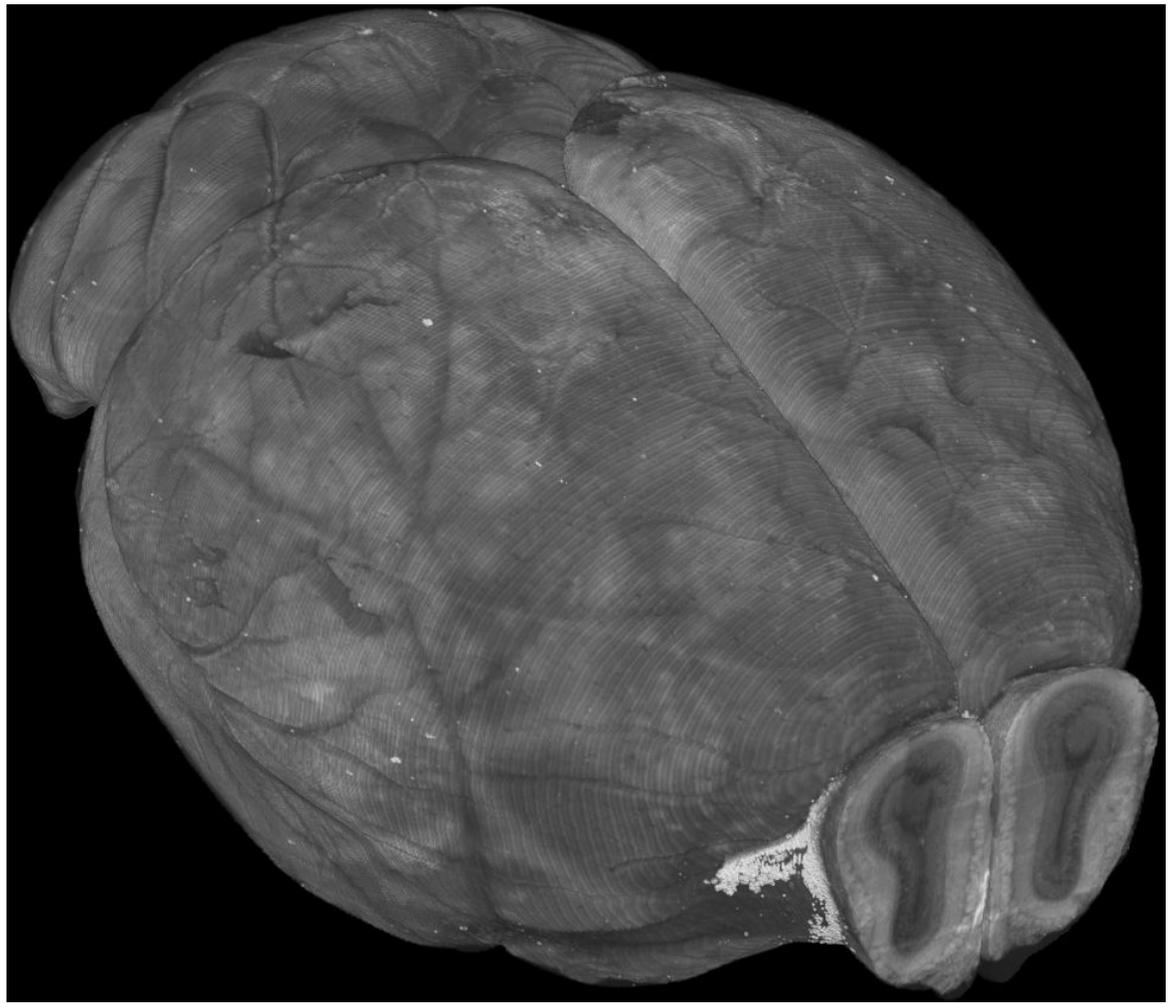


Photo: <https://www.flickr.com/photos/jimg944/5085398602/>

# Viral tracing

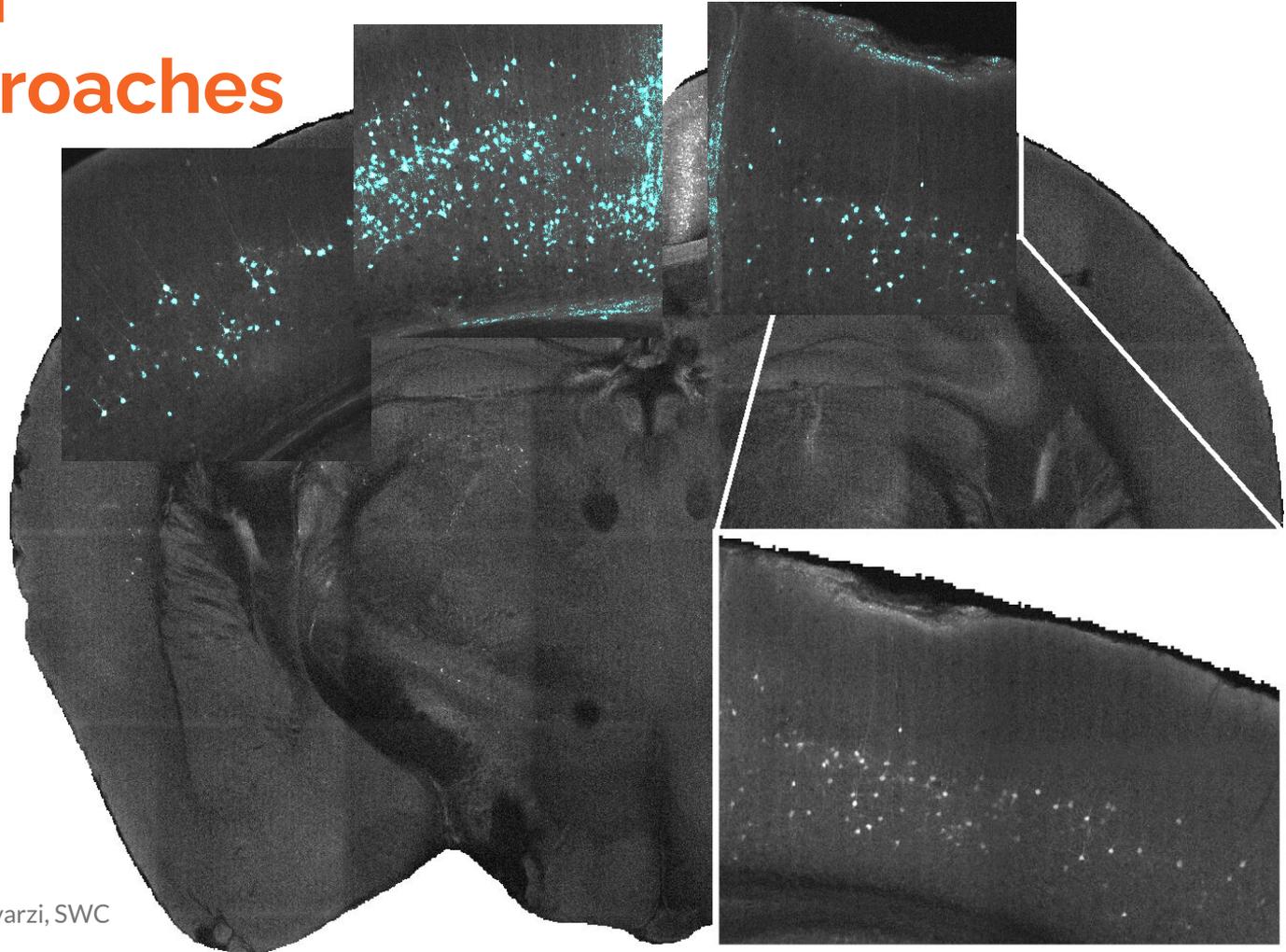


# Whole-brain microscopy

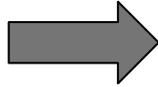
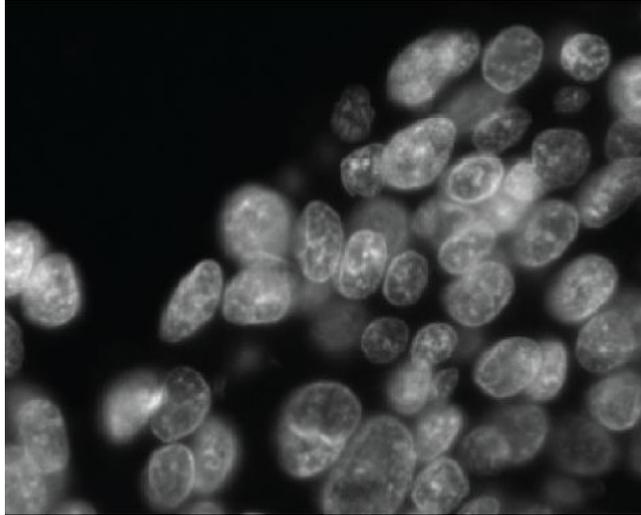


