



## **The EBRAINS infrastructure:**

# integrated services addressing current and future challenges in brain research

Workshop EBRAINS – a platform for collaboration in digital neuroscience <u>Centro para el Desarrollo Tecnológico Industrial</u> <u>Madrid Spain</u> November 12, 2019 Jan Bjaalie, University of Oslo U1O

Director, HBP Infrastructure Operations Platform leader, HBP Neuroinformatics Co-chair, International Brain Initiative (IBI) Head, Norwegian Neuroinformatics Node (INCF Norway)





## **Advancing neuroscience:**

## Collaborative data sharing, reproducible research, and workflows from data to models of brain function

Workshop EBRAINS – a platform for collaboration in digital neuroscience <u>Centro para el Desarrollo Tecnológico Industrial</u> <u>Madrid Spain</u> November 12, 2019 Jan Bjaalie, University of Oslo UiC

Director, HBP Infrastructure Operations Platform leader, HBP Neuroinformatics Co-chair, International Brain Initiative (IBI) Head, Norwegian Neuroinformatics Node (INCF Norway)



#### «Humans are brains»



- "Understanding the brain"
- "Healing the brain"
- "Modeling the brain"

### Basic science, Health, Technology

The strongest driving force for neuroscience today is the societal demand for treatments for brain disease

- Cost of dementia alone will surpass cost of all of cancer
- 2015: number of people in age group 60-64 surpassed number in age group 20-24



# Human Brain Project (HBP): the first in a wave of large brain projects



#### Start of Brain Projects

		Human Brain Project
		US BRAIN
		Japanese Brain/MINDS
		Israel Brain Technologies
		Korean Brain Initiative
		Australian Brain Alliance
Allen Institue for Brain Science		China Brain Project
Blue Brain Project		
	Human Connectome Project	International Brain Laboratory
		International Brain Initiatives
		Canada Brain Research Strategy
2003 2004 2005 2006 2007 2008	8 2009 2010 2011 2012 2013 2	2014 2015 2016 2017 2018 2019 2020 2021 2022

4



- At the interface of neuroscience, computing, technology and brain medicine
- Research & Infrastructure development
- Driven by "co-design projects" and "use cases"

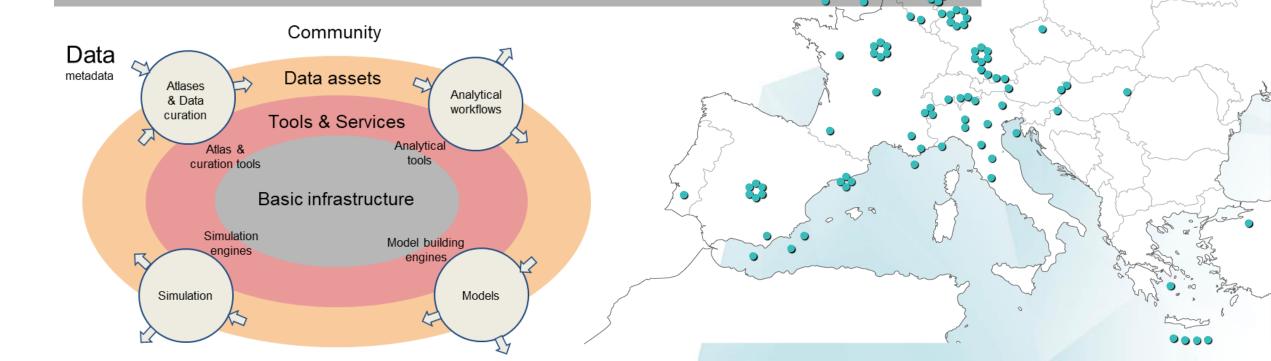
#### Missions:

- >Understanding the multi-level brain organization in space and time
- Transfer the acquired knowledge to brain-derived applications in health, computing, and technology
- > Developing and maintaining a European HBP Research Infrastructure
- >Create a benefit to the European society and ensure an ethical approach in research

H

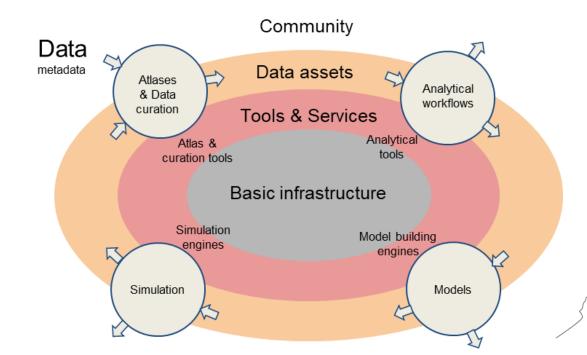


The Human Brain Project Flagship has developed and will release a federated ICT infrastructure – a research e-infrastructure for brain research – helping the research community collect, analyse, share, integrate and model data about the brain with the aim of better understanding the functioning of the human brain and its diseases.









#### The EBRAINS infrastructure will include

- FAIR data services
- Brain atlases for data integration
- Brain modeling and simulation
- Closed-loop AI and robotics
- Medical brain activity data
- Interactive High-Performance Computing and Neuromorphic Computing





### European Brain ReseArch INfrastructureS

- > supporting a wide range of research methodologies
- > facilitating collaborative research and data sharing
- integrating a large variety of ICT services addressing current and future challenges in the neurosciences
- > providing working solutions for experimental, computational, and clinical neuroscientists

https://ebrains.eu Pre-launch October 18, 2019

The Project and work-in-progress

www.ebrains.eu Powered by HBP - Working Solutions for the scientific community





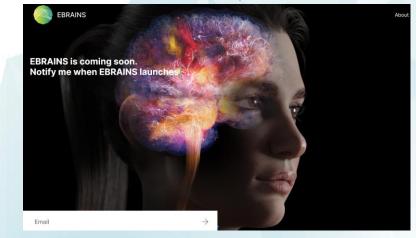
#### European Brain ReseArch INfrastructureS

- > supporting a wide range of research methodologies
- > facilitating collaborative research and data sharing
- integrating a large variety of ICT services addressing current and future challenges in the neurosciences
- > providing working solutions for experimental, computational, and clinical neuroscientists

