



#### Introduction to the Brain Image Library: Designated Repository for NIH BRAIN Initiative Microscopy Data



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www.brainimagelibrary.org

bil-support@psc.edu

### Workshop Agenda

- What is the Brain Image Library:
  - The mission and scope of the archive
  - How to access/getting started
  - Who to contact/how to get additional information
  - Input on use cases/planned future features
- Exploring data in BIL:
  - Finding, accessing, downloading and using contributed data
  - Tools to interact with the data:
  - Input on use cases/planned future features
- Contributing data to BIL:
  - How to submit your image data to BIL
  - Input on data contribution/pre-submission processing
- Open time with BIL Staff



# The Brain Image Library

**<u>Mission</u>**: National public resource enabling researchers to deposit, analyze, mine, share and interact with microscopy datasets of the brain.

#### Scope:

- Permanent repository for high-quality brain microscopy datasets
  - Whole brain images of mouse, rat, other mammals and model organisms
  - Targeted experiments Including connectivity between cells and spatial transcriptomics ( \*FISH)
  - Historical collections
- Provide HPC computing capability local to the data for pre-submission data processing and post-submission exploration
  - Enclave access to pre-release data
  - Research access to restricted-access, secured data
- Provide user access and support







University of Pittsburgh

#### Designated repository for NIH BRAIN Initiative Data

- Requirements for investigators outlined in NIH NOT-MH-19-010 Notice of Data Sharing **Policy for the BRAIN Initiative**
- BRAIN Initiative applications after March 1, 2020, are required to share the data they • collect
- General expectation is that data will be • submitted to the archives every 6 months
- Data will be shared with the research community no later than:
  - When papers using the data have been published
  - The end of the award period

#### Notice of Data Sharing Policy for the BRAIN Initiative

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ational Eye Institute (NEI)
ational Institute on Aging (NIA)
ational Institute on Alcohol Abuse and Alcoholism (NIAAA)
ational Institute of Biomedical Imaging and Bioengineering (NIBIB)
unice Kennedy Shriver National Institute of Child Health and Human Development (NICHD)
ational Institute on Deafness and Other Communication Disorders (NIDCD)
ational Institute on Drug Abuse (NIDA)
ational Institute of Neurological Disorders and Stroke (NINDS)

#### National Center for Complementary and Integrative Health (NCCII Purpose

Initiative. Specifically, this Notice clarifies the expectation that applicants to BRAIN Initiative funding opportunity announcements; 1) submit their data to one of the BRAIN data archives for ents in the Resource Sharing Plan as further detailed below; and 3) include costs attributed to data preparation and submission to a data archive i sharing: 2) include specific required grant application

#### **BRAIN Initiative Overvier**

The Brain Research through Advancing Inr and application of inprovative technologies, researchers will be able to produce a new dynamic picture of the brain that, for the first time, will show how individual cells and complex neuro circuits interact in both time and space. It is expected that the application of these new tools and technologies will ultimately lead to new ways to treat and prevent brain disorder

NIH is one of several federal agencies involved in the BRAIN Initiative. Planning for the NIH component of the BRAIN initiative is guided by the long-term scientific plan. "BRAIN 2025: A Scientific Vision." which details seven high-priority research areas and calls for a sustained federal commitment of S4.5 billion over 12 years. This Notice and related FOAs issued as part o the BRAIN initiative are based on careful consideration by the NIH of the recommendations of the BRAIN 2025 Report, and input from the NIH BRAIN Multi-Council Working Group Videocasts of the NIH BRAIN Multi-council Working Group are available athttp://www.braini

To enable rapid progress in development of new technologies as well as in theory and data analysis, the BRAIN Initiative encourages collaborations b from statistics, physics, mathematics, engineering, and computer and information science

#### BRAIN Initiative Data Sharing

The BRAIN 2025 report calls for establishing platforms for sharing data related to the BRAIN Initiative. In response, NIH has released RFAs related to data archives (RFA-MH-1 145 and RFA-MH-17-255), data standards (RFA-MH-19-146 and RFA-MH-17-256), and software to integrate and analyze data (RFA-MH-19-147 and RFA-MH-17-257). Each of those effort is focused on data from a "sub-domain" which are usually defined by the data collection methodology

Data archives that have been established include

1) The Neuroscience Multi-omic Data Archive (https://nemoarchive.org/about.php, R24MH114788) to hold data from -omics experiment: (www.brainimagelibrary.org/index.html. R24MH114793) to hold microscopy data