Data: Chryssanthi Tsitoura & Sepiedeh Keshavarzi, SWC

# Cell detection -classical approaches

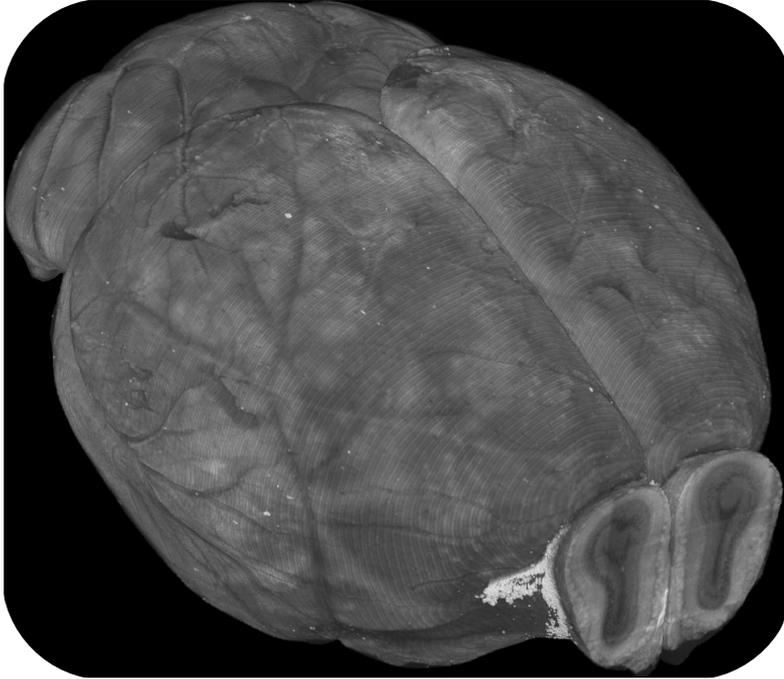


# Cell detection - deep learning



StarDist - Weigert et al. (2020) WACV

# Cell detection - deep learning

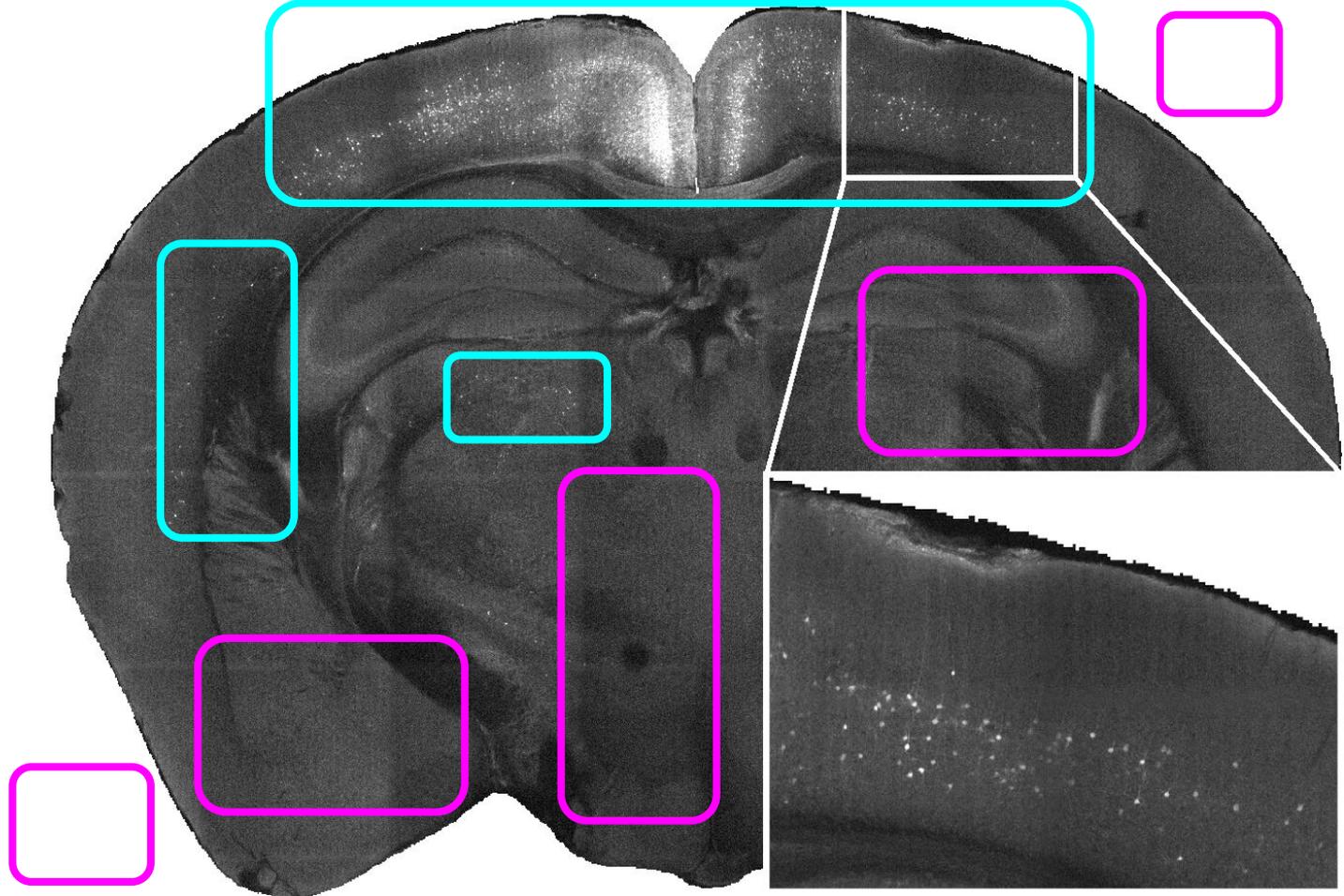