https://ebrains.eu Pre-launch October 18, 2019

The Project and work-in-progress

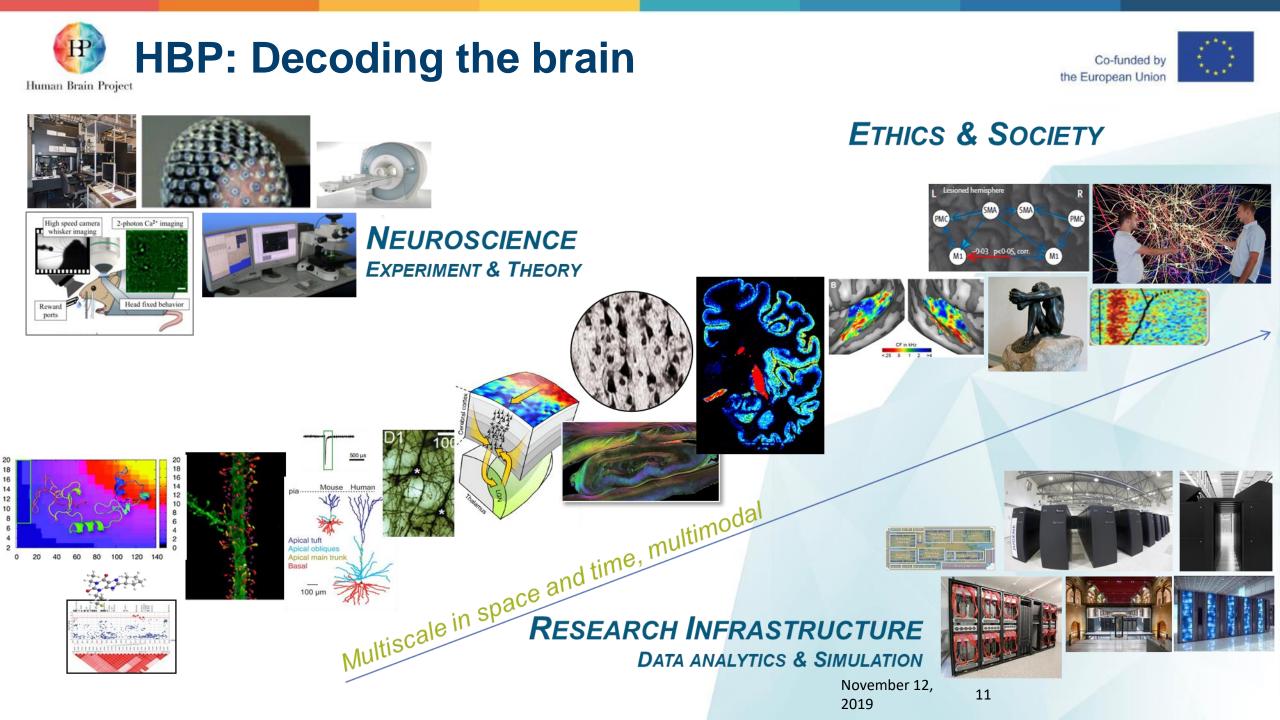
#### www.ebrains.eu Powered by HBP



## Making the infrastructure sustainable



- In the context of preparing for the sustainability of the infrastructure developed by the project, 6 countries participating in HBP have established EBRAINS as a governing Legal Entity / Association internationale sans but lucratif, in Belgium (August 2019)
- Recruitment of EBRAINS CEO just started.







Co-funded by the European Union

#### **Neuroscience data**

- Multiple modalities
- Multiple spatial scales
- Multiple temporal scales







## The Challenge

The Human Brain has astronomical complexity and resilience DATA INTEGRATION CHALLENGE

## The HBP Answer

EBRAINS: a research e-infrastructure for neuroscience - helping the neuroscience community collect, analyse, share, integrate and model data about the brain with the aim of better understanding the functioning of the human brain and its diseases.





- Combining data residing in different sources and providing users with a unified view of them
- Increasingly important as the volume of data explodes
- Focus in many scientific disciplines and in other sectors of society
- Numerous challenges

Databases **Multilevel** and multiscale integration Tools **Models Neuroinformatics** 

Co-funded by the European Unior





- Culture of Knowledge sharing is well developed
- Knowledge builds on data: Culture of Data sharing is lagging behind
- Transformative HBP Neurodata management:
  - Creating a large repository of organized and curated data, unique at the level of containing heterogenous multi-level and multi-modality data
  - Data from HBP and other sources of shared data
  - Accompanied by efficient workflows for organizing, curating, and analysing neuroscience data in the context of brain atlases
- Key aspect: HBP 3-tier curation process for data and models
  - 1. Basic metadata
  - 2. Location metadata registration to reference atlas
  - 3. Neural activity metadata deep integration





## Data sharing

FAIR data service for neuroscience

Brain atlases as key tools in the FAIR data service





## Data sharing

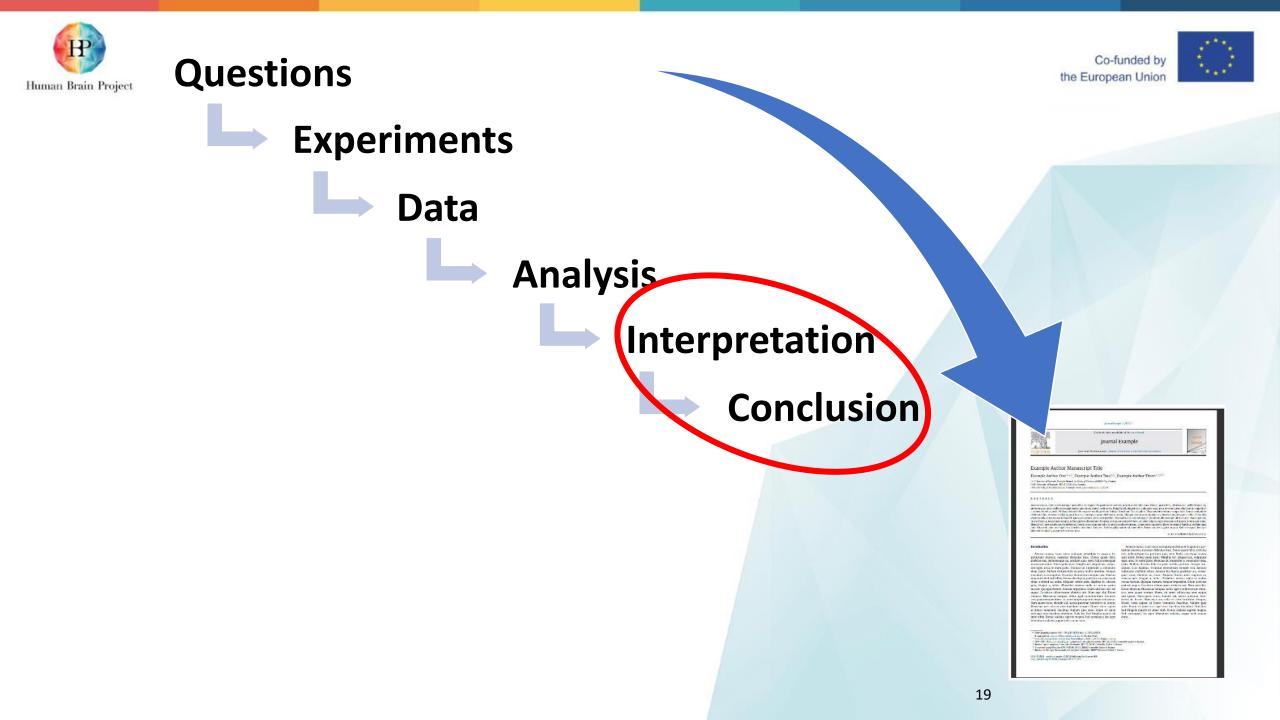
- Why sharing? What are others doing?
- FAIR data service for neuroscience
  - What is it? Is it relevant for you?
- Brain atlases as key tools in the FAIR data service
  - What would geography be without atlases of the planet Earth?
  - What will future neuroscience be without tools for navigating the brain?