3) Data Archive for the BRAIN Initiative (https://dabi.loni.usc.edu, R24MH114796) to hold data related to human electrophysiology experiments

4) OpenNeuro (https://openneuro.org/, R24MH117179) to hold magnetic resonance imaging dat

5) Block and Object Storage Service (https://bossdb.org/, R24MH114785) to hold electron microscopy data

A few additional data archives in distinct sub-domains are expected to be funded. The complete list of BRAIN Initiative infrastructure awards can be found here. The awardees for the BEA







#### **Publicly Accessible Data**

# Over 1100 public mouse brain datasets are available at BIL



Techniques used to generate datasets in BIL





#### Notable Data Contributors

Hongkui Zeng Allen Institute for Brain Science





Hongwei Dong University of California

**Guoqiang Bi** University of Science and Technology China



Z. Josh Huang Cold Spring Harbor Laboratory





X. William Yang University of California Los Angeles

**Pavel Osten** Cold Spring Harbor Laboratory









#### Exemplar Dataset



**Contributor:** Hongkui Zeng Allen Institute for Brain Science

We feature in this poster one of the 34 fMOST datasets received from the Zeng U19 project at the Allen Institute for Brain Science. Imaging was performed at Huazhong University of Science and Technology. This dataset represents wholebrain imaging of a transgenic mouse (Gnb4-IRES2-CreERT2/wt;AI82(TITL-GFP)/Ai140(TIT2L-GFP-ICL-TTA2)) in which a small sampling of neurons, including claustrum projection neurons, are fluorescently labeled [4]. The green channel shows where GFP is expressed, and can be used for tracing these neurons. It is important that only a sparse subset be labeled; otherwise, it would become much more difficult to trace individual neurons with high confidence. For this specimen BIL currently has available 4 neuron morphologies in .swc format, of which we illustrate one (17781\_00001.swc). Potentially, thousands of other morphologies could be reconstructed from this datase



Image 05920 (of 11031), red channel, 34721 x 54600 pixels x, y resolution = 0.35µm, z spacing = 1µm; field-of-view: 19.11 x 12.15 mm.



Combined red and green channels. Zoom factor = 1.28; field-of-view: 14.93 x 9.47 mm.

Image 05920 (of 11031), green channel, 34722x54600 pixels.

fMOST (fluorescence micro-optical sectioning tomography) dataset showing one of several neuron morphologies reconstructed from this dataset.



Zoom factor = 5.24; field-of-view: 3.64 x 2.36 mm



Zoom factor = 10.51; field-of-view: 1.82 x 1.18 mm.



Zoom factor = 2.61; field-of-view: 7.32 x 4.73 mm.



Zoom factor = 34.00; field-of-view: 0.56 x 0.38 mm. (Pixel resolution is the same as the acquisition resolution - 0.35 x 0.35 µm.

x, y resolution = 0.35µm, z spacing = 1µm; field-of-view: 19.11 x 12.15 mm





Axons and dendrites of an individual labeled neuron can be traced by starting at a soma and following the green lines through successive images in z. This figure was generated from the 17781\_00001.swc file uploaded to BIL [7] and rendered with the SharkViewer tool [8]





#### **Computational Services**

 Computational Systems to process and explore BIL data in-place available at no charge for open research and to support courses

#### BIL Hardware:

- VM computational system:
  - Remote Desktop Capable
  - NVIDIA V100 GPU's (shared)
- 4 PB storage (will grow to 10PB)
- Tape drives capable of receiving data on LTO 7/8 tapes
- "Brainball": 30TB portable drive systems
- Dedicated data transfer nodes

#### Bridges (HPC System):

- 28,628 Intel Xeon cores
- GPUS:
  - 64 NVIDIA K80
  - 64 NVIDIA P100
- 17 PB storage:
  - 10PB persistent
  - 7.3PB node-local

- Nodes:
  - 800x128GB RAM
  - 42x3TB RAM
  - 4x12TB RAM
- 88 NVIDIA Volta GPUs:
  - *9 Nodes,* with 8 NVIDIA V100
  - 1 NVIDIA DGX-2, 16 Volta with 2.4 TB/s NVSwitch



#### Coming Soon:

• Bridges-II: New \$10,000,000 HPC System arriving in June 2020

# Web Portal Hosting

- Access data in unique ways
  - Access data in-place
  - Commercial software capable
  - Can take advantage of HPC to compute/deliver custom results on-the-fly

