

Master de Sciences et Technologies Mention Biologie Intégrative et Physiologie Parcours : Neurosciences Responsable : Professeur Régis Lambert

Internship Proposal Academic Year 2019-2020

1. Host team :

Research Unit (e.g. Department or Institute) : Neuroscience Paris Seine, IBPS, Sorbonne Université, INSERM U1130 – CNRS UMR8246 Research Unit Director : Hervé Chneiweiss Research Team Director : Mangin/Legendre Team name : Dsco Address : 7 quai Saint-Bernard, 75005 Paris Supervisor of the Research Intern for this project : Dongdong LI Telephone : 01 44 27 33 57 E-mail : dongdong.li@inserm.fr

2. Internship project title:

Seeing and Understanding Glial Activities in Brain Neural Network

3. Internship Description :

Human brain comprises ~100 billion neurons, as well as a comparable number of glial cells⁽¹⁾. Their dynamic communications control our cognition, emotion and behavior. While neurons are long viewed to be the information-processing cell, ever increasing evidence reveals unprecedented roles of glial cells in brain activity⁽²⁾. Notably, **astrocytes**, a major type of glial cell, actively regulate the function, therefore the health of neural circuits. The concept of *'tripartite'* synapse has emerged, which includes the neuronal pre- and post-synaptic elements and surrounding astrocyte processes, all interacting in a reciprocal manner⁽³⁾.

Astrocyte activity is encoded by dynamic **calcium** signals, which tightly regulate the microenvironment at neuronal synapses, and has also been implicated in animal behaviors such as body energy balance, sleep, memory consolidation and addiction⁽⁴⁾. Aberrant astrocyte calcium activity stands as a prominent feature of neurological disorders such as Alzheimer's disease. Nowadays, deciphering the cellular and molecular mechanisms that regulate astrocyte calcium activity is an essential challenge for neuro-medical research. It will help to understand, so to normalize glial abnormality in brain diseases.

By a multidisciplinary approach combining mouse genetic/chemical model, optical microscopy and pharmacology, this short-term project aims to enable the **Master-2 student to**

- 1) obtain the state-of-art knowledge on the significance of astrocytes in brain function;
- 2) image directly astrocyte calcium activity in neural networks in living brain tissues;
- 3) probe cellular pathways that regulate astrocyte calcium kinetics in situ.

The supervisor possesses experiences in optical imaging of neuronal and glial activities, optogenetics, and the cellular mechanism of astrocyte activities (<u>http://cvscience.aviesan.fr/cv/1811/dongdong-li</u>), and will work closely with the student on both theoretical and experimental aspects. There is no technical hindrance



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This project is readily to be started without difficulty, while all technical requirements are already established in the host lab. Not only will the up-to-date training on brain neuron-glia interaction be delivered, novel observations on glial activity regulation are also expected. This project is coherent on itself, but can also pave the basis for a potential long-term study (like a PhD study).

References:

- 1. Herculano-Houzel, S. (2014) The glia/neuron ratio: how it varies uniformly across brain structures and species and what that means for brain physiology and evolution. *Glia* **62**, 1377-1391
- 2. Barres, B. A. (2008) The mystery and magic of glia: a perspective on their roles in health and disease. *Neuron* **60**, 430-440
- 3. Dallerac, G., Zapata, J., and Rouach, N. (2018) Versatile control of synaptic circuits by astrocytes: where, when and how? *Nat Rev Neurosci* **19**, 729-743
- 4. Santello, M., Toni, N., and Volterra, A. (2019) Astrocyte function from information processing to cognition and cognitive impairment. *Nat Neurosci* **22**, 154-166