200-1000 GB

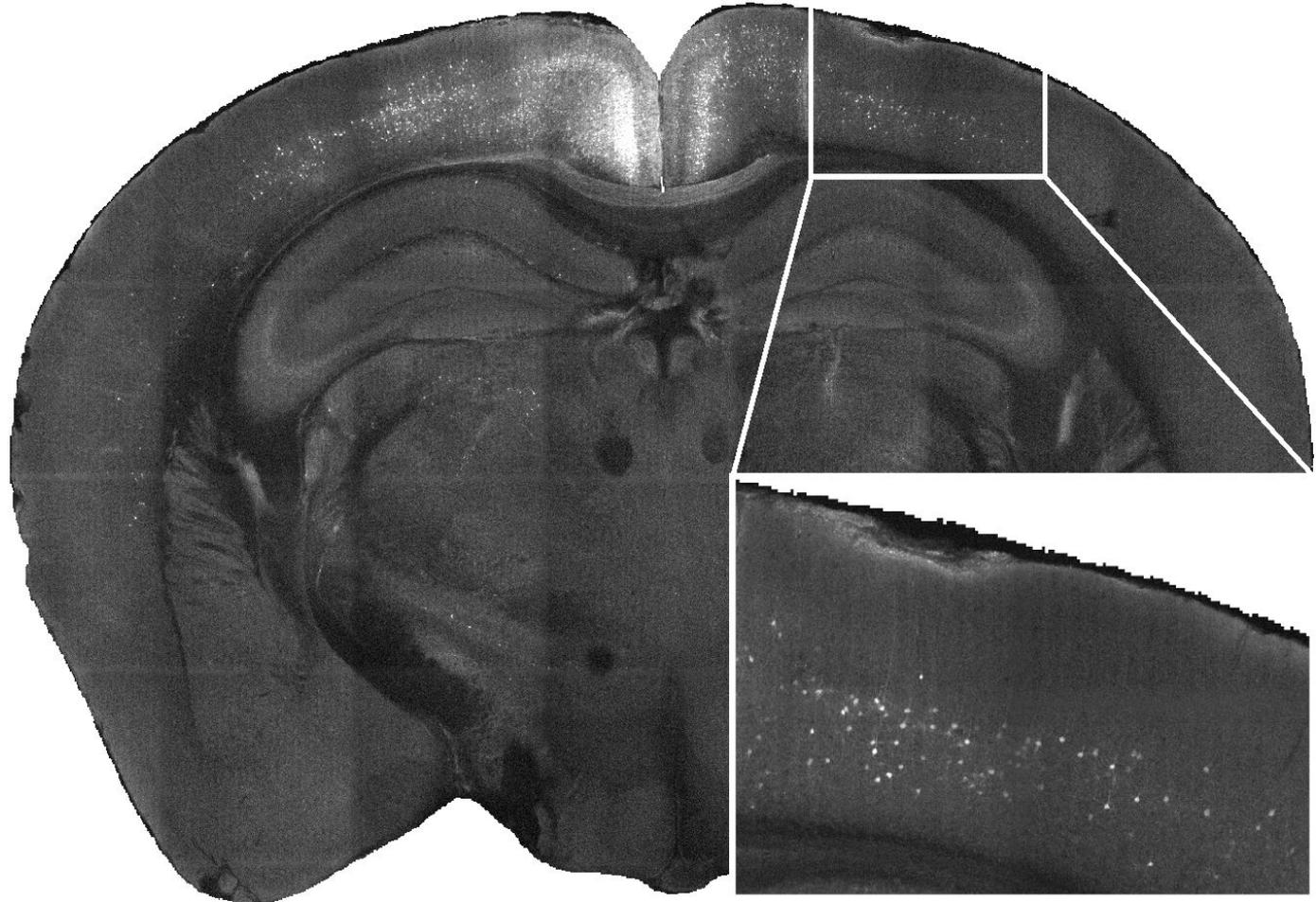


24GB VRAM

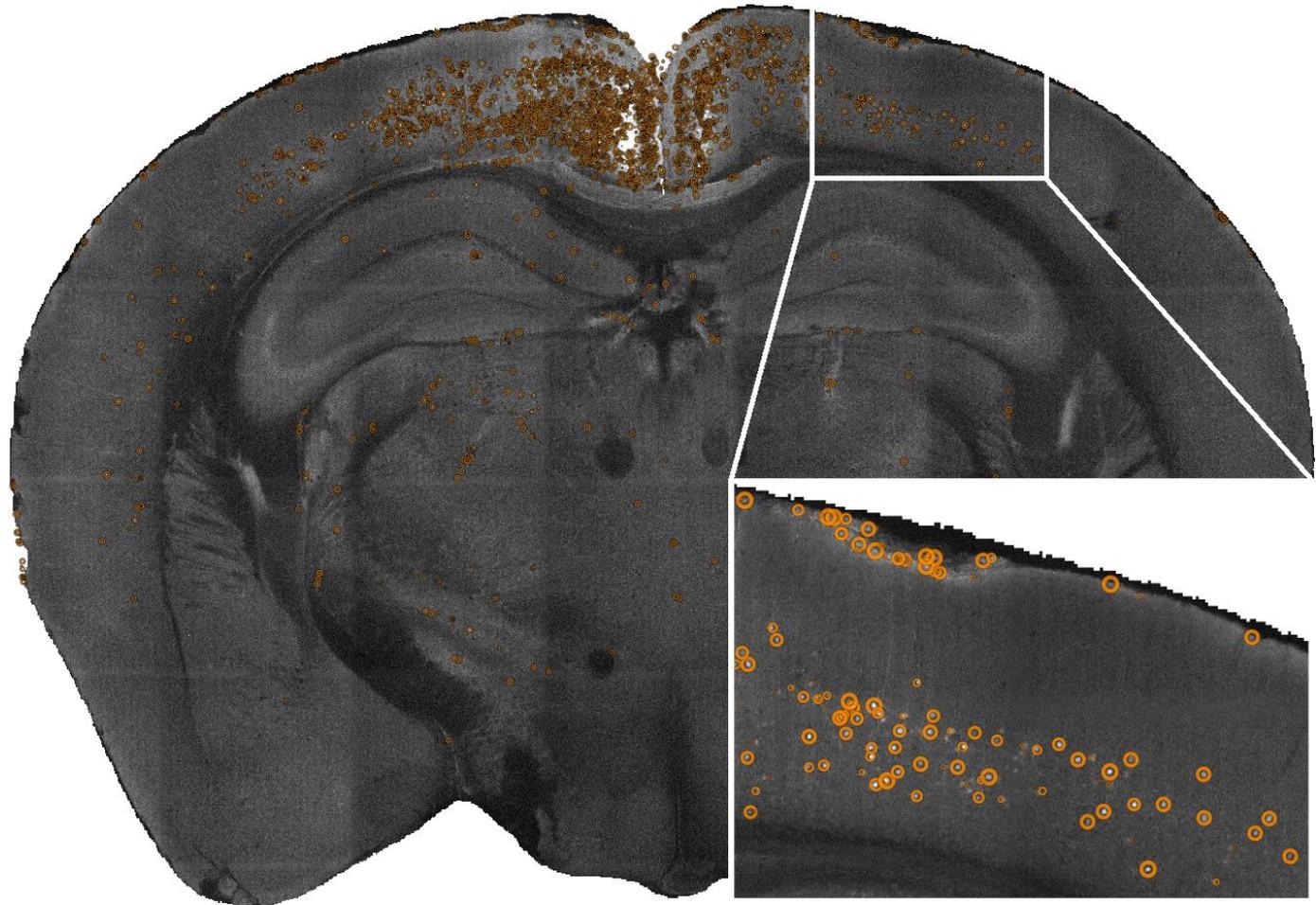
# Cell detection - data reduction



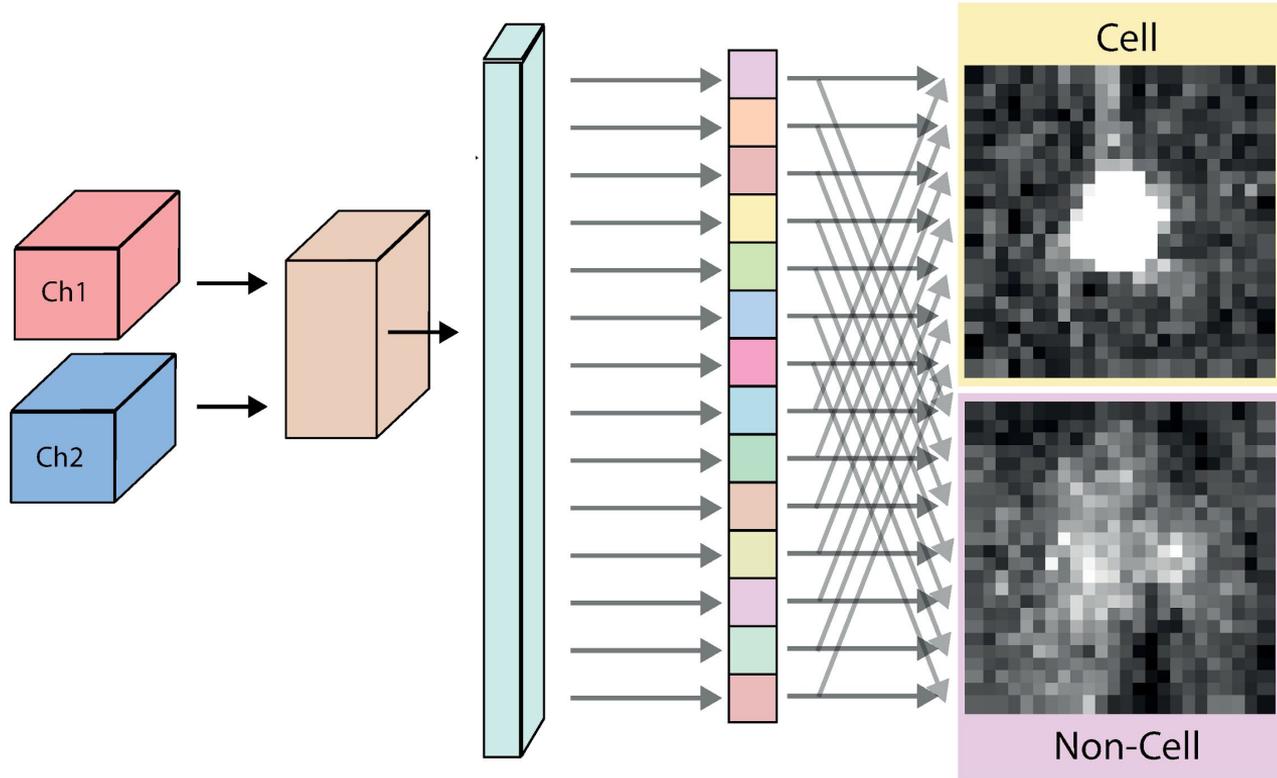
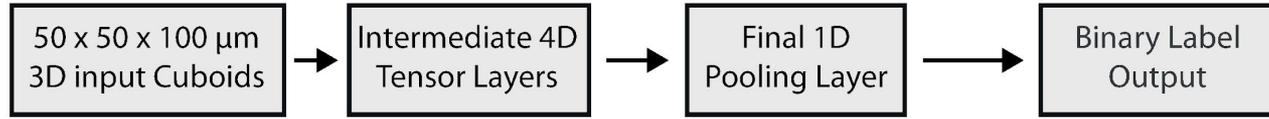
# Cell detection - data reduction