## Data sharing

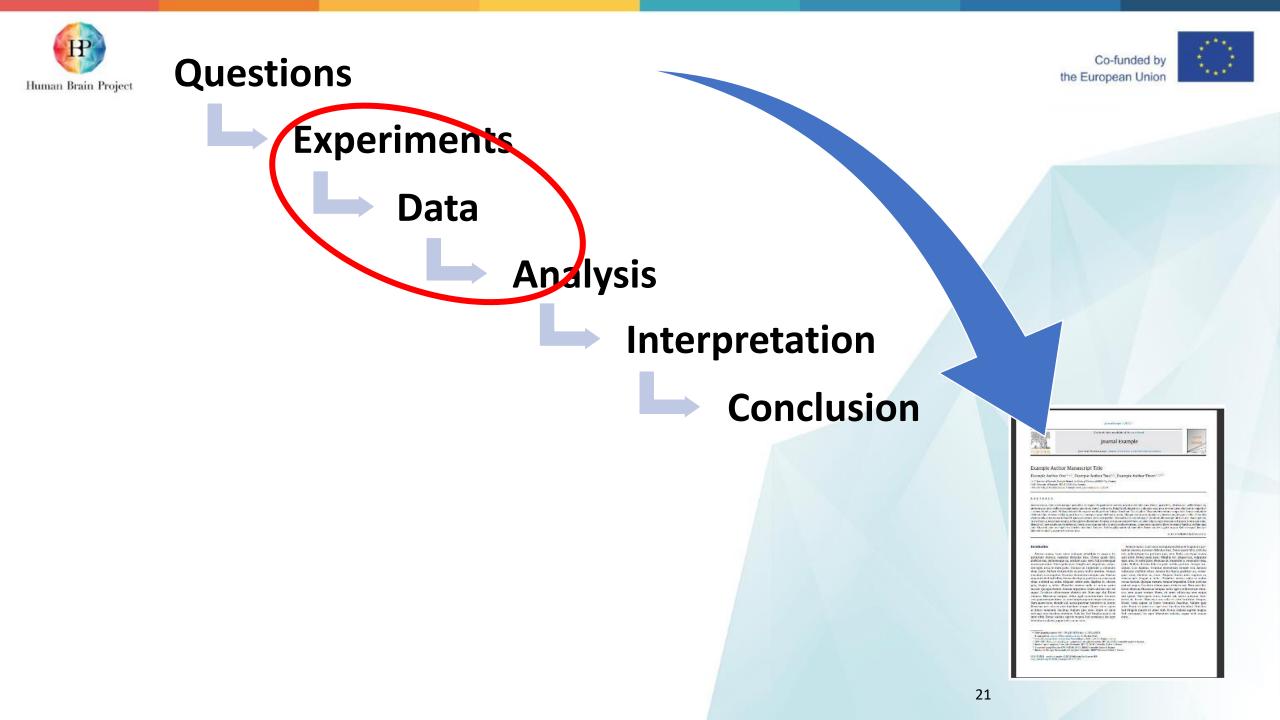
- Why sharing? What are others doing?
- FAIR data service for neuroscience
  What is it? Is it relevant for you?
- Brain atlases as key tools in the FAIR data service
  - What would geography be without atlases of the planet Earth?
  - What will future neuroscience be without tools for navigating the brain?



## **Knowledge** sharing through journals



- The history of scientific journals dates from 1665, when the French <u>Journal des sçavans</u> and the English <u>Philosophical</u> <u>Transactions of the Royal Society</u> first began systematically publishing research results.
- Over a thousand scientific journals were founded in the 18th century, and the number has increased rapidly after that.
- ~ 40.000 journals today
- ~ 60 million articles since 1665
- ~ 2 million new articles per year
- Prior to mid-20th century, peer review was not compulsory. Nature instituted formal peer review in 1967.









- In Journal articles?
- In Supplementary material provided together with journal articles?
- In Data repositories coupled to journals?
- In other Data repositories?

#### What do others do?



Brussels, 14.3.2018 SWD(2018) 83 final

#### COMMISSION STAFF WORKING DOCUMENT

Implementation Roadmap for the European Open Science Cloud

... to create a fit for purpose pan-European federation of research data infrastructures, with a view to moving from the current fragmentation to a situation where data is easy to store, find, share and re-use.





Launched November 2018



EUROPEAN COMMISSION

The EOSC will offer 1.7 million European researchers and 70 million professionals in science, technology, the humanities and social sciences a virtual environment with open and seamless services for storage, management, analysis and re-use of research data, across borders and scientific disciplines by federating existing scientific data infrastructures, currently dispersed across disciplines and the EU Member States.

The European Open Science Cloud (EOSC) initiative has been proposed in 2016 by the European Commission as part of the European Cloud Initiative to build a competitive data and knowledge economy in Europe.



#### EUROPEAN OPEN SCIENCE CLOUD

Launched November 2018





# Data sharing Why sharing? What are others doing?

## FAIR data service for neuroscience

- What is it? Is it relevant for you?
- Brain atlases as key tools in the FAIR data service
  - What would geography be without atlases of the planet Earth?
  - What will future neuroscience be without tools for navigating the brain?





- Computational neuroscientists will benefit from having access to «primary data» or more data from experiments: extract key features relevant for modeling and simulation
- Experimental and clinical neuroscientists will benefit from having access to data from other laboratories: improved analysis, new combinations of data, adding data
- Groups producing and sharing data will benefit from future data sharing impact factors
- By providing well organized and interpretable data, accompanied by well defined conditions for access and use, HBP Neurodata management will build trust and professionalize the sharing of data



nber 12, 27

2019



Co-funded by the European Union

Human Brain Project

ct Science -

ce - Platforms -

Collaborate + Follow HBP +

About - Education & Training -

Welcome to the Human Brain Project

The Human Brain Project aims to put in place a cutting-edge research infrastructure that will allow scientific and industrial researchers to advance our knowledge in the fields of neuroscience, computing, and brain-related medicine

Learn more about the project

November 12,

2019

28

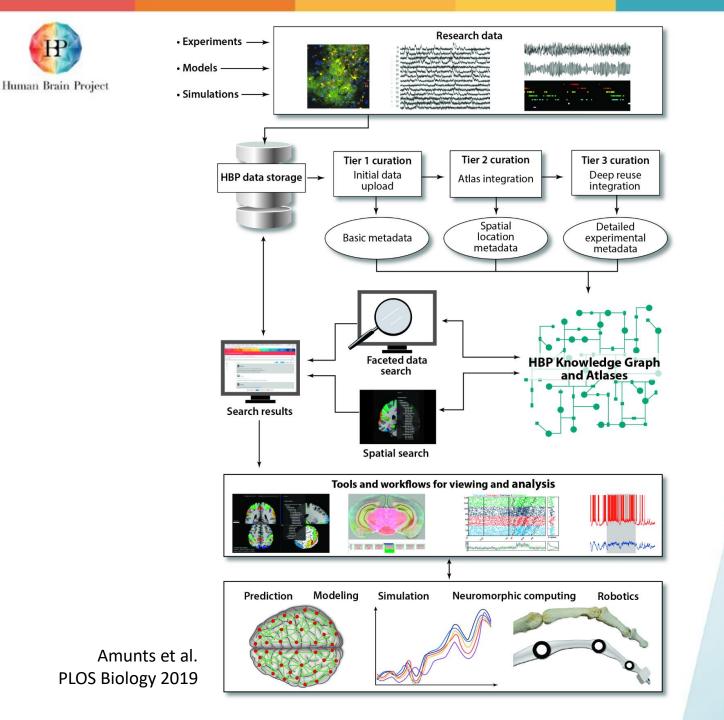
3D-Polarized Light Imaging of the human hippocampus. Image: Axer, Amunts and team, Jülich.

Social,

Ethical,

Reflective

Co \$ EBRAINS Explore the Brain Medicine Brain Simulation EBRAINS is coming soon. Notify me when EBRAINS laund News FRIDAY, 29 MARCH 2019 New brain atlas of transgenic mouse disease models shared via ... Email Events



#### Data to knowledge

Co-funded by the European Union

Research data are uploaded to data storage at the HBP High Performance Computing centers

The data are tagged with metadata through a 3-tier curation process, INCLUDING ETHICS CURATION

The data are made accessible through searches for metadata in the online HBP Knowledge Graph and HBP Atlases

Users can analyze data using tools and workflows for visualization and analysis available through the HBP infrastructure

The multiple-scale data are used for informing modeling and simulation, and for extracting principles of relevance for development of brain-inspired technologies November 12, 2019 29



#### Explore interactive 3-D anatomical brain atlases

Navigate to your favourite brain region and inspect the structure of the brain in any plane of orientation.

