

INSTITUT COCHIN ENTITY DMP - HISTIM Core Facility

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▶ To cite this version:

Maryline Favier, Thomas Guilbert. INSTITUT COCHIN ENTITY DMP - HISTIM Core Facility. Institut Cochin, Inserm U1016, CNRS UMR 8104, Université Paris Cité, F-75014 Paris, France. 2024, https://institutcochin.fr/plateformes/histim. hal-04835929v2

HAL Id: hal-04835929 https://hal.science/hal-04835929v2

Submitted on 18 Dec 2024

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INSTITUT COCHIN ENTITY DMP - HISTIM

Plan de gestion de données créé à l'aide de DMP OPIDoR, basé sur le modèle "Science Europe: structured template for research entities" fourni par DMP OPIDoR.

Plan Details

Plan title INSTITUT COCHIN ENTITY DMP - HISTIM

Deliverable

Version First version

Plan purpose/scope The aim of this DMP is to describe as precisely as possible the various

research products produced by the services offered by the core facility. This DMP is part of a wider framework, that of making available all the

DMPs of the 9 Institut Cochin core facilities (MOUSET'IC,

PROTEOM'IC, PIV, PIME, METABOL'IC, CYBIO, IMAG'IC, GENOM'IC and HISTIM) as well as the DMP of the CID database dedicated to the transfer, storage, consultation and sometimes processing of most of the

data produced by the Institut Cochin's core facilities.

The purpose of this DMP entity is to provide drafting assistance for researchers wishing to draw up a DMP for their research projects and

wishing to call on the expertise of our core facilities.

Fields of science and technology (from OECD classification)

1.6 Biological sciences

Language eng

Creation date 2024-07-18

Last modification

date

2024-12-18

Identifier https://doi.org/10.57889/histim cf

Identifier type DOI

License

Name Creative Commons Attribution 4.0 International

URL http://spdx.org/licenses/CC-BY-4.0.json

Structure Details

Entity's name Institut Cochin Core Facility - HISTIM

Acronym HISTIM

Identifier https://institutcochin.fr/plateformes/histim

Identifier type URL

DescriptionThe HISTIM core facility is open to teams from the Institut Cochin as well as to teams from outside the institute, both institutional and

private. It offers:

• A technical service for animal and human anatomy and pathology. It can handle your samples at any stage of the histological process, depending on your requirements: impregnation and embedding of tissues in paraffin, assistance with the preparation of frozen tissues, microtome or cryostat sectioning, standard staining (HE, HES, etc...) or specific staining (Masson, Pearls, PAS, etc...) virtual slides

- Immunostaining service: conventionnal immunostaining: in brightfield or fluorescence, 1 to 3 markers. Multiplex immunostainingwith: OPAL, ultivue (4 to 8 markers). Hyper-plex immunostaining: MACSima technology (1 to 200 markers in the same tissue section.
- A training service for the equipment needed for histology: microtome, cryostat, laser microdissector, fluorescence microscope and slide scanner.
- Support at every stage of your scientific project: funding application, implementation, follow-up, results

The HISTIM core facility is networked with the PARCC 'Morphology and Histology' core facility (manager: Corinne Lesaffre). The core facilities have become a single one with 2 divisions. This single CF has been accredited by the former Paris Descartes University. This merger means that skills and expertise can be pooled and equipment and services shared, making them accessible at the same rate to all teams from a Université Paris Cité entity.

The HistIM core facility is networked with Rés'Hist which which brings together 14 core facilities in the Paris region.

Since 2010, the Institut Cochin's core facilities, with the help of Yousra Lottin, Quality Engineer, have been committed to a quality approach. In 2012, they obtained ISO 9001: 2008 version certification. This certification is renewed every year after an audit, and in 2018, the platforms were certified to Iso9001: Version 2015. The HISTIM core facility has also been awarded IBISA certification for 2022.

Creation date 2009-09-01

Research outputs:

- 1. Physical Slides (Physical object)
- 2. Photonic Microscopy Virtual Slides (Image)

Contributors

Name	Affiliation	Roles
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INSTITUT COCHIN ENTITY DMP - HISTIM

Data description and collection or re-use of existing data

Physical Slides

Research output description

Name Physical Slides

Description Tissue sections are cut on a microtome for FFPE tissue (4µm thick) and on a

cryostat for frozen tissue (7µm thick).

They are then stained with SPECTRA, an automated staining machine that

handles standard and specific morphological staining.

