Introductory Remarks to Symposium 27

Brain Organoids for Modelling Immune-Neural Interactions in Epilepsy

Andreas G. Chiocchetti, Frankfurt/Main

Epilepsy is a central nervous system (CNS) disorder hallmarked by seizures and abnormal brain activity. Current anti-seizure drugs block seizures in only ~70% of patients, do not address the underlying pathology and do not impact the progression of the disorder. Hippocampal sclerosis (HS) as well as malformations of cortical development (MCD) such as focal cortical dysplasia (FCD) and lissencephaly as well as low-grade epilepsy-associated tumors (LEAT) such as ganglioglioma are among the most frequent causes for pharmacoresistant focal epilepsy. All these types of brain lesions harbor innate and adaptive immune cell infiltrations which likely contribute to and modulate their epileptogenicity

Understanding the specific mechanisms involved in the interaction of immune cells and cells of the brain parenchyma for the generation and progression of seizures and epilepsy in these disorders will allow the development of novel drugs that modify the process of epileptic neural network transformation itself.

Human induced pluripotent stem cells (hiPSCs) reprogrammed from patient somatic cells have proven as powerful tool to model human diseases including epilepsies. Platforms in which neurons, astrocytes, oligodendrocytes and microglia derived from healthy or diseased subjects mature in a single system represent a robust method to model human brain disorders.

In this symposium we will discuss the potentials and pitfalls of using hiPSC-derived glioneuronal cell cultures (2D) and brain organoids (3D) to address role the development and mechanisms of epileptic neural network aberration and the role of immune-neural interactions therein.

Symposium 27

Friday, March 28, 2025 14:30 - 16:30, Lecture Hall 104

Chairs: Andreas G. Chiocchetti, Frankfurt/Main

- 14:30 **Opening Remarks**Andreas G. Chiocchetti and Nico Melzer
- 14:35 Nico Melzer, Duesseldorf STUDYING IMMUNE-NEURAL INTERACTIONS IN A MODEL OF T-CELL DRIVEN HIPPOCAMPAL SCLEROSIS: PITFALLS AND TRANSLATIONAL VALUE (S27-1)
- 15:00 Denise Haslinger, Frankfurt
 IN VITRO MODELLING OF MATERNAL IMMUNE
 ACTIVATION (MIA) IN CEREBRAL ORGANOIDS
 (\$27-2)
- 15:25 Andrea Rossi, Duesseldorf
 GENOME-EDITING TO MODEL SELECTIVE
 SOMATIC MUTATIONS ASSOCIATED WITH
 FOCAL EPILEPTOGENIC LESIONS (S27-3)
- 15:50 Julia Ladewig, Mannheim
 OMICS INSIGHTS INTO LIS1-PATIENT-DERIVED
 CEREBRAL ORGANOIDS UNRAVEL NOVEL
 MOLECULAR PATHWAYS UNDERLYING DISEASE
 SEVERITY AND SUGGEST THERAPEUTIC STRATEGIES (S27-4)
- 16:15 Discussion / Concluding Remarks