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#### Help Desk Services

- Contact
  - Please email: bil-support@psc.edu.
  - Time-critical, call PSC hotline at: 412-268-6350.
  - Please note that we are located in the Eastern Time zone.
- Help Desk Scope:
  - Answer questions
  - Facilitate software installation to enable processing and visualization.
  - Networking and data transfer support
    - Diagnose and facilitate resolution of last mile and routing problems



#### How to get started – Important Documentation

- BIL website: <u>www.brainimagelibrary.org</u>
- Locating public BIL data:
  - <u>http://www.brainimagelibrary.org/datasets.html</u>
- Downloading public BIL data:
  - <u>http://www.brainimagelibrary.org/download.html</u>
- Compute on the data without downloading:
  - <u>http://www.brainimagelibrary.org/compute.html</u>
- Contribute data to BIL:
  - <u>http://www.brainimagelibrary.org/submission.html</u>
- Help:
  - <u>http://www.brainimagelibrary.org/contact.html</u>
  - email: <u>bil-support@psc.edu</u> or call: 412-268-6350



A public resource enabling researchers to deposit, analyze, mine, share and interact with large brain image datasets

The Brain Image Library is supported by the National Institutes of Mental Health of the National Institutes of Health under award number R24-MH-114793. The content is solely the responsibility of the authors and does not necessarily represent the official views of the National Institutes of Health.



## Help Us Meet Your Needs

- How can we make BIL more accessible and usable by the research community?
  - What information would you expect to find within three mouse clicks?
  - What use cases should be supported?
  - What external resources should be linked to? Why?
  - What exploration & visualization software should be supported?
- Training/outreach needs and uses
- Feedback on future directions as well as general feedback



#### **Data Contributors:**

Results:

 Understanding your future computational and data storage needs including tools/software that are needed for your work





# Exploring Data in the Brain Image Library

www.brainimagelibrary.org

bil-support@psc.edu

# Wouldn't it be great if.....

- We had a collection of thousands of large, high-quality, highresolution microscopy datasets of the brain...
- And this collection was attached to significant high performance computing resources...







Watson et. al. https://doi.org/10.1371/journal.pone.0180486

#### Then we might be able to.....

- Confirm that neurons grown in tissue culture are "similar" to neurons found in whole brains. (Do they make the right connections with their neighbors?)
- Have sufficient training and validation sets to generate realistic models of key cellular-level brain structures
- Study variation across genetically similar (inbred) populations
- Develop special purpose maps joining the work from individual laboratories to elucidate broader connectivity patterns of localized circuits.
- Do several other interesting things......



#### Finding Data: BIL Data Inventory

#### www.brainimagelibrary.org Menu: Data Access -> Brain Inventory

#### 🛞 Brain Image Library

About ▼ Data Submission ▼ Data Access ▼ Contact

Search				
bil_uuid	sample_id	organism_type	organism_ncbi_taxonomy_id	transgenetic_line_information
72f3f3e249a07462	Ntsr1_Ai75_M_361838	mouse	NCBI:txid10090	Strain_Name='C57BL6';Cre_Driver_Line='Ntsr1';Reporter
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#### Finding Data: Future

/		About ▼ Data Submission ▼ Data Access ▼ Contact	
		Investigator: Josh Huang Collection:697a07d24603a697 Dataset: 180715_KM001_AdcyapHGfemale_processed	
	$(\mathbf{f})$	Investigator: Hongkui Zeng Collection:7f537a62e521a26a Sample: 712844138 Dataset: mouseID_297974	
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Investigator: Josh Huang Lab Name: Huang Lab Organization: Cold Spring Harbor Laboratory Collection: 697a07d24603a697 Dataset: 180715\_KM001\_AdcyapHGfemale\_processed Channels: 2 Dimension: 11377 x 8557 Size: 6.4G

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Metadata

Brain Image Library

Field	Value
Sample ID:	180715_KM001_AdcyapHGfemale_processed
NCBI Taxonomy ID:	NCBI:txid10090
Location:	/bil/data/69/7a/697a07d24603a697/180715_KM001_AdcyapHGfemale_processed/ Firefox/Chrome/Safari ftp link Explorer/Edge ftp link
Modality:	cell counting
Transgenetic Line:	Strain_Name='C57BL6/Agouti/Swiss'; Pynline_Driver_InjectionSite='AdcyapHG'
Method:	imaging
Organization:	Cold Spring Harbor Laboratory
Collection Name:	697a07d24603a697
Collection Description:	Collection 697a07d24603a697
Technique:	STPT
Collection UUID:	697a07d24603a697
Organism:	mouse