The Big Brain  $\Rightarrow$  JuBrain  $\Rightarrow$  WHS rat brain atlas  $\Rightarrow$  Allen Mouse CCF  $\Rightarrow$  More informations on HBP Atlases  $\Rightarrow$ 

SHARE data

HBP offers a comprehensive **management and validation** of all data and metadata before it is uploaded and made available in the Knowledge Graph search. FIND data

Explore neuroscience datasets shared through HBP's Knowledge Graph **data sharing repository**. Browse through a collection of **HBP supported tools** (reference atlases, elephant, ilastik) to visualise, combine and investigate data.

**USE** data

How to share my data

Knowledge Graph Search

How to use our data







Fill out request form:

HBP partners: [Contact HBP data curation team]

Non-HBP scientists [Request data sharing service]





Next deadline: March 31st



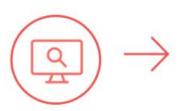
Deliver your data and metadata



1-2 weeks after acceptance



View data in the HBP Knowledge Graph



1-2 months after acceptance

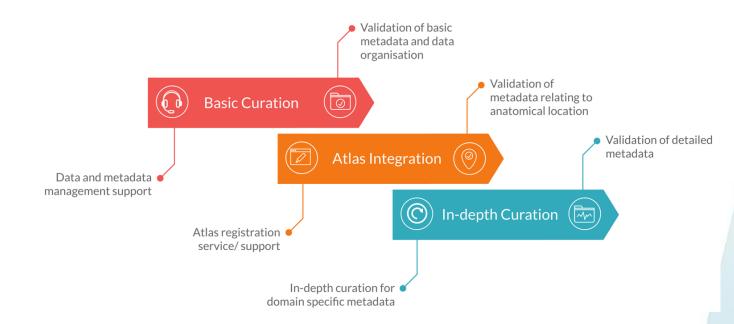






= the organisation and integration of data from various sources into a collection

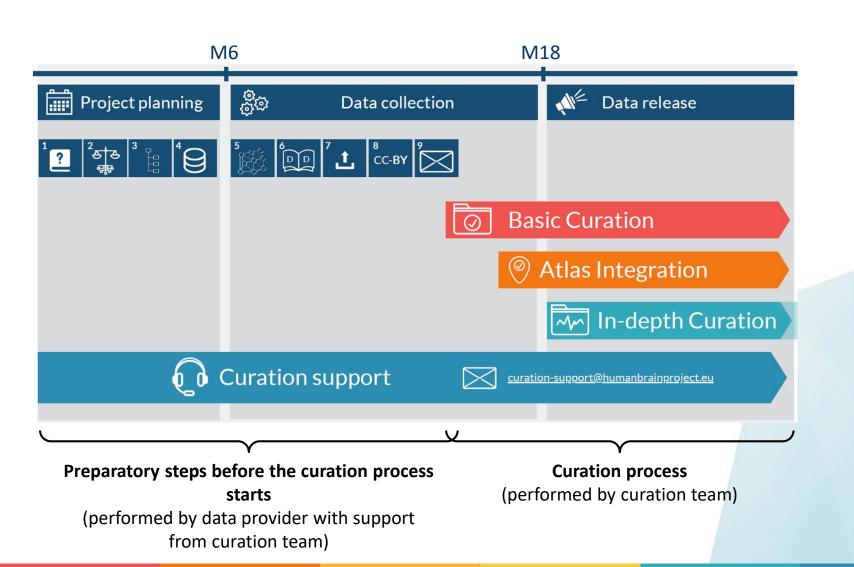
## The HBP's data curation process is organised in three tiers: Basic Curation, Atlas Integration and In-depth Curation.



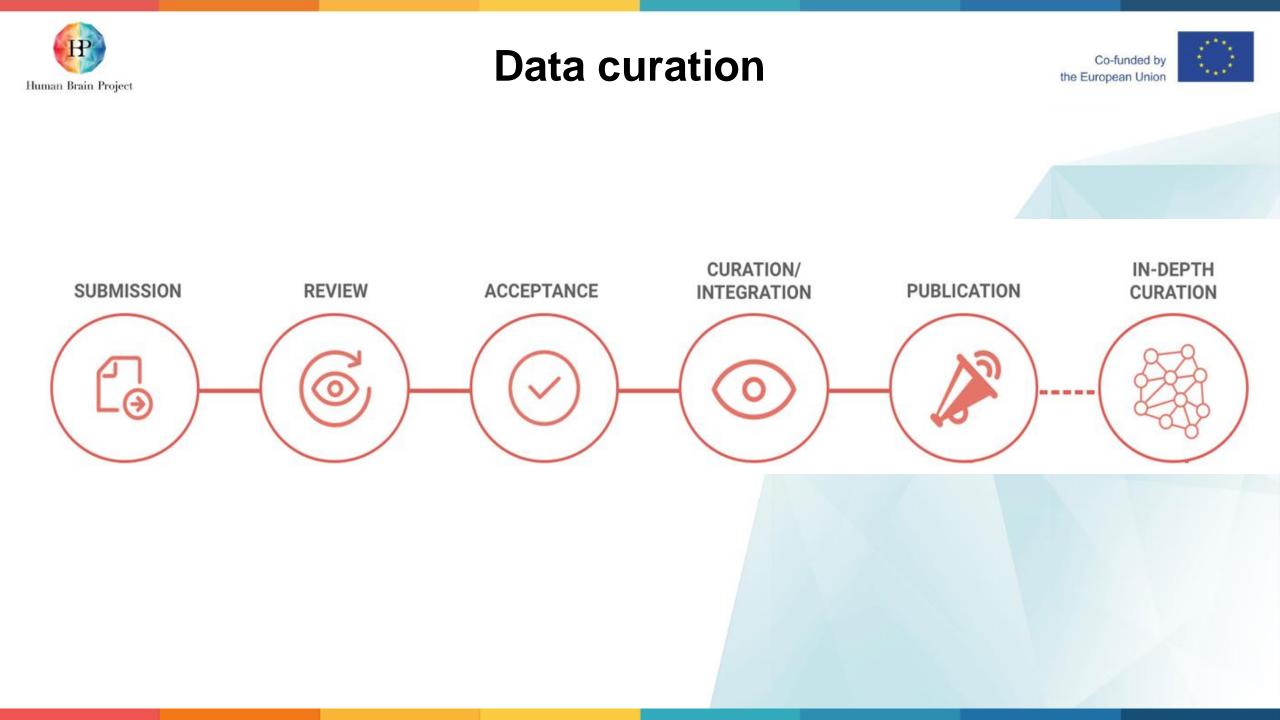
Ensures that... ...data and metadata are organised / managed / stored and integrated in a collection of other neuroscience data ...data are made FAIR