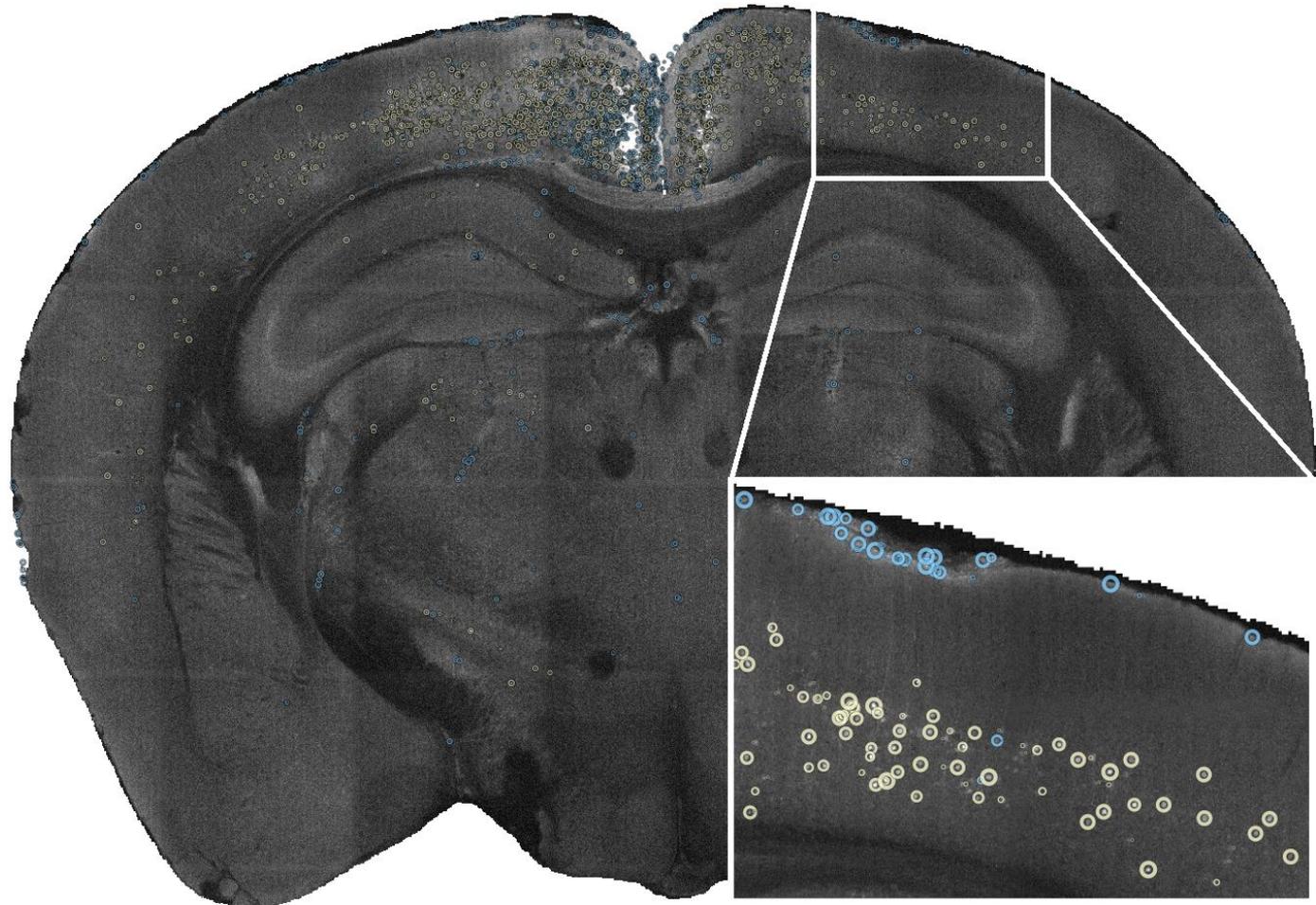
# Cell detection - detection



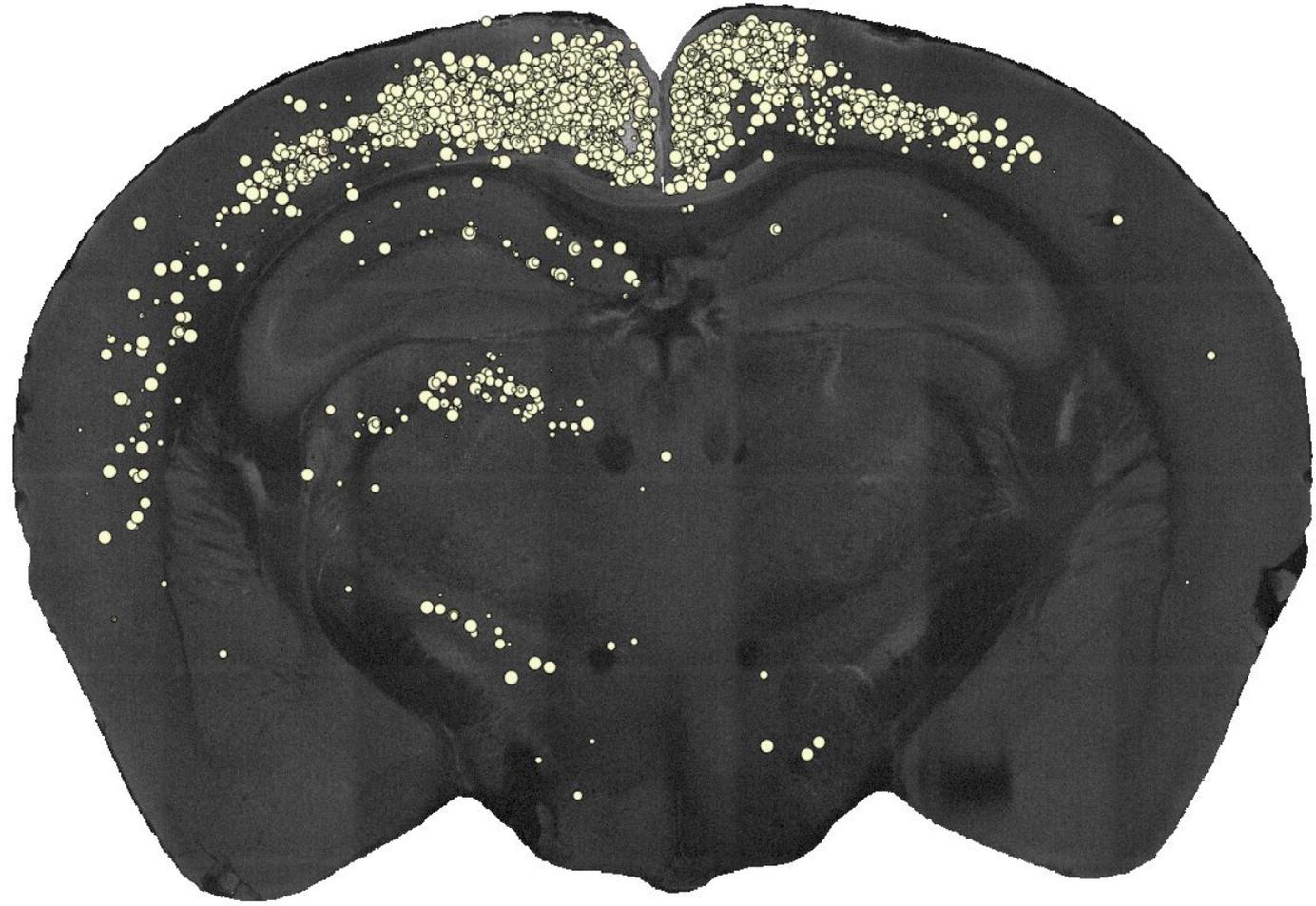
# Cell detection - classification



# Cell detection - classification

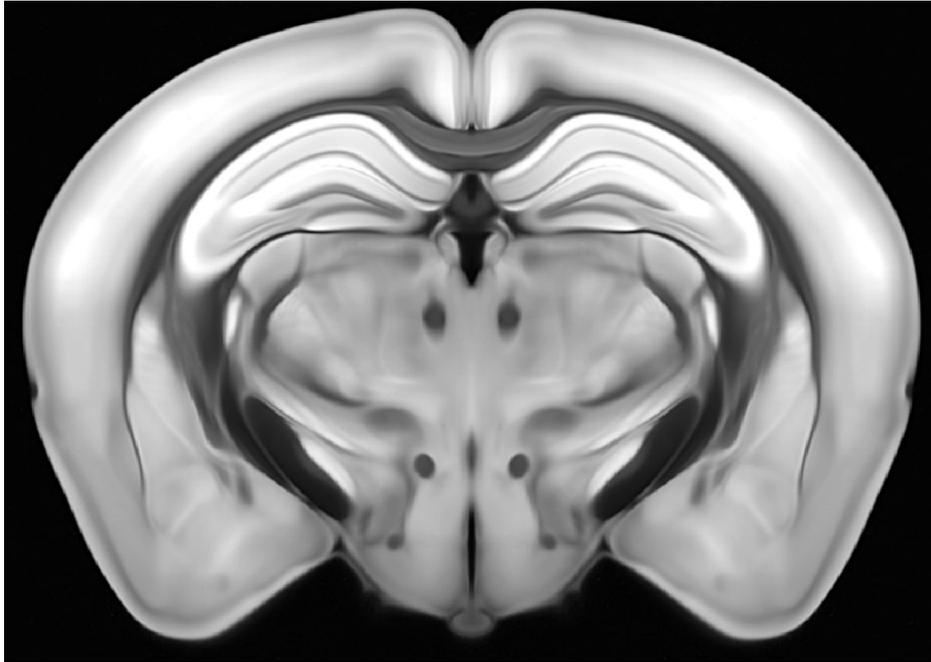


# Brain segmentation



# Registration

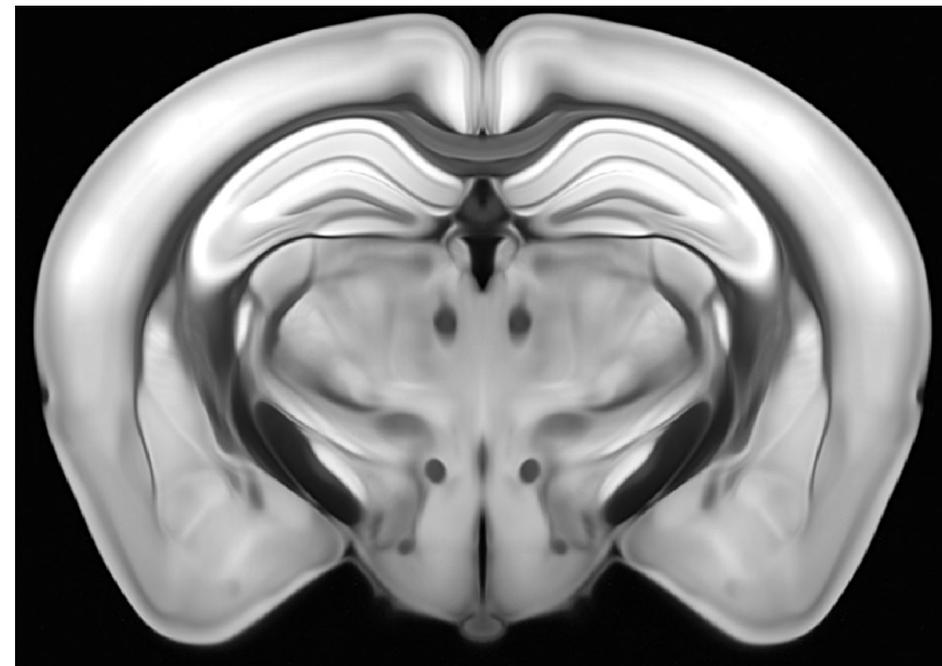
Allen mouse brain reference atlas



Reference image

# Registration

Allen mouse brain reference atlas

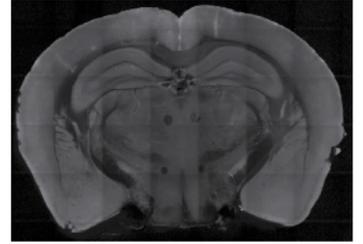
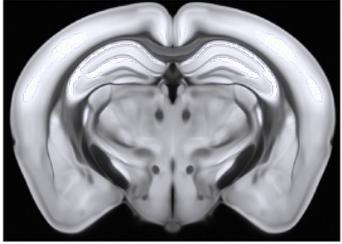