Sample Images from the dataset (click to zoom)

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#### <u>What would you like to see?</u> <u>How would you like to search?</u>

View Info

🍘 Brain Image Librar

**Results:** 

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## Finding Data: File Systems

- First, check the data inventory for the file location
- On BIL computational resources, the file path to the BIL entry is:
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    - where <c1c2> are replaced by the first and second characters of the unique identifier and <c3c4> are replaced by the third and fourth characters of the unique identifier.
  - Example bil\_uuid: abcded0123456789, sample\_name: example\_dataset\_01
    - /bil/data/ab/cd/abcdef0123456789/example\_dataset\_01/
- In the future, DOI's will also be used to directly access datasets



#### Downloading Data

- No Login Required:
  - Globus: bil#download
    - Preferred method, due to length of typical data transfers
    - Restart/resume capable
  - Anonymous FTP: download.brainimagelibrary.org
- Download Considerations:
  - Some BIL datasets can exceed 10TB and may be difficult to download over slow network connections.
    - 10 Terabyte mouse brain @ 1Gbit/sec ~ 1day (25hrs)
    - Same brain would take over 4 days @ 250 Mbits/sec
  - Consider compute/visualize in-place.



- Login to the Globus transfer web service at: www.globus.org.
- Select an Endpoint field
  - Select 'Brain Image Library Download' (bil#download).
- Upon successful connection you will land in the /bil/data directory
- Navigate to the directory containing the dataset(s) that you wish to download.

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- In the other Endpoint field, select your destination server.
- Select the appropriate transfer settings that apply to your download needs, e.g., "preserve source file modification times", "verify file integrity after transfer", etc.

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 Select the dataset(s) and/or files that you wish to download by highlighting their directory and/or file names.

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- Click the arrow button in the direction of the transfer.
- This will schedule your transfer to begin.
- At this point you may log out of the Globus user interface and let the transfer proceed.





- To check on the progress of your transfer, select the Activity tab
- Then select the job corresponding to your transfer.

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Condition	ACTIVE			Cancelled	0	
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- Upon successful completion of your transfer, the status page for your transfer will show 'Condition SUCCEEDED'
- You may also receive an e-mail to notify you if your Globus account settings are configured to do so.



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### Tools for Interacting with Data in-Place

- Rich suite of software available for • use on BIL VMs and Bridges
- VMs capable to bring-your own • commercial software

Singularity

SPICE

Contact us @ <u>bil-support@psc.edu</u> 







# Require Significant Computational Resources?

- Apply for an XSEDE resource allocation on Bridges:
  - There is **NO CHARGE** for accessing these computational resources.
  - Normally a research starter allocation provides up to:
    - 50,000 Regular memory core-hours (50,000 Service Units)
    - 1000 Large Memory terabyte-hours. (1,000 LM Service Units)
    - 2,500 K80 (or 1000 P100) GPU hours (2,500 Service Units)
    - 1500 (Volta,DGX-2) AI GPU hours (1500 Service Units)
  - Research allocations can be for substantially more service units
  - Educational/instructional allocations also available.
  - We can help groups with the process contact: bil-support@psc.edu
- Multiple users can be added to and share a single resource allocation:
  - Pl's collaborators (including non-US) students



# Help Us Meet Your Data Exploration Needs

- How can we make BIL more accessible and usable by the research community?
  - What information would you expect to find within three mouse clicks?
  - What use cases should be supported?
  - What external resources should be linked to? Why?
  - What exploration & visualization software should be supported?
- Training/outreach needs and uses
- Feedback on future directions as well as general feedback





View Info

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# Contributing Data to the Brain Image Library

www.brainimagelibrary.org

bil-support@psc.edu

# Contributing Data to BIL

- There is no-charge to contribute data to BIL or access BIL services.
- BIL can handle (and we encourage!) the submission of very large datasets
  - If you will be submitting more than 100TB of data, please let us know in advance.
- We can also provide letters of collaboration for proposals submitted to funding agencies
- Detailed data submission instructions are at:
  - www.brainimagelibrary.org/submission.html



The Brain Image Library (BIL) is an NIH funded public resource (R24-MH-114793) to analyze, mine, share and interact with large high-resolution brain image datasets. BIL relies on contributions of brain image data from researchers like you. There is no charge for contributing data to the library and a help desk is available to provide data submission assistance.



bil-support@psc.edu
www.brainimagelibrary.org





#### Moving Data to BIL

- Preferred method is over the internet
  - BIL is connected to multiple internet providers, peering networks, and a highspeed link to the internet2 backbone.
  - We can help debug networking issues.
- Special arrangements can be made to receive data on disk or tape
  - BrainBall
    - Portable 30TB RAID system
  - Tape Cartridge:
    - Capability to receive data on LTO7 and LTO8 tapes.