### **Data curation**

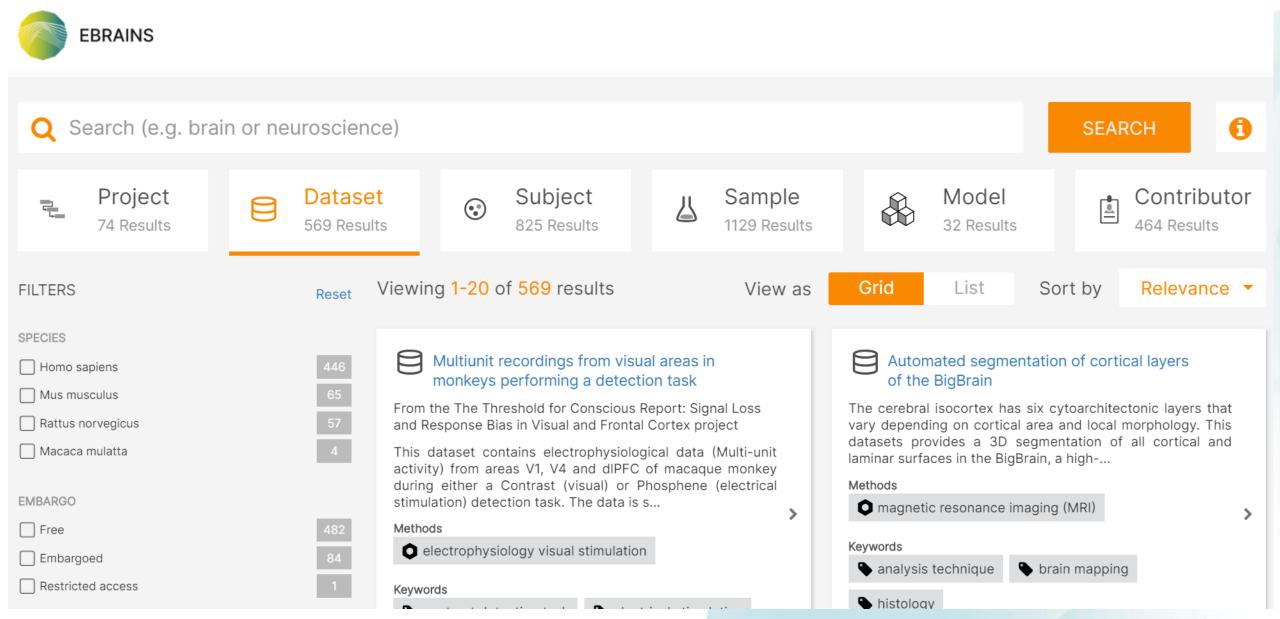


- Download and read the data integration guide
- Complete the **ethics survey**
- ំ Organise your data consistently
- Screate an account for the HBP storage
- Download and fill in the metadata template MINDS
- Download and fill in the DataDescriptor
- **Upload** your data to the HBP storage
- Choose a licence for your dataset(s)
- Contact the curation support to inform us about your **submission**









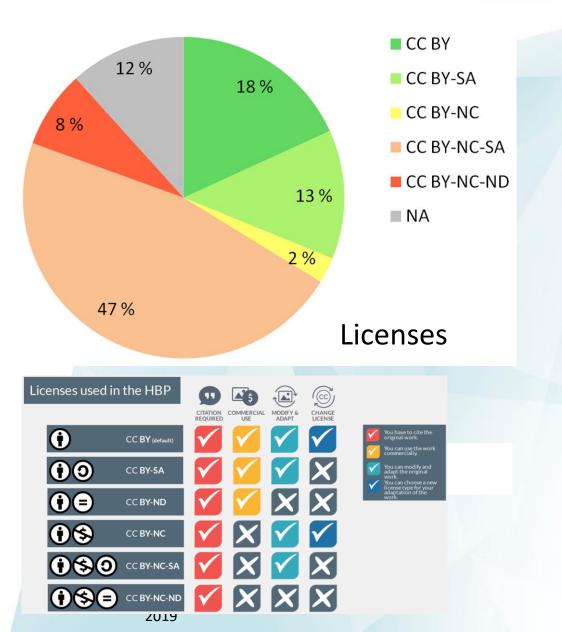
Find data

Human Brain Project

HBP Knowledge Graph Highest FAIRness score HBP Knowledge Graph Lowest FAIRness score







Co-funded by the European Union

1	No.	
100	102	10

Huma

		Do you allow commercial uses of your work?		
		Yes	No	
Do you allow adaptations of your work to be shared?	Yes	Attribution 4.0 International (this is the default)	Attribution- NonCommercial 4.0 International	
	No	Attribution- NoDerivatives 4.0 International	Attribution- NonCommercial- NoDerivatives 4.0 International	
	Yes, as long as others share alike	Attribution-ShareAlike 4.0 International	Selected License Attribution- NonCommercial- ShareAlike 4.0 International	

## © creative commons

- Helps licensors (creators) retain copyright while allowing others to copy, distribute, and make some uses of their work
- Ensures licensors get the credit for their work they deserve
- Works around the world and lasts as long as applicable copyright lasts (because they are built on copyright)





SEARCH

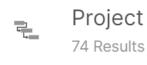
A

Contributor

464 Results









SPECIES

Homo sapiens

Mus musculus

Rattus norvegicus

Macaca mulatta

EMBARGO

Free

Embargoed

Restricted access

YouTube video: Multiscale Integration of Brain Data https://www.youtube.com/watch?v=atCdA2H45iU&t=3s







### HBP Collaboratory

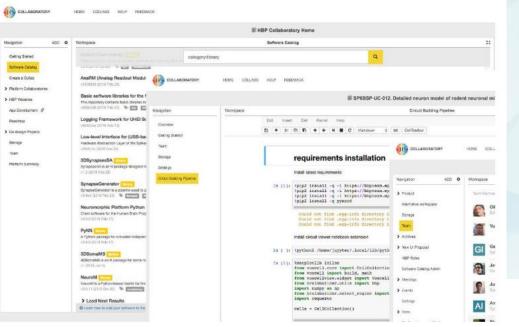
The HBP Collaboratory is divided into collaborative workspaces or *collabs* which helps you share your analyses and collaborate with others on data driven, reproducible scientific workflows.

The collaboratory collects tools from the HBP Platforms in one place.

- Neuroinformatics Platform
- Brain Simulation Platform
- High Performance Analytics and Computing Platform
- Medical Informatics Platform
- Neuromorphic Computing Platform
- Neurorobotics Platform

### The Collaboratory Jupyter notebook

Additionally, numerous standard analysis packages are installed by default into the Collaboratory Jupyter notebook, making your scientific work easier. See the growing list of software packages and services registered here.



#### Visit the HBP Collaboratory 🔶





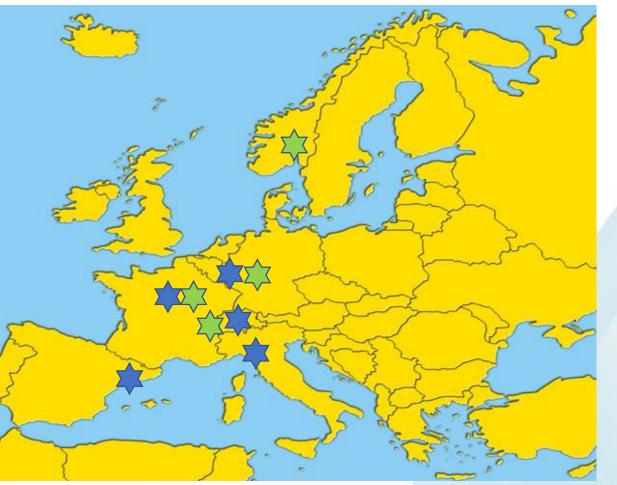












Storage and computing







Centro Svizzero di Calcolo Scientifico Swiss National Supercomputing Centre



BSC Barcelona Supercomputing Center

Centro Nacional de Supercomputación



40





















Storage and computing





Centro Svizzero di Calcolo Scientifico Swiss National Supercomputing Centre







41

Barcelona Supercomputing Center Centro Nacional de Supercomputación





### **Publications with HBP Data Citations**



#### www.nature.com/scientificdata

- 🔁 DATASET 35. Zu, T. et al. Recovery from polyglutamine-induced r 8853-8861 (2004).
- 36. Sgobio, C. et al. Optogenetic measurement of presyn Description: Tabular overview (csv format) of brain-wide reporter expression mapped in five
- indicator expression in dopaminergic neurons. PLoS 37. Echigo, R., Nakao, K., Fukaya, M., Watanabe, M. & E272-E278 (2009).