Reference image

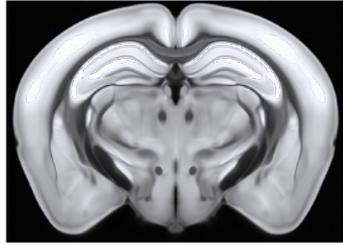


Annotation image

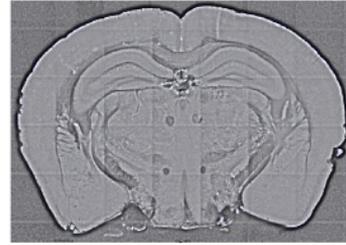
# Registration



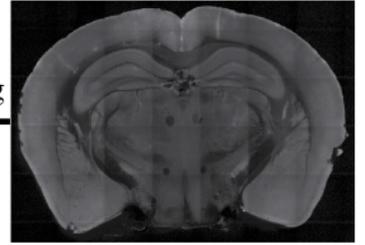
# Registration



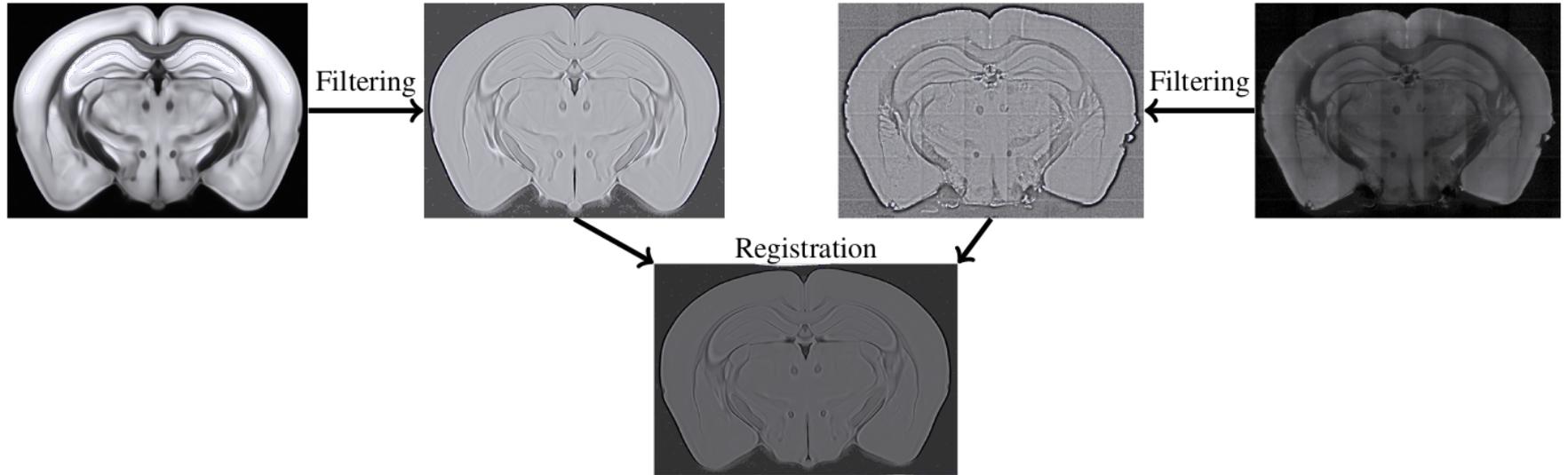
Filtering



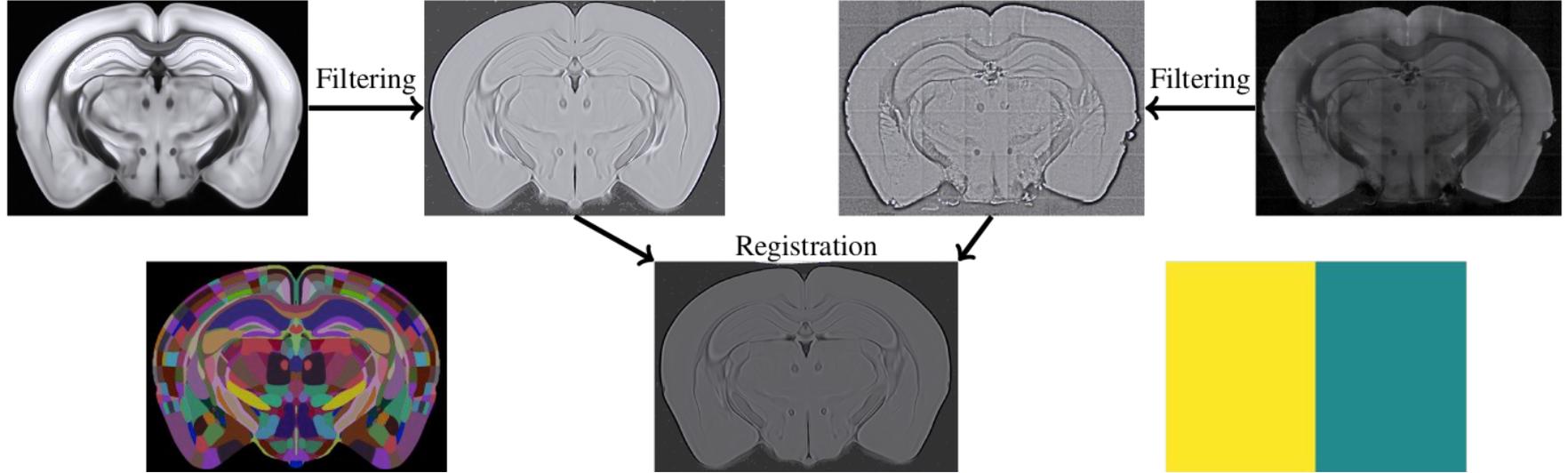
Filtering



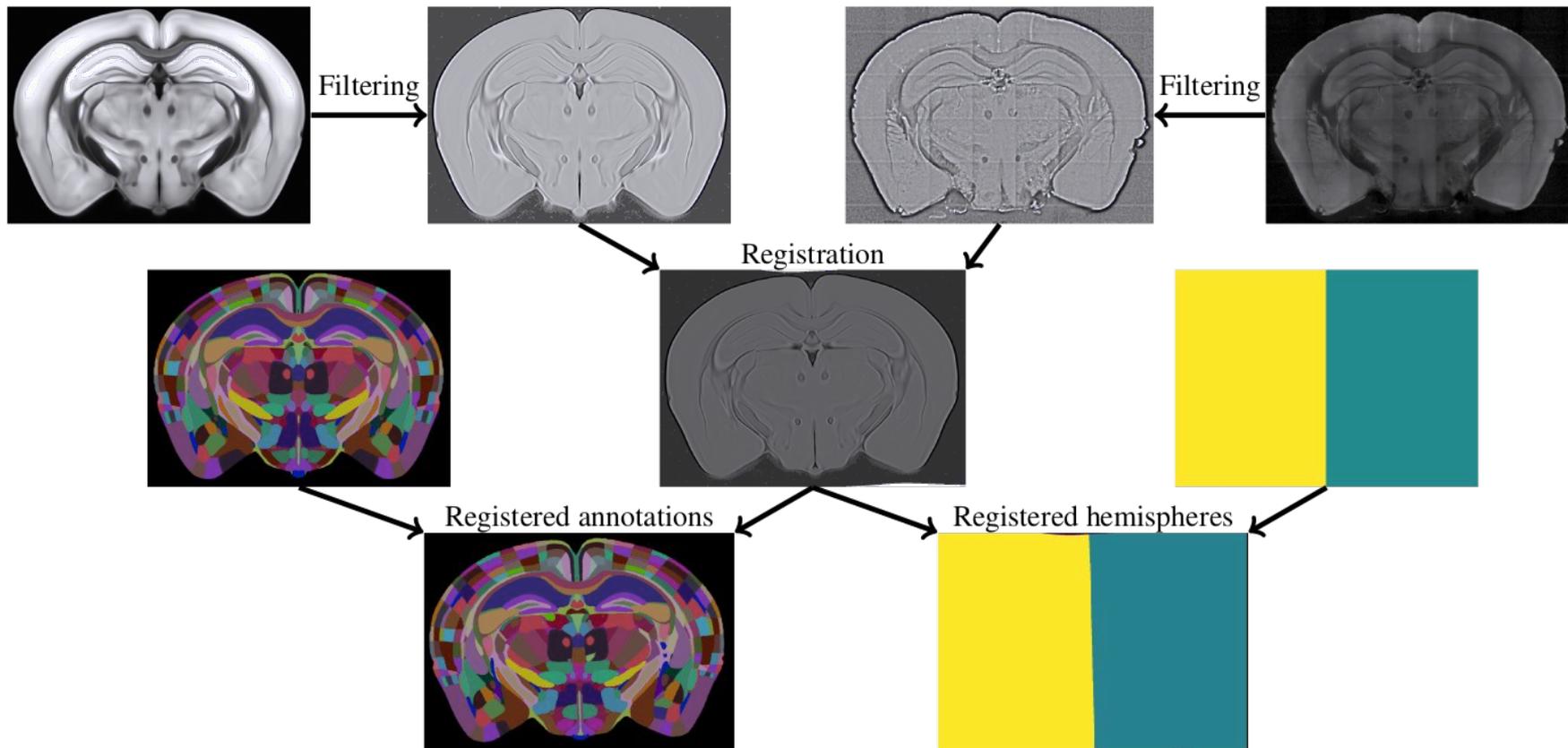
# Registration



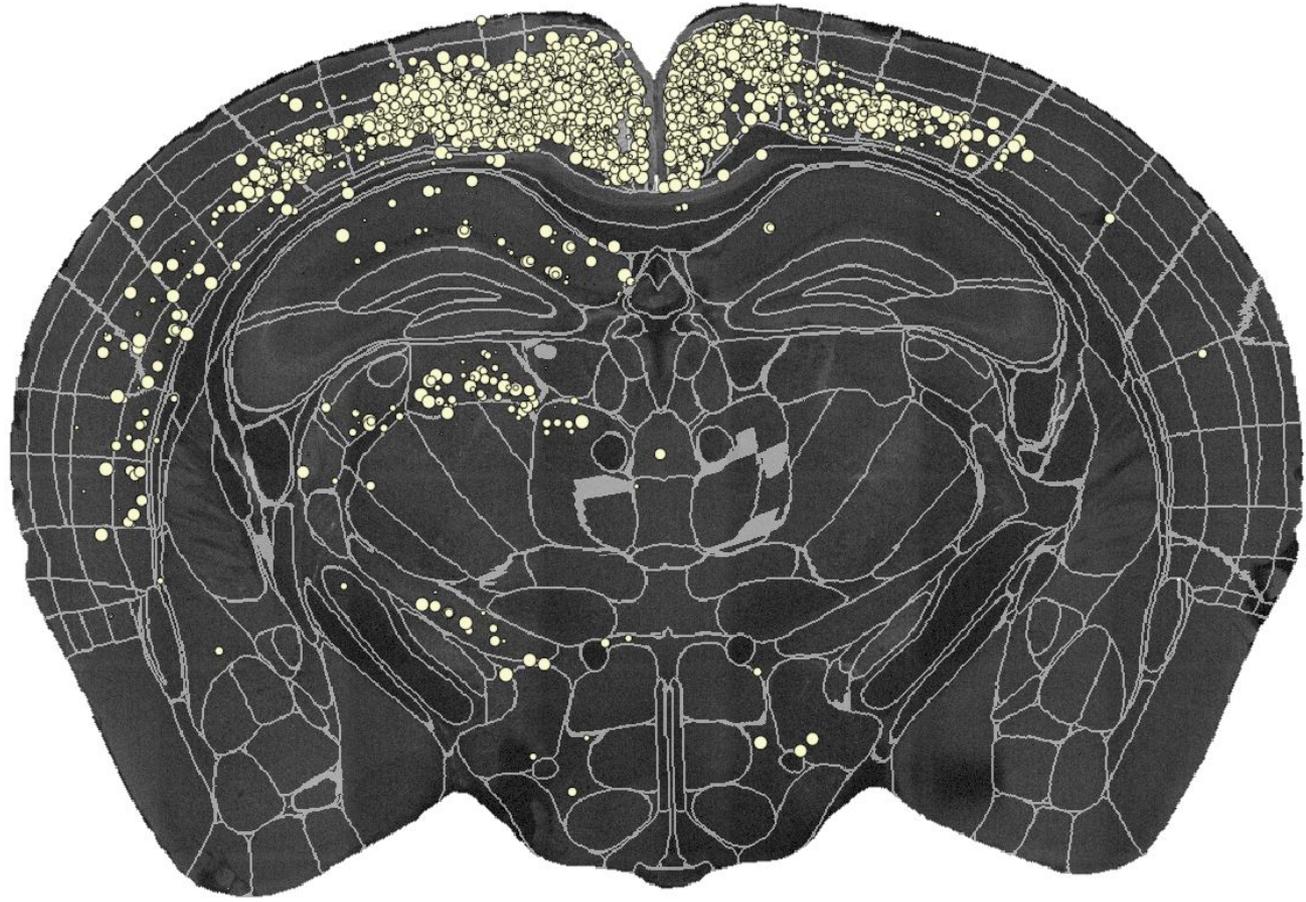
# Registration



# Registration



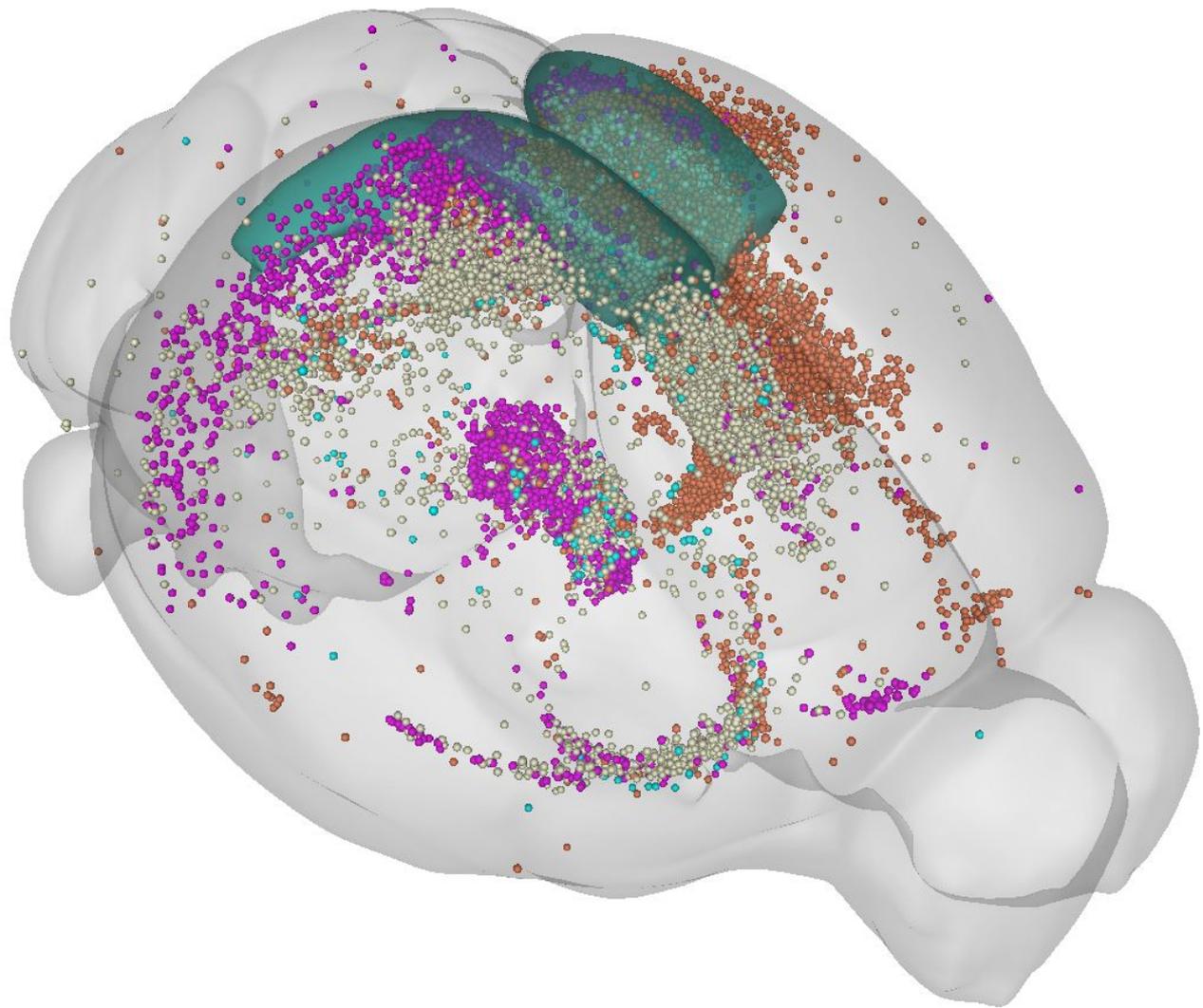
# Registration



# Registration

Brain structure name	Left hemisphere count	Right hemisphere count	Total count
