



Felicidades a la nueva graduada de doctorado del ICFO

La Dra. Marina Cunquero Navarro se graduo con una tesis titulada *Use of Advanced Imaging Tools for Assessing the Performance of Novel Therapeutic Techniques for Restoring Vision*

September 16, 2024

Felicidades a la Dra. Marina Cunquero Navarro, quien defendio su tesis esta tarde en el Auditorio del ICFO.

La Dra. Cunquero obtuvo su Master en Laboratorio de Analisis Clinicos por la Universitat Pompeu Fabra y se unio al equipo de investigacion de Microscopia Optica de Superresolucion y Nanoscopia (SLN) dirigido por el Dr. Pablo Loza-Alvarez. Su tesis titulada *Uso de herramientas de imagen avanzadas para evaluar el rendimiento de nuevas tecnicas terapeuticas para restaurar la vision* fue supervisada por el Dr. Loza-Alva

RESUMEN:

Studying vision restoration is paramount for addressing degenerative blinding diseases, which significantly impact quality of life and public health. With more than 230M people

worldwide affected by moderate to severe vision loss, and an estimated increase in blindness from 38M to 115M by 2050, the urgency for effective treatments is clear. Vision, being the most complex and crucial human sense, relies on an intricate network of structures. Light is captured by photoreceptors in the retina and translated into neural signals processed by the brain, enabling sight. Degenerative diseases often involve the progressive deterioration of photoreceptors, leading to blindness. Currently, there is no cure but various approaches are being researched to restore sight. These include gene and cell therapies targeting diseased tissue, as well as methods like optogenetics and neuroprosthetics to modulate neuron activity and bypass dysfunction. By understanding and manipulating neural activity, scientists aim to restore vision or slow down the degenerative processes.

The results of this PhD thesis highlight progress across four areas for vision restoration research. Chapter 2a explores a new retinal implant using biocompatible reduced graphene oxide microelectrodes, demonstrating its ability to record neural signals from retinal tissue and capture light-induced firing patterns in retinal ganglion cells. It also details a protocol for retinal calcium imaging, facilitating future studies combining electrical stimulation with optical techniques. Chapter 2b focuses on adapting an ophthalmoscope for *in vivo*, single-cell resolution imaging of the retina in rodents, aiming to set the basis for future implementations to monitor the functionality of implanted retinal prostheses. Chapter 3 delves into the potential of human-induced pluripotent stem cell-derived retinal organoids for vision research. Using advanced imaging techniques, researchers successfully observed the formation of distinct retinal cell types within the organoids and identified differences in calcium dynamics between healthy and diseased models. Future work aims to refine MEA recordings and investigate the link between retinal organoid structure and function. Chapter 4 introduces a novel three-photon excitation technique offering deeper brain penetration and higher precision compared to traditional methods. This approach successfully regulates neuronal activity in zebrafish with minimal light exposure, showcasing its potential for revolutionizing the study of neural circuits and development of neuromodulation therapies. Taken together, these advancements across retinal implant design, *in vivo* monitoring of the retina, studying the potential of patient-derived organoid models, and development of non-invasive three-photon brain stimulation techniques, pave the way for future development of more effective vision restoration therapies.

Comite de Tesis:

Prof. Dr. Serge Picaud, Institut de la Vision Paris

Prof. Dr. Michael Krieg, ICFO

Dr. Niccolo Accanto, Institut de la Vision Paris