#### **Pre-Submission Processing**

- Processing:
  - Remote desktop VMs
  - HPC Access to Bridges
- Individual and project storage space
- Rich suite of software available on BIL VMs and Bridges
- Contact us @ bil-support@psc.edu





**BIO-FORMATS** 







#### **Contributing Data: Process**

- 1. <u>Set up Submission Account</u>: Use the XSEDE portal to set up a submission account, sending that portal account name to bil-support@psc.edu
- 2. <u>Enter the Submission Portal</u>: Log into the data submission portal.
- **3.** <u>**Create a Collection:**</u> A collection is a wrapper associating image data with metadata.
- **4.** <u>Upload Image Data</u>: Because this is the time-limiting step, we suggest this step be started before uploading metadata for the collection.
- 5. <u>Upload Metadata</u>: Collection metadata can be uploaded in spreadsheet format.
- 6. <u>Validate Data</u>: Validate the data and correct any errors discovered.
- 7. <u>Review and Submit Data</u>: Correct all errors and check all warnings in the validation log prior to submission



• Visit

portal.xsede.org

- Click on:
   Create Account
- Fill in requested information





- Fill in personal information
- Choose a passkey (this is something you make up)
- System will send you email with a verification code
- You will use the verification code together with passkey in the next step to finish creating your account

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- Verify your account using email address, passkey, and verification code
- Choose username
- Choose password (that complies with password rules)

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VERIFY YOUR ACCOU	NT REQUEST	
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CHOOSE YOUR TERAG	GRID USER NAME AND PASSWORD         User names must be between 3 and 8 characters in length, must         contain only lowercase letters and numbers, and must begin with a         letter.         Passwords must be a non dictionary word, at least 8 characters in         length and must contain at least 3 of the following character classes:            • lowercase letters             • uppercase letters             • numbers	



SUBMIT



- Email <u>bil-support@psc.edu</u> with the following:
  - Your XSEDE Portal Account
  - Information about the project that you are associated with (e.g. NIH or NSF Grant number & Title)
- You will receive automated email messages:
  - That you have been added to the system
  - One with your username + instructions to set your password



#### Enter Submission Portal (Step2)

- With a web browser, visit: submit.brainimagelibrary.org
- Log in with your
  - Username
  - Password
- Problems?
  - <u>bil-support@psc.edu</u>
  - 412-268-



The Brain Image Library (BIL) is a public resource enabling researchers to deposit, analyze, mine, share and interact with large brain image datasets. More information about the Brain Image Library can be found on our informational website: www.brainimagelibrary.org. The BIL is supported by the National Institute of Mental Health of the National Institutes of Health under award number R24MH114793.

Please log in or sign up to begin creating collections for publication in the BIL.

University of Pittsburgh



#### Create a Collection (Step 3)

- Menu: New -> Collection
- A "Collection" is a flexible ingest wrapper to tie data and metadata together:
  - One per brain
  - One per experiment
- Creating a collection also creates a "Landing Zone" (a private space for you to upload this data to)
  - /bil/lz/user/collection

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#### Upload Image Data (Step 4)

- Upload endpoints support Globus, rsync, sftp, and scp:
  - upload.brainimagelibrary.org (rsync,sftp,scp)
  - #bil-upload (globus)
- Authentication is required
- Once authenticated, change to the landing-zone directory

#### \$ sftp testuser@upload.brainimagelibrary.org

The authenticity of host 'upload.brainimagelibrary.org (128.182.108.164)' can't be established. ECDSA key fingerprint is 32:cf:46:44:3d:9c:8e:b2:1d:14:03:66:45:0b:11:29. Are you sure you want to continue connecting (yes/no)? **yes** Warning: Permanently added 'upload.brainimagelibrary.org,128.182.108.164' (ECDSA) to the list of known hosts. testuser@upload.brainimagelibrary.org's password: Connected to upload.brainimagelibrary.org. sftp> cd /bil/lz/testuser/1234567890abcdef