Type Physical object

Workpackage

Keywords (free-text) Histology

Language eng

Issued Date

Persistent identifier User initials, a number, the first letter of the considered tissue

Identifier type Identifiant local

May contain personal

data?

Nο

May contain sensible

data?

No

May take ethical issues Yes

into account?

Will existing data be reused?

Justification FFPE blocks and frozen tissues can be stored at room temperature or at -80]

respectively so that they can be reused and recut at a later date.

Permanently mounted stained slides can be stored for several years at room

temperature and can be anlayzed again.

How new data will be collected or produced?

Name of the method

Fixation, dehydratation, impregnation and paraffin blocks embedding and cutting

Description

The tissues are prepared and preserved in 2 ways:

- fixation with 4% paraformaldehyde for 24 to 72 hours, followed by dehydration in growing ethanol and paraffin impregnation on a Logos One (MMF) machine, followed by inclusion in FFPE paraffin blocks,
- or frozen in nitrogen-cooled isopentane (with or without prior fixation), in a mould with OCT (cryopreserver) for preservation at -80°.

Tissue embedded in FFPE is microtome-cut into thin $4\mu m$ sections and placed on a glass slide which is stored at room temperature.

Frozen tissues were cut using a cryostat into $7\mu m$ thin sections on a glass slide stored at -20° in the medium term and -80° in the long term.

The slides can be analysed using various techniques:

- standard or specific morphological staining (Leica SPCTRA automated stainer)
- immunolabelling to reveal proteins: conventional (single or double marking in visible DAB and/or red and/or green), fluorescence (single, double or more complex OPAL or Ultivue type)
- multiplex with revelation of dozens of proteins by sequential fluorescent immunostaining (MACSima, Miltenyi)
- spatial transcriptomics (Visium technology, 10x genomics)
- laser microdissection (Arcturus XT, excilone)

The slides are then digitised using the P250 slide scanner and stored as virtual slides in the CID (Cochin Image Database).

Data Nature

Experimental Data

Equipments, technical platforms used

• HIST'IM: https://cat.opidor.fr/index.php/HIST'IM

Photonic Microscopy Virtual Slides

Research output description

Name Photonic Microscopy Virtual Slides

Description HISTIM provides users virtual slides acquisition from samples prepared at the

core facility. These images are used by the biology community for

fundamental and applied research.

The virtual slides are obtained using the Histech P250 3D (MM France),

formerly the Lamina Slide Scanner (Perkin Elmer).

Multidimensional images are available:

• 2D images (x, y),

· histological pyramidal images of tissues,

File format is .mrxs, openable through BioFormat software, enable us to read,

collect and use metadata of all experiments.

Type Image

Workpackage

Keywords (free-text) photonic microscopy, slide scanner

Language eng

Issued Date

Persistent identifier Named after users needs

Identifier type Identifiant local

May contain personal

data?

No

May contain sensible

No

May take ethical issues Yes

into account?

Will existing data be reused?

Justification Images from photonic microscopes can be reused in a number of contexts:

- 1. analysis
- 2. complementary analyses
- 3. reuse by third-party researchers, provided that the image metadata are sufficiently enhanced

How new data will be collected or produced?

Name of the method

Slide canner digitalization

Description

The images are obtained using proprietary acquisition software. The data collected is saved locally, on the acquisition computer, on a data hard disk physically separate from the system hard disk, in a folder named after the team leader (Institut Cochin), or in a folder named after the unit external to

the Institut Cochin, associated with the name of the team leader.

Data Nature

Experimental Data

Equipments, technical platforms used

• HIST'IM: https://cat.opidor.fr/index.php/HIST'IM

Documentation and data quality

Physical Slides

What metadata and documentation (for example way of organising data) will accompagny the data?

Description

The documents and software used to track and trace samples are as follows:

- Open Iris: service request management website
- Service request files
- Core facility charter
- Sample traceability procedures
- Non-conformity procedures
- e-mail communication

Samples follow a formalised path on the platform, making it possible to identify the stage the sample has reached in the histological process.

The nomenclature of the slides created by the platform follows that indicated in the service request file.

Related references

 Guide for collecting and reporting metadata on protocol variables and parameters from slide-based histotechnology assays to enhance reproducibility: https://doi.org/10.1080/01478885.2022.2134022

What methods will be used to ensure their scientific quality?