Retrosplenial area, ventral part, layer 5	1853	814	2667
Lateral dorsal nucleus of thalamus	1541	0	1541
Retrosplenial area, ventral part, layer 2/3	163	686	849
Retrosplenial area, dorsal part, layer 5	561	82	643
Retrosplenial area, dorsal part, layer 2/3	194	245	439
Ventral anterior-lateral complex of the thalamus	412	0	412
Anterior cingulate area, dorsal part, layer 5	340	40	380
Anteroventral nucleus of thalamus	374	0	374
...	...	...	...

# Registration

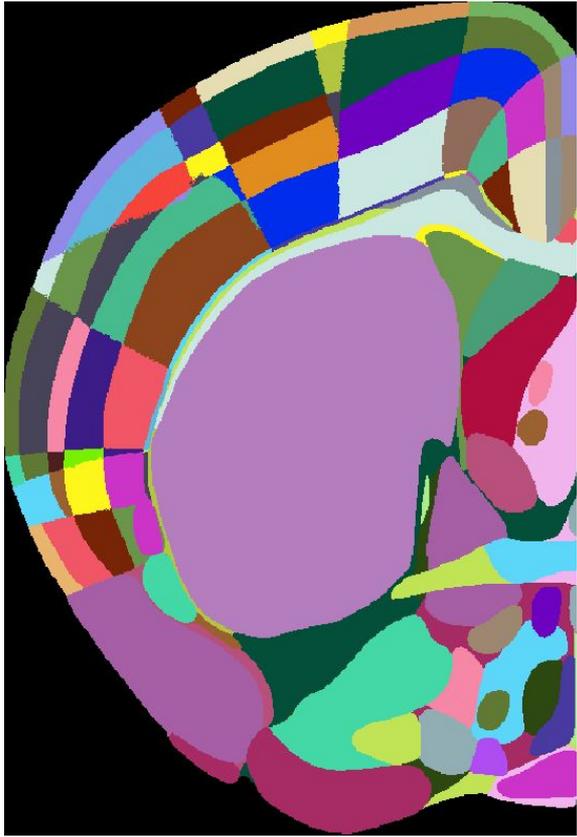


# Brain atlases - mouse



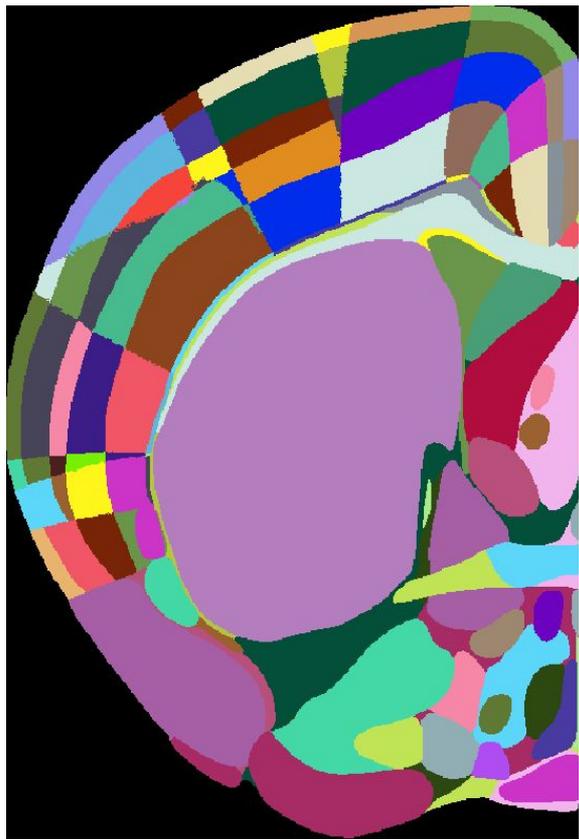
Allen Mouse Brain CCFv3  
(Wang et al., 2020)