### **Data Citations**

microscopic images that were spatially registered to the Allen Mouse Common 1. Lillehaug, S. et al. Human Brain Project Neuroinforn Coordinate Framework (v2; mouse.brain-map.org). The density of labeling was OPE assessed by two independent researchers using a semi-quantitative grading 2. Lillehaug, S. et al. Human Brain Project Neuroinforn system from 0 - 4, introduced in Yetman et al., Brain Struct Funct 221:2231-49, 3. Lillehaug, S. et al. Human Brain Project Neuroinforn 2016. Here grade 0 represents absence of labeled cells (less than 1 per 0.01 mm2), grade 1 - low density (few cells, possible to count), grade 2 - medium density 4. Lillehaug, S. et al. Human Brain Project Neuroinforn (several cells that can be individually discerned, but not readily counted), grade 3 high density (many labeled cells with large degree of overlap), and grade 4 - very 5. Lillehaug, S. et al. Human Brain Project Neuroinforn high density (where individual cells cannot be discerned). For each promoter-tTA line one representative case was semi-quantitatively scored and results verified in 6. Lillehaug, S. et al. Human Brain Project Neuroinforn the other cases. Scores did not vary more than 1 grade between cases or researchers in any regions. The highest numbers were reported. If the density of 7. Lillehaug, S. et al. Human Brain Project Neuroinforn labeling was found to vary substantially within a region, the highest observed 8. Lillehaug, S. et al. Human Brain Project Neuroinform score was recorded. 9. Yetman, M., Lillehaug, S., Bjaalie, J., Leergaard, T. & org/10.25493/5H13-1Q0 (2018). Received: 20 February 20 10. Yetman, M., Lillehaug, S., Bjaalie, J., Leergaard, T. & Accepted: 19 December 20 org/10.25493/WB6K-V72 (2018). Published: 26 February 20 11. Yetman, M., Lillehaug, S., Bjaalie, J., Leergaard, T. & License: Creative Commons Attribution 4.0 International org/10.25493/AYBB-BXV (2018). 12. Odeh, F. et al. Human Brain Project Neuroinformati hbp-00170 Comparative DataDescriptor v1p1.txt 9 Terms of use 13. Boy, J. et al. Human Brain Project Neuroinformatics

Comparative overview of brain-wide tetracycline-tran	isactivator expression	
Description :	Project: Transgene expression in tetracycline-transactivator driver lines	
Tabular overview (csv format) of brain-wide reporter expression mapped in five	Custodians: 0 Leergaard, Trygve B.	
commonly used tetracycline-transactivator (tTA) driver lines: neuropsin (Nop); L7/Purkinje cell protein 2 (Pcp2); Pituitary homeobox 3 (Pitx3); cellular prion	Preparation: In silico	
protein (Prnp), and Ca2+/calmodulin-dependent protein kinase IIa (Camk2a). This	Methods	
analysis is derived from microscopic brain images taken from 12 driver-reporter	Anchoring Analysis Density measurment	
constructs (DOIs for source data sets are specified in data descriptor file listed		
below; hbp-00170_Comparative_DataDescriptor_v1p1.txt) in which the different	Keywords	
promoters regulate the expression of the E. coli derived LacZ reporter gene encoding $\beta$ -galactosidase, visualized histologically using X-gal (5-Bromo-4-chloro-3-indolyl $\beta$ -d-galactopyranoside) as a substrate. Labelling was observed in	Atlas B6.Cg-Tg(Camk2a-tTA)1Mmay/DboJ	
	B6.Cg-Tg(Klk8-tTA)QMmay/MullMmmh	
microscopic images that were spatially registered to the Allen Mouse Common		
Coordinate Framework (v2; mouse.brain-map.org). The density of labeling was assessed by two independent researchers using a semi-quantitative grading	Ca2+/calmodulin-dependent protein kinase II promoter	
system from 0 – 4 introduced in Vetman et al. Brain Struct Funct 221:2231-49	view more	

#### DOI for these data: 0

S. Lillehaug, M. Yetman, M. Puchades, M. Checinska, H. Kleven, J. Jankowsky, ... T. Leergaard. (2018). Comparative overview of brain-wide tetracycline-transactivator expression [Data set]. Human Brain Project Neuroinformatics Platform. DOI: 10.25493/ARKS-R7H

Contributors: Lillehaug, S.; Yetman, M.; Puchades, M.; Checinska, M.; Kleven, H.; Jankowsky, J.; Bjaalie, J.; Leergaard, T.

Files (3)

Subjects (1) (1)

Table tTA-distributions.csv 🛛 Terms of use

2019

42



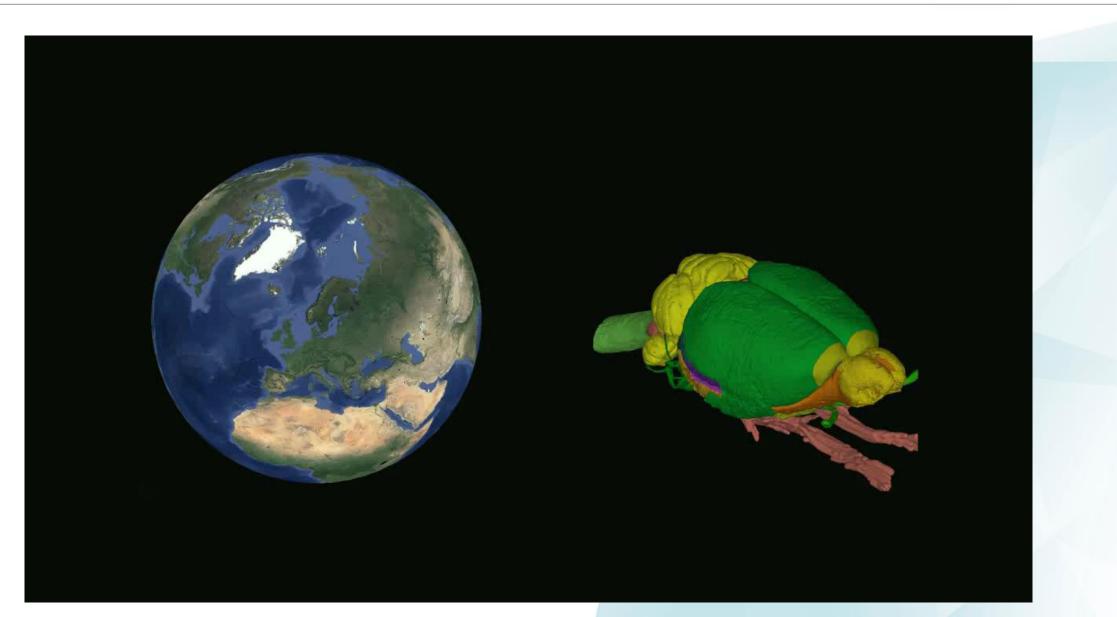


- Data sharing
  Why sharing? What are others doing?
- FAIR data service for neuroscience
  What is it? Is it relevant for you?
- Brain atlases as key tools in the FAIR data service
  - What would geography be without atlases of the planet Earth?
  - What will future neuroscience be without tools for navigating the brain?