# Upload Metadata (Step 5)

- Menu: New-> Descriptive Metadata
- Download and fill out the metadata spreadsheet
  - Be sure to enter in the spreadsheet the collection subdirectory where the data is located.
- Select the collection the metadata is for
- Upload filled-in metadata spreadsheet

Brain Image Library (🛞) New - View - Valida	te → Submit → 👤 ropelews C+Log out
Upload Descriptive Metad	lata Spreadsheet
Download and fill out either the Excel or LibreOffice Calc temp	olate.
Choose a collection and upload your metadata.	
Associated collection:	Ø Open
	Organize · New folder · · · · · · · · · · · · · · · · · · ·
Upload Metadata	<ul> <li>This PC</li> <li>3 D Objects</li> <li>Desktop</li> <li>Occuments</li> </ul>
	Bownloads     V     S       File name:     V     All Files       Open     Cancel

#### Your Descriptive Metadata

+ Create Metadata

Cancel

Descriptive Metadata successfully uploaded

	Collection	Date Created	Last Edited	Locked	Sample Id	Organism Type	Organism Ncbi Taxonomy Id	Transgenetic Line Information	Modality	Method	Technique	Anatomical Structure	r T C
Detail	Tues	10/20/2019 4:20 p.m.	10/20/2019 4:20 p.m.	<b>₽</b>	abcdef	mouse	NCBI:txid10090	_	Anatomy/Morphology	cell type distribution	Ribbon	Whole brain	•
Detail	Whole Brain - September 2018	09/04/2018 5:21 p.m.	03/25/2019 3:29 p.m.	•	abcdef	mouse	NCBI:txid10090	_	Anatomy/Morphology	cell type distribution	Ribbon	Whole brain	-
Detail	Test of new acls	09/11/2018 10:29 p.m.	03/25/2019 3:03 p.m.	ſ	abcdef	mouse	NCBI:txid10090	_	Anatomy/Morphology	cell type distribution	Ribbon	Whole brain	
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# Validate Data (Step 6)



- Menu: Validate -> Validate Collection
- Checks performed on files in directories identified in the metadata spreadsheet (+ subdirectories of those directories):
  - Zero length file
  - For image files, bioformats can successfully read image file header information
  - Removal of (unix) problematic special characters from filenames (e.g. space, quotes asterisk, non-printable characters)
- Checks not performed:
  - Any directory that is not identified in the metadata spreadsheet



### Submit Data (Step 6)

- Submitting data will make the data publicly available
- Current:
  - To submit data and make the data publicly available, send email to <u>bil-support@psc.edu</u>
  - Additional manual checks are performed
- Future:
  - Automated submit with embargo periods



#### Data Contribution Workflow



# Help Us Meet Your Data Contribution Needs

- To submit data that you have now, how much disk space would you need?
- What software and visualization tools are needed for your pre-analysis work?
- What validation/QA/QC feedback would you find useful for your datasets?
- What higher-level use cases might your data support if similar datasets were found in BIL?
- Training/outreach needs
- Feedback on future directions as well as general feedback







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#### For More Information

- Contact us at:
  - Email: <u>bil-support@psc.edu</u>
  - Telephone : 412-268-6350
- Web Portals
  - Contributor Portal and Data Upload
    - submit.brainimagelibrary.org (Metadata)
    - upload.brainimagelibrary.org (Image Data)
  - Data Access
    - download.brainimagelibrary.org
  - Informational Website
    - www.brainimagelibrary.org



CONTACT

bil-support@psc.edu
www.brainimagelibrary.org

Acknowledgements: BIL is operated as a partnership between the Biomedical Applications Group at the PSC, the Center for Biological Imaging at the University of Pittsburgh and the Molecular Biosensor and Imaging Center at Carnegie Mellon University. BIL is supported by the National Institute of Mental Health under award number R24MH114793. The content is solely the responsibility of the authors and does not necessarily represent the official views of the National Institutes of Health. The Bridges supercomputing system at the PSC was acquired through NSF award ACI-1445606 [2]. User access to Bridges is supported through the Extreme Science and Engineering Discovery Environment (XSEDE) which is supported through NSF grant OCI-1053575 [5]. Any opinions findings and conclusions or recommendations expressed in this material are those of the authors and do not necessarily reflect the views of the NSF. The authors wish to acknowledge the Brain Initiative Cell Census Network (BICCN) for their data contributions to BIL.