# Brain atlases - mouse



Allen Mouse Brain CCFv3  
(Wang et al., 2020)

# Brain atlases - mouse

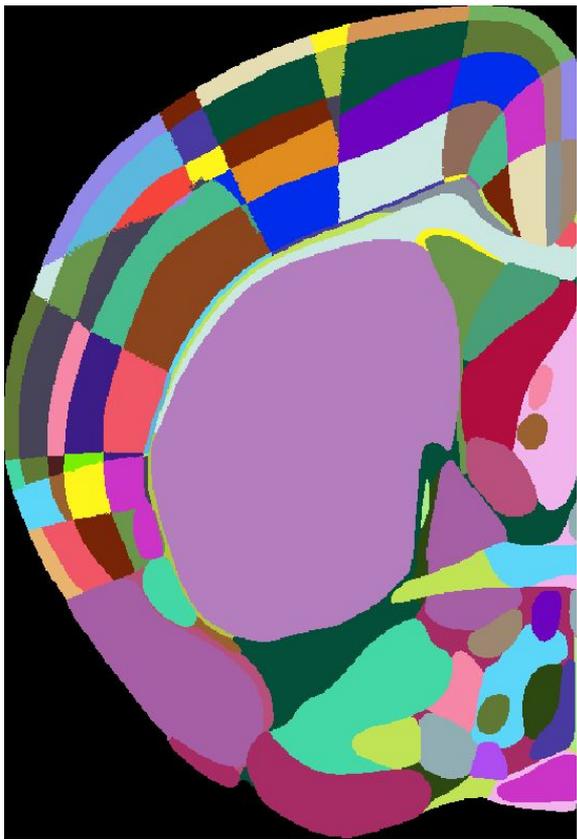


Allen Mouse Brain CCFv3  
(Wang et al., 2020)



Enhanced and Unified Mouse  
Brain Atlas (Chon et al., 2019).

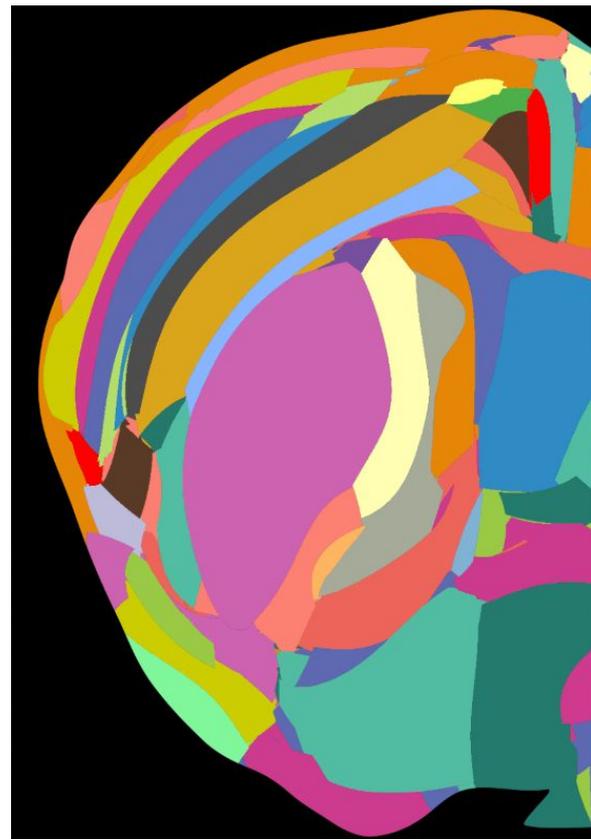
# Brain atlases - mouse



Allen Mouse Brain CCFv3  
(Wang et al., 2020)

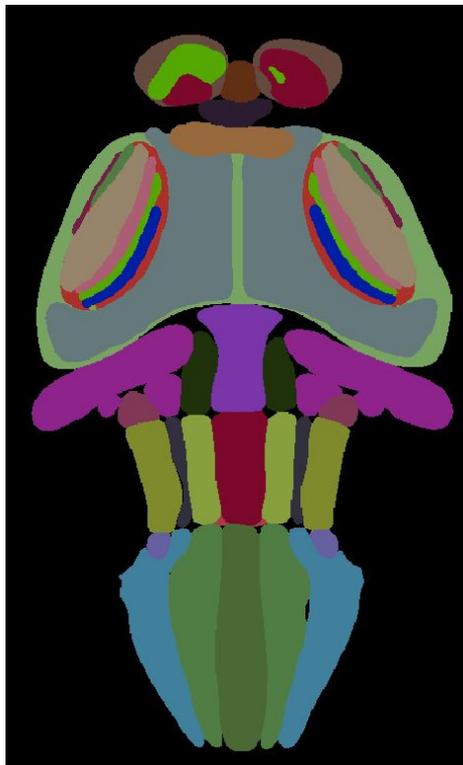


Enhanced and Unified Mouse  
Brain Atlas (Chon et al., 2019).

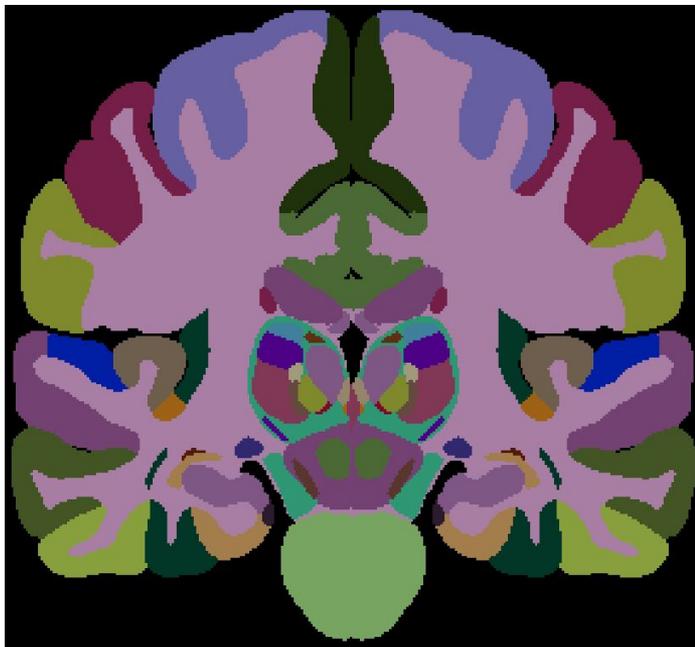


Molecular atlas of the adult  
mouse brain (Ortiz et al., 2020)

# Brain atlases - other species



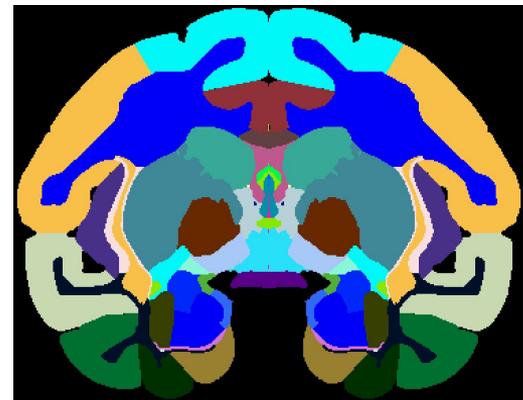
Max Planck Larval Zebrafish Atlas  
(Kunst et al., 2019)



Allen human atlas  
(Ding et al., 2020)

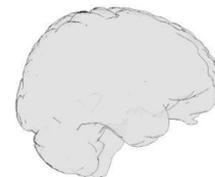
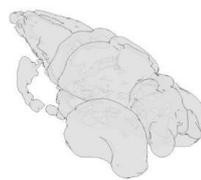


Waxholm Space atlas of the Sprague  
Dawley rat brain (Papp et al., 2014).