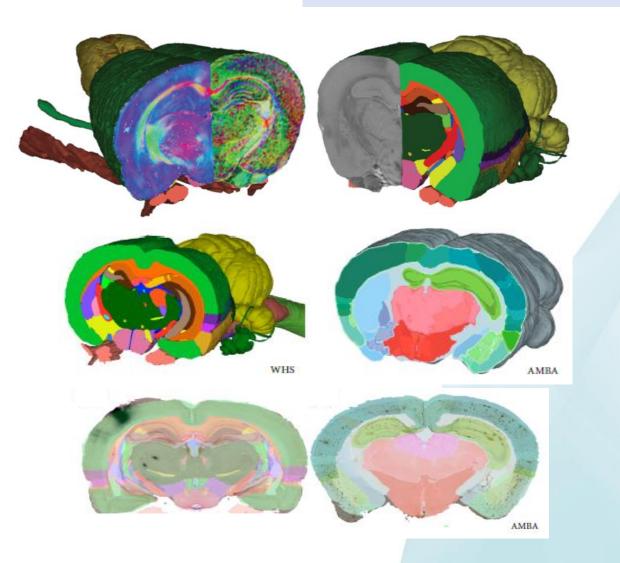
Tier 2 curation: Location in the brain

Co-funded by the European Union



# Human Brain Project Atlas integration

- 3D reference atlases
- Tools for viewing and registration
- Curation service

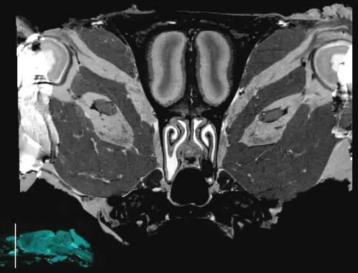






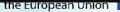
- 3D reference atlases
- Tools for viewing and registration
- Curation service

YouTube video: Next generation rodent brain atlases https://www.youtube.com/watch?v=d-FwWfdDR\_8



Waxholm Space Rat Brain Atlas 3-D atlas template: Microscopic MRI and Diffusion Tensor Imaging data

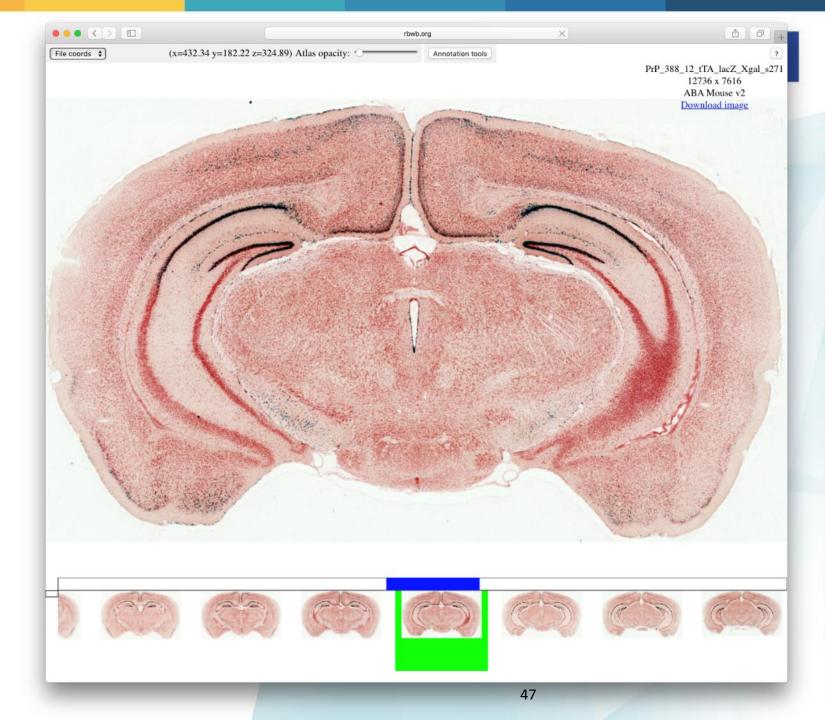








# Interactive viewer for series of 2D images

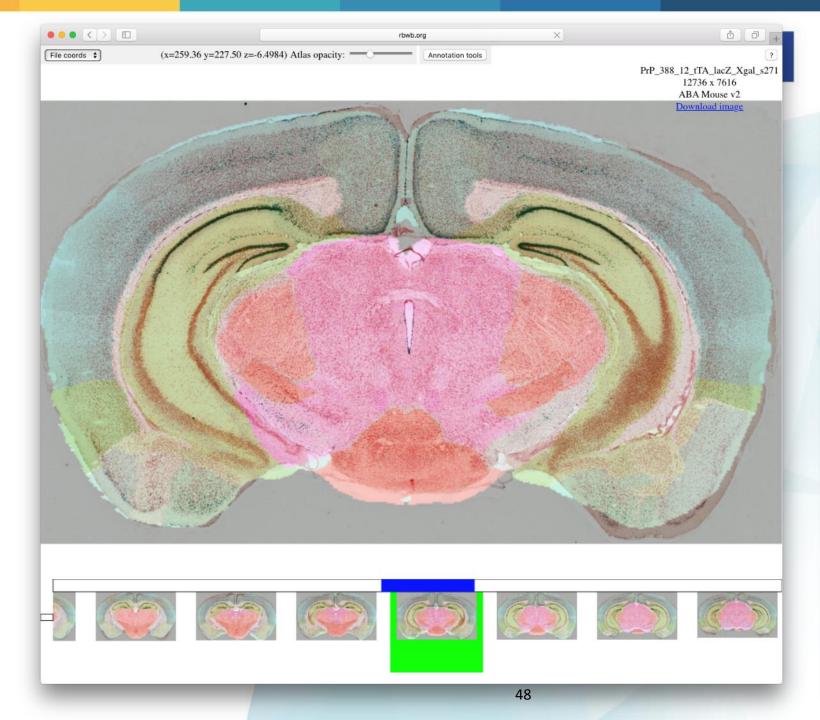




Viewer for series of 2D images

The spatial metadata allow viewing of images with overlay of the reference atlas

User chooses image, adjust level of transparancy of atlas overlay, and navigates the image





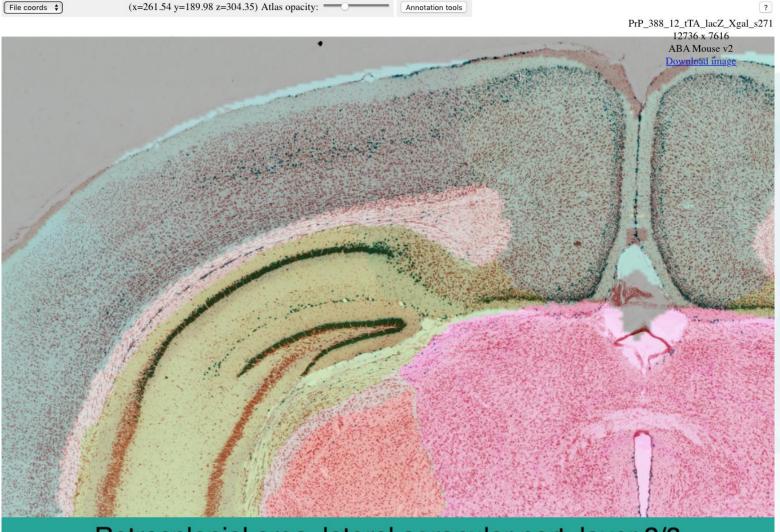
Viewer for series of 2D images

The spatial metadata allow viewing of images with overlay of the reference atlas

User chooses image, adjust level of transparancy of atlas overlay, and navigates the image

