Laboratory for Interdisciplinary Physics (LIPhy, Grenoble) 140 rue de la Physique , F-38402 St Martin d'Hères



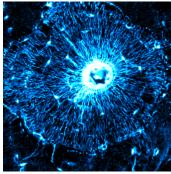
Master thesis offer (4–6 months) Period : January to July 2024 Location : LIPhy, University Grenoble Alpes campus, St martin d'Hères, France Context : Human Frontiers Science Program (HFSP) grant award 2021-2024 Contact : aurelien.gourrier@univ-grenoble-alpes.fr

## Connectomic analysis of multiscale biological cellular networks in mineralized tissues

Your profile: you are currently completing a Master of Science or Engineering degree in one of the following topics : physics of complex systems, bio/medical imaging and signaling or applied mathematics with a solid training in Python programming and image processing ?

Beyond your master, you wish to pursue a PhD to tackle applied biomedical questions in an international context of scientific excellence ?

**Our offer:** we are currently seeking a talented Master student to develop new analytical tools to better understand the role of complex cellular networks in mineralized tissues (bones & teeth). The analysis of complex networks is a fascinating field with incredibly varied applications: social networks interactions, logistic transport, telecom and computing infrastructures etc. It is also a cornerstone of neuroscience,



where biologists seek to understand the relations between measured functions and the brain organization. At the most fundamental level, this implies understanding how the topology/connectivity of the neuron network determines higher scale functions and *vice versa*. Interestingly, few people know, that **there are as many cells in the human skeleton as there are in the brain and that in both cases cells are highly interconnected**, as seen on the fluorescence microscopy image on the left showing interconnected cells surrounding a blood capillary in a human femur. **The precise biological role of this network and its topology is still very poorly understood** and it is therefore tempting to apply similar ideas

developed in the general framework of complex networks analysis to bones. Such approaches are expected to have a strong impact in the biomedical community.

In our group (Structural & Functional Bioimaging – <u>Optics & Imaging team</u>) we developed a unique expertise for optical imaging of cellular networks in mineralized tissues over the recent years and we are now focusing on the image processing and analysis aspects. Together with colleagues from **McMaster University (Canada)** and the **City College of New-York (USA)**, we were recently awarded a prestigious **Human Frontiers Science Program research grant** to tackle this issue.

The key aspects of this project reside in: 1) the huge number of microscopic cells and connections to consider at the organ level, resulting in a large scale dimensionality of the network graph; 2) the specific symmetries and interfaces that tend to split the network into sub-graphs depending on tissue type and 3) the connection to other (e.g. vascular) networks and surfaces, resulting in a true multiscale problem.

Your missions: during this Master internship, you will be in charge of :

- acquiring confocal fluorescence microscopy data of bone samples from the HFSP project
- optimizing image processing pipelines using morphological models and machine learning tools
- implementing graph analysis including geometry and symmetry network characteristics

The "dimensionality curse" is a strong aspect of this part of our project in terms of the amount of images that needs to be handled as input, of the potentially huge network size and the need to provide statistical (reduced) metrics of the whole network for biomedical diagnosis. All of these aspects relate to "**Big Data**" issues that require both **smart approaches from mathematics or physics and programming code efficiency**.

Because of the highly interdisciplinary aspect of the project, you will benefit from the support and training available in our group and in the exceptional scientific environment of Grenoble in Physics, Applied Mathematics and AI.

This Master internship will benefit from a ongoing PhD project which allowed setting up the basic set of tools necessary for the project. The major biomedical application of this project is the **analysis of the cellular porosity network evolution as a function of mineral depletion in bone in a lactating mouse model**.

It is expected that the internship will be followed by a PhD thesis which precise content is opened for discussion around the following suggestions:

- proposing strategies adapted to multiscale studies, e.g. coarse-graining, wavelets...
- providing dynamical analysis of network evolution as a function of biological constraints
- **Your working environment:** this work will be based at the LIPHY, located on the <u>University Grenoble Alpes</u> <u>campus</u> in an exceptional mountain scenery. Our research lab offers unique interdisciplinary expertise at an international level and hosts numerous collaborators from various parts of the world in a sportive and relaxed atmosphere with state of the art technical and scientific support.

The project will be performed in close collaboration with experts in image and complex systems analysis from the LARIS (University of Angers) and GIPSA Lab (Grenoble) and colleagues in Canada and the USA.