We have 3 types of characterisation on the platform, using different techniques each time, and each time we have quality controls to ensure the reliability of the results:

- For standard or specific histological staining linked to tissue characterisation at morphological level, we use reference slides which are gold slides, internal control slides and positive control slides. A satisfactory morphological stain is one that is identical to the gold slide, which is the so-called 'perfect' stain in terms of contrast and colour intensity. CIQ slides are used to certify the quality of stains, because
 - reliable staining starts with reliable stains. For specific stains, we use positive controls that are known to be positive for the stain used.
- We use positive and negative controls to characterise proteins and therefore immunolabelling. For some antibodies, we also perform a validation of the antibody prior to the experiment.
- For gene expression characterisation and spatial transcriptomics studies, a reliable result depends first and foremost on a reliable biological sample. We first carry out 2 quality controls: morphology and RNA quality. After handling, a PCR is carried out to check the quality of the RNA extracted. In addition, the reagents used for each type of experiment are monitored for change and renewal, which is done either in the monitoring notebook or directly in the machines.

Photonic Microscopy Virtual Slides

What metadata and documentation (for example way of organising data) will accompagny the data?

Description

For quality, interpretation, reproducibility and sharing value, science data should be accompanied by detailed descriptions of the conditions that were used to produce them. For light microscopy, Micro-Meta App is an intuitive, highly interoperable, open-source software tool that was developed in the context of the 4D Nucleome (4DN) consortium and is designed to facilitate the extraction and collection of relevant microscopy metadata as specified by the recent 4DN-BINA-OME tiered-system of Microscopy Metadata specifications. For the IMAG'IC core facility, when images are imported to the CID database, the machine metadata are automatically recorded as tags for archiving: [Microscope name, Acquisition software, Objectives, Lasers, Filtres, Tube, Acquisition date, User name].

Users can also add sample tags: [Cell type, Tissu type, Staining, Protein, molecule, ...].

It is thus possible to retrieve any data imported into the CID by searching for these machine or sample tags.

Metadata/data standards

• NBO-Q (4DN-BINA-OME-QUAREP Microscopy Metadata Specifications) : https://rdamsc.bath.ac.uk/api2/m111

Metadata language code

eng

Documentation software

Bio-Formats

What methods will be used to ensure their scientific quality?

The data generated on the HISTIM core facility comes from a machine that is regularely checked by the constructor.

The quality metadata are automatically calculated when each data is imported.

Legal and ethical requirements, codes of conduct

Physical Slides

How will other legal issues, such as intellectual property rights and ownership, be managed? What legislation is applicable?

Description

Public: the blocks belong to the platform but are returned at the end of the project.

Private: the blocks are returned at the end of the project.

What ethical issues and codes of conduct are there, and how will they be taken into account?

Description

The charter describes upstream that these ethical codes are respected:

- for animal experimentation projects, the charter requires users to have their APAFIS application in order, with validation by the ethics committees
- for GMOs, users must have their containment declaration in order and the platform undertakes to use them in suitable containment facilities
- for human samples: if patient tissue is used, it must be anonymised before it is sent to the platform. They must be fixed beforehand and frozen tissues are used exclusively in L2.

The rules of our supervisory bodies INSERM, CNRS and UPC are respected.

Photonic Microscopy Virtual Slides

How will other legal issues, such as intellectual property rights and ownership, be managed? What legislation is applicable?

Promulgated in 2016, the Law for a Digital Republic (LRN) includes several measures on which the national open science policy is based. These measures are transposed, in particular, into the Research Code and the Code governing relations between the public and the administration (CRPA).

HISTIM core facility follows these recommendations.

Related references

 Guide d'application de la Loi pour une République numérique pour les données de la recherche : 10.52949/31

What ethical issues and codes of conduct are there, and how will they be taken into account?

Description

The vast majority of samples imaged on our platform are fixed. Imaging is performed under the ethical conditions laid down by law. User are responsible about their APAFIS certification.

Data processing and analysis

Physical Slides

How and with what resources will the data be processed / analyzed?

Description

The slides can be analysed using various techniques:

- standard or specific morphological staining (Leica SPCTRA automated stainer)
- immunolabelling for protein revelation: conventional (single or double marking in visible DAB and/or red and/or green), fluorescence (single, double or more complex OPAL or Ultivue type)
- multiplex with revelation of dozens of proteins by sequential fluorescent immunostaining (MACSima, Miltenyi)
- spatial transcriptomics (Visium technology, 10x genomics)
- laser microdissection (Arcturus XT, excilone)

The slides are then digitised using the P250 slide scanner and stored as virtual slides in the CID (Cochin Image Database).