NeuroNames macaque atlas  
(Rohlfing et al., 2012)

# BrainGlobe Atlas API



## Python API

```
from bg_atlasapi.bg_atlas import BrainGlobeAtlas
atlas = BrainGlobeAtlas("allen_mouse_25um")

# reference image
reference_image = atlas.reference
print(reference_image.shape)
# (528, 320, 456)

# hemispheres image (value 1 in left hemisphere, 2 in right)
hemispheres_image = atlas.hemispheres
print(hemispheres_image.shape)
# (528, 320, 456)

from pprint import pprint
VISp = atlas.structures["VISp"]
pprint(VISp)
# {'acronym': 'VISp',
#  'id': 385,
#  'mesh': None,
#  'mesh_filename': PosixPath('/home/user/.brainglobe/allen_mouse_25um_v0.3/meshes/385.obj'),
#  'name': 'Primary visual area',
#  'rgb_triplet': [8, 133, 140],
#  'structure_id_path': [997, 8, 567, 688, 695, 315, 669, 385]}
```

## Command line interface

```
adam@garfield: ~
File Edit View Search Terminal Help
(brainglobe) adam@garfield:~$ brainglobe list

Brainglobe Atlases

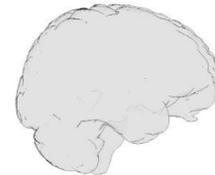
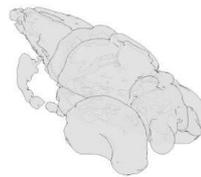


| Name                | Downloaded | Local version | Latest version |
|---------------------|------------|---------------|----------------|
| allen_human_500um   | ✓          | 0.1           | 0.1            |
| ngin_zfish_1um      | ✓          | 0.3           | 1.0            |
| allen_mouse_100um   | ✓          | 1.1           | 1.1            |
| allen_mouse_50um    | ✓          | 0.3           | 0.3            |
| kin_unified_25um    | ✓          | 0.1           | 0.1            |
| allen_mouse_10um    | ✓          | 1.1           | 1.1            |
| example_mouse_100um | ✓          | 1.0           | 1.1            |
| allen_mouse_25um    | ✓          | 1.1           | 1.1            |



(brainglobe) adam@garfield:~$ brainglobe update -a allen_mouse_10um
allen mouse atlas (res. 10um)
From: http://www.brain-map.com (Wang et al 2020, https://doi.org/10.1016/j.cell.2020.04.007 )
bg_atlasapi: allen_mouse_10um is already updated (version: 1.1)
(brainglobe) adam@garfield:~$
```

# BrainGlobe Atlas API



## Currently implemented atlases

- The [Allen Mouse Brain Atlas](#) (10, 25, 50 and 100 micron)
- The [Allen Human Brain Atlas](#) (100 micron)
- The [Max Planck Zebrafish Brain Atlas](#) (1 micron)
- The [Enhanced and Unified Mouse Brain Atlas](#) (10, 25, 50 and 100 micron)
- The [Smoothed Kim et al. Mouse Brain Atlas](#) (10, 25, 50 and 100 micron)

# BrainGlobe

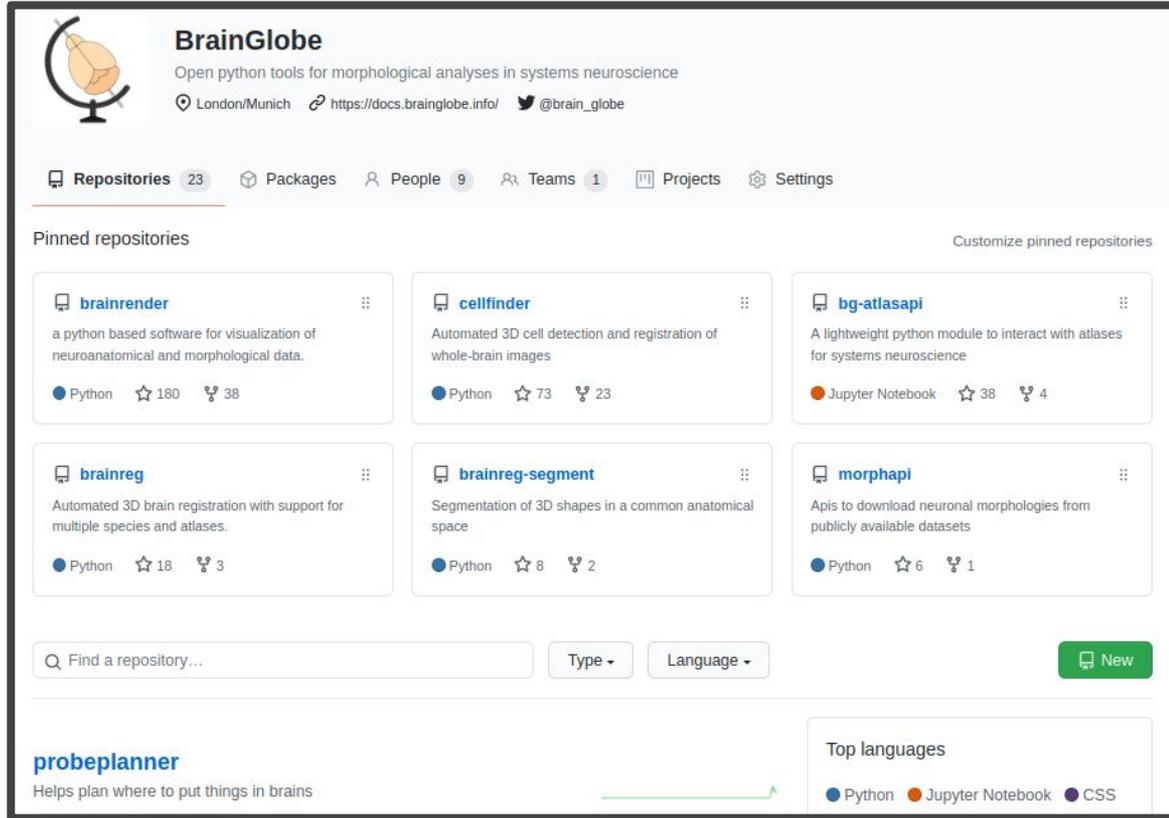
## Building an open-source community

### Currently

- 23 repositories
- 17 total contributors
- ~2.5k downloads/month

### Future

- New contributors
- Biologists adopt tools
- Developers use core tools



The screenshot displays the GitHub profile for BrainGlobe, which is described as "Open python tools for morphological analyses in systems neuroscience". The profile includes a location of London/Munich, a website link to https://docs.brainglobe.info/, and a Twitter handle @brain\_globe. The navigation bar shows 23 repositories, 9 people, 1 team, and settings. The pinned repositories section features:

- brainrender**: A Python-based software for visualization of neuroanatomical and morphological data. 180 stars, 38 forks.
- cellfinder**: Automated 3D cell detection and registration of whole-brain images. 73 stars, 23 forks.
- bg-atlasapi**: A lightweight Python module to interact with atlases for systems neuroscience. 38 stars, 4 forks.
- brainreg**: Automated 3D brain registration with support for multiple species and atlases. 18 stars, 3 forks.
- brainreg-segment**: Segmentation of 3D shapes in a common anatomical space. 8 stars, 2 forks.
- morphapi**: Apis to download neuronal morphologies from publicly available datasets. 6 stars, 1 fork.

At the bottom, there is a search bar for repositories, a "New" button, and a "Top languages" section showing Python, Jupyter Notebook, and CSS.

# Acknowledgements



## Cell detection

Charly Rousseau<sup>1,2</sup>  
Christian Niedworok<sup>1</sup>  
Sepiedeh Keshavarzi<sup>1</sup>  
Chryssanthi Tsitoura<sup>1</sup>  
Lee Cossell<sup>1</sup>  
Molly Strom<sup>1</sup>  
Troy Margrie<sup>1</sup>

## 3D Atlas registration

Charly Rousseau<sup>1,2</sup>  
Christian Niedworok<sup>1</sup>  
Troy Margrie<sup>1</sup>

## Funding



## 3D Visualisation

Federico Claudi<sup>1</sup>  
Luigi Petrucco<sup>2,3</sup>  
Ruben Portuges<sup>2,3,4</sup>  
Troy Margrie<sup>1</sup>  
Tiago Branco<sup>1</sup>

## Atlas API

Federico Claudi<sup>1</sup>  
Luigi Petrucco<sup>3,4</sup>  
Ruben  
Portuges<sup>3,4,5</sup>  
Troy Margrie<sup>1</sup>  
Tiago Branco<sup>1</sup>

1) Sainsbury Wellcome Centre, UCL

2) Sorbonne Université, Institut du Cerveau - Paris Brain Institute - ICM, Inserm, CNRS

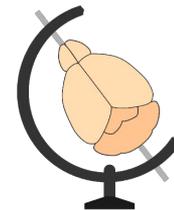
3) Institute of Neuroscience, Technical University of Munich

4) Max Planck Institute of Neurobiology

5) Munich Cluster for Systems Neurology (SyNergy)

# Software

All available as part of the BrainGlobe computational neuroanatomy software suite:



## Cell detection cellfinder

 [github.com/brainlobe/cellfinder](https://github.com/brainlobe/cellfinder)

 [@findingcells](https://twitter.com/findingcells)

Tyson, A. L., Rousseau, C. V., Niedworok, C. J., et al. (2021) "A deep learning algorithm for 3D cell detection in whole mouse brain image datasets" bioRxiv, [doi.org/10.1101/2020.10.21.348771](https://doi.org/10.1101/2020.10.21.348771)

 [github.com/brainlobe](https://github.com/brainlobe)

[docs.brainlobe.info](https://docs.brainlobe.info)

 [@brain\\_globe](https://twitter.com/brain_globe)

[gitter.im/BrainGlobe](https://gitter.im/BrainGlobe)

## 3D atlas registration brainreg

 [github.com/brainlobe/brainreg](https://github.com/brainlobe/brainreg)

## 3D visualisation brainrender

 [github.com/brainlobe/brainrender](https://github.com/brainlobe/brainrender)

Claudi, F., et al. (2021) "Visualizing anatomically registered data with Brainrender" [doi.org/10.7554/eLife.65751](https://doi.org/10.7554/eLife.65751)

## Atlas API BrainGlobe Atlas API

 [github.com/brainlobe/bg-atlasapi](https://github.com/brainlobe/bg-atlasapi)

Claudi, F., Petrucco, L., Tyson, A. L., et al. (2020) "BrainGlobe Atlas API: a common interface for neuroanatomical atlases" Journal of Open Source Software, v5(54), 2668 [doi.org/10.21105/joss.02668](https://doi.org/10.21105/joss.02668)

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 [@adamltyson](https://twitter.com/adamltyson)