Reference atlas coordinates and structure names are available

Spatial information preserved: workflows for feature extraction

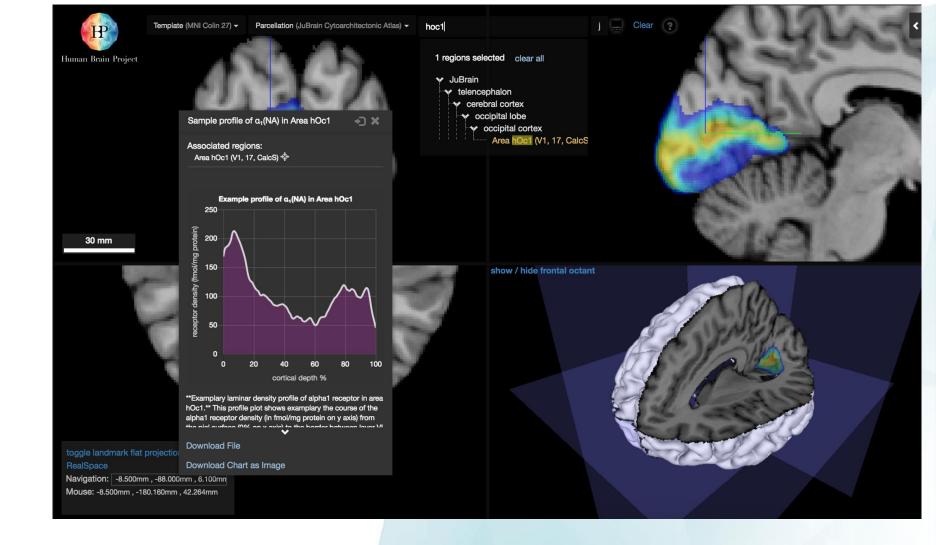


### Retrosplenial area, lateral agranular part, layer 2/3





### Receptor data linked to brain areas

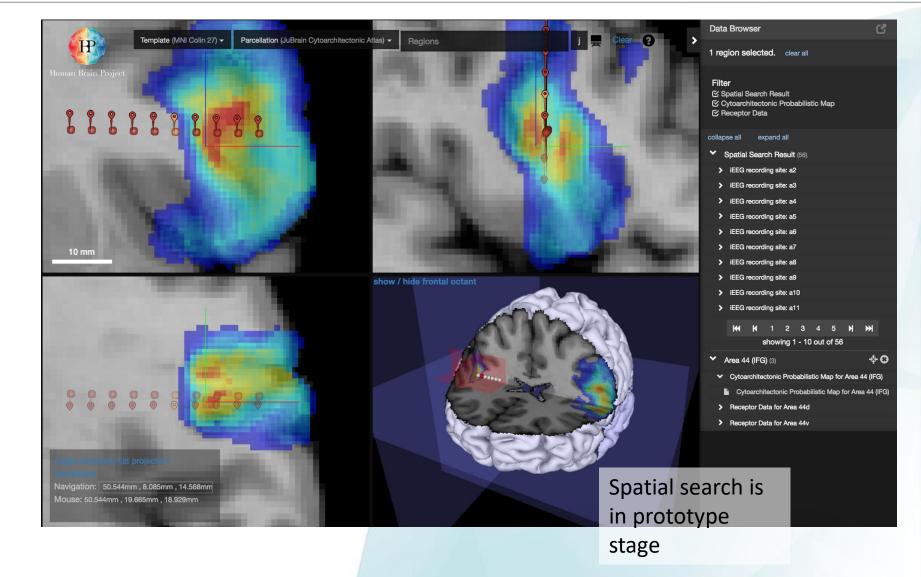








Find iEEG recordings by atlas coordinates









SC1: FAIR data services: Curated and shared data - neuroscience data publishing

SC2: Brain atlas services: navigate the brain in 3D - find, contribute and analyse brain data, based on location

SC3: Brain modelling and simulation workflows: integrated tools to create and investigate models of the brain

SC4: Closed loop Al and robotics workflows: design, test and implement robotic and Al solutions

SC5: Medical brain activity data platform: human intracerebral EEG database and analysis service

SC6: Interactive workflows on HPC or NMC: Europe-wide access to scalable and interactive compute services





# Interactive workflows on HPC or NMC: Europe-wide access to scalable and interactive compute services

- High-performance computing (HPC) has become an important aspect in neuroscience research
  - process and analyse high-resolution data sets
  - simulating large and complex neuronal network models analysing the simulation and/or experimentation results
- Neuromorphic Computing providing access to different kinds of specialized hardware systems, targeting aspects of the emulation of spiking network models difficult to approach by standard simulation methods



53



- **Openness measure** to attract new groups/projects to the HBP IT infrastructure
- Calls to invite external researcher to submit ideas and request HBP engineering solutions
- Target groups: academic & clinical research, pharma and industry

February 2019: 15 Voucher Projects funded





Graphene Flagship Seeking Partners

for Core 3 Project

MAY

- Currently 23 Partnering Projects contributing to the implementation of the HBP roadmap
- Primarily recruited through FLAG-ERA, a new funding mechanism gathering most regional and national funding organisations (NRFOs) in Europe with the goal of supporting the Future G-ERA and Emerging Technologies ABOUT 🗸 FLAG-ERA CALLS FUNDED PROIECTS HUMAN BRAIN PROJECT 🗸 GRAPHENE FLAGSHIP (FET) Flagship concept and FLAGSHIP PILOTS more specifically, the FET Highlights News Flagship initiatives Graphene and Human Brain Project The HBP Research Infrastructure H FLAG-ERA JTC 2019 evaluation 30 Voucher Programme Call 2019 IUL first step: 65 pre-proposals selected (HBP) May 15, 2019 – The first step of the FLAG-ERA JTC 2019
  - May 15, 2019 The first step of the FLAG-ERA JTC 2019 evaluation has been completed. The evaluation was conducted by an independent international scientific evaluation panel for each of the three call topics. Out of 82 submitted pre-proposals, 65 have been invited to submit a full proposal. These represent a total requested funding of ... Continue reading



### **High-Level Support Team**



Extended data curation support

Jupyter notebook user support

Functional data analysis workflow support

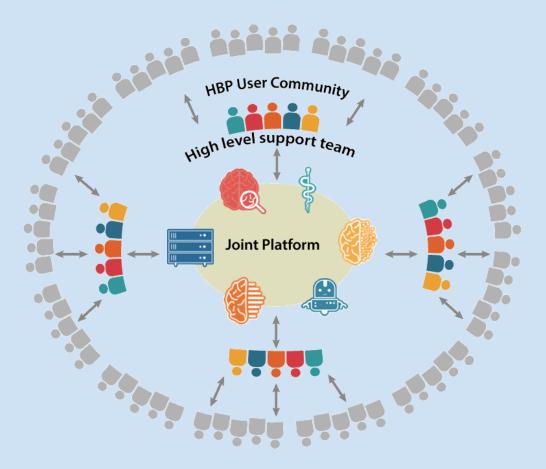
Brain simulation functionality and deep integration support

Medical informatics community management and deep integration support

Neuromorphic computing advanced user support

Neurorobotics documenter and community management

Simulation and data analytics workflow support







- EBRAINS Open Day, Heidelberg, November 25
- HBP Summit, Athens, February 2020
- Questions: support@ebrains.eu



# Thank you

www.humanbrainproject.eu

www.ebrains.eu