Equipments, technical platforms

• HIST'IM: https://cat.opidor.fr/index.php/HIST'IM

Photonic Microscopy Virtual Slides

How and with what resources will the data be processed / analyzed?

To ensure the FAIR principle, an image processing workflow is copied to allow the exact same processing to be reproduced despite the constant evolution of software and its updates.

Online processing

Some processing can be done directly by the CID database.

Image processings are available online and are dedicated to denoising, debluring, upsampling and segmentation using CellPose3. Whether for conventional-sized fluorescence images or for large pyramidal coded images, the CID can process these cases with CellPose either directly or in a QuPath environment.

Quantitative analysis

All the methods described before are also available on personal request.

Related references

• QuPath : https://qupath.github.io/

• Cellpose3: one-click image restoration for improved cellular segmentation: 10.1101/2024.02.10.579780

Equipments, technical platforms

• HIST'IM: https://cat.opidor.fr/index.php/HIST'IM

Storage and backup during the research process

Physical Slides

How will data be stored and backed up during the research?

Storage needs No need for storage, users get everything back.

Measures taken for data security

N/A

Photonic Microscopy Virtual Slides

How will data be stored and backed up during the research?

Storage needs

For obvious security reasons, the connection of USB sticks or external hard disks is forbidden on the data acquisition machines.

As soon as the data is imported into the CID, it is archived and can be searched using the different tags accumulated during the import and the different processing and analysis. In all cases the raw data is kept intact, just as the results of processing or analysis will be new data linked to the raw data.

- Data that have been used to publish research must be accessible at any time.
- It is very important to keep the data that is usable at the state of the art of current research. The lack of short-term storage means that we must consider the notion of data quality and therefore retain only those data that have

reliable and rich metadata, and those that allow us to answer a properly posed scientific question.

It is therefore urgent to find a solution for storing data that has not been consulted for more than two years, for example, on a medium that does not clog up the storage of useful data in common use (storage on magnetic tapes, optical disks, etc.).

One of the main purposes of the CID DB is to allow the long-term archiving of this type of data. If the WEB server is attacked, it can be rebuilt at any time by the Image server. All the data are stored on an Image server in the premises of the Institut Cochin. A complete backup is made every day in one of the 4 other buildings of the Institut Cochin. If the Image server is damaged, a restoration is launched via this copy.

To summarize, the IMAG'IC IT architecture runs on 3 servers:

- Data server (backed up every day to a storage bay in another building)
- DMZ: for the imaging website (backed up every day on a storage bay in another building)
- MZ: Database server (no backup) a solution will have to be found here All images are saved to the Cochin Image Database.

cf. INSTITUT COCHIN ENTITY DMP - CID

Storage requirements are indicated for an average year of data generation.

Estimated volume of

data

2

TB

Unit

Equipments, technical platforms

• CID: https://cat.opidor.fr/index.php/CID

Measures taken for data security

CID access by institutional login and personal password.

Data sharing and long-term preservation

Physical Slides

How will data be shared?

Modalities of sharing As physical object, samples are directly given to the users.

YES

Reusability

How will data be long-term preserved? Which data?

Justification The blocks keep well and can therefore be reused. No blocks are physically

stored at the core facility.

Start date End date

Final dispositions

Photonic Microscopy Virtual Slides

How will data be shared?

Modalities of sharing Raw data from research is increasingly in demand by publishers.

Reproducibility in science is of utmost importance and providing raw data is one way to make the results of an experiment more reliable. Institut Cochin's researchers are therefore invited to deposit their raw data from a publication work on the CID (openData of Cochin Image Database). A direct download web page will then be generated and this link can be protected by a unique DOI (Digital Object Identifier). The IMAG'IC and GENOM'IC core facilities are responsible for generating this link. A contract has been signed with the

OPIDoR service of the Inist-CNRS via the DataCite consortium: https://opidor.fr/identifier/

Also, once the images has been used for scientific publication, they are shared through pulbic archive website BioImage Archive. The link is then identify with a DOI link that we are able to create.

Reusability YES, with no condition

Data

repository/catalogs

• BioImage Archive : https://cat.opidor.fr/index.php/BioImage Archive ()

• CID: https://cat.opidor.fr/index.php/CID()

How will data be long-term preserved? Which data?

Justification Every image is available on CID database for an amount of time which has not

been defined yet.

Estimated volume of

data

2

Unit TB

Start date End date

Final dispositions