

## **2nd Channelopathy Meeting Tübingen**

## Genetic epilepsies and other neuronal ion channel disorders: Mechanisms and therapeutic perspectives

4<sup>th</sup>-6<sup>th</sup> October 2023, Tübingen, Germany

## **Preliminary Program**

| Wednesday, 4 October 2023 |   |  |
|---------------------------|---|--|
| 7:30 pm                   | Get-Together Schloss Hohentübingen  |  |
| Thursday, 5 October 2023  |   |  |
| 8:30 am                   | Introduction: Holger Lerche (Tübingen)  |  |
| 8:40-11:40 am             | Session 1: Sodium channelopathies   |  |
|                           | Chair: Thomas Wuttke & Moran Rubinstein   |  |
| 8:40                      | Andreas Brunklaus (Glasgow): Emerging phenotypes in sodium channelopathies                                    |  |
|                           | - the role of functional prediction and therapeutic advances  |  |
| 9:00                      | Evgeniia Rusina (Nice): A homeostatic response boosted as therapeutic approach                                |  |
|                           | in <i>Scn1a</i> <sup>+/-</sup> Dravet syndrome mice   |  |
| 9:20                      | Nikolas Layer (Tübingen): Early electrophysiological and transcriptomic                                       |  |
|                           | alterations drive epileptogenesis in Dravet syndrome  |  |
| 9:35                      | Elena Gardella (Dianalund): SCN8A: Clinical complexity and natural history study                              |  |
| 9:55                      | Heinz Beck (Bonn): Targeting aberrant dendritic integration to treat cognitive                                |  |
|                           | comorbidities of epilepsy   |  |
| 10:15, virtual            | Sanjay Sisodiya (London): Genetic epilepsies: complexities to come - SCN1A as an                              |  |
|                           | example   |  |
| 10:35-11:05 am            | Coffee Break  |  |
| 11:05                     | Angelika Lampert (Aachen): Precision Therapy in sodium channel related  |  |
|                           | neuropathic pain  |  |
| 11:25                     | JP Johnson (Xenon, Burnaby): A Selective Na <sub>V</sub> 1.1 Potentiator Enhances                             |  |
|                           | Interneuron Excitability to Normalize Motor Performance in a Dravet Syndrome                                  |  |
|                           | Mouse Model   |  |
| 11:45 am - 1:00 pm        | Session 2: Potassium channelopathies  |  |
| 44.45                     | Chair: Gabriele Lignani & Stephan Lauxmann  |  |
| 11:45                     | Jennifer Kearney (Chicago): Modeling KCNB1-associated developmental and                                       |  |
| 12.05                     | epileptic encephalopathy in mice  |  |
| 12:05                     | Peter Müller (Tübingen): Transcriptomic insights into epileptogenesis and                                     |  |
| 12.20                     | compensatory changes in a KCNA2 loss-of-function mouse model  |  |
| 12:20                     | Konstantin Khodosevich (Copenhagen): Understanding mechanisms of  |  |
| 12:40, virtual            | epileptogenesis using single cell omics  Ann Poduri (Boston): Modeling potassium channelopathies in zebrafish |  |
| 1:00-2:00 pm              | Lunch Break   |  |
| 2:00-3:50 pm              | Session 3: HCN, calcium and ligand-gated channelopathies  |  |
| 2.00-3.50 pm              | Chair: Dirk Isbrandt & Ulrike Hedrich-Klimosch  |  |
| 2:00                      | Bina Santoro (New York): Molecule, cell, organism: multi-level analysis of a sick                             |  |
|                           | ion channel   |  |
| 2:20                      | Christopher Reid (Parkville): A small-molecule precision medicine for HCN1 DEE                                |  |
| 2:40                      | Stephan Marguet (Cologne): HCN/h channel modulation of excitation/inhibition                                  |  |
|                           | balance in the MEC is crucial for cortico-hippocampal information flow and                                    |  |
|                           | intrahippocampal dynamics   |  |
| 3:00                      | Jörg Striessnig (Innsbruck): CACNA1D L-type calcium channelopathies in  |  |
|                           | neurodevelopmental disorders: opportunities for drug repurposing  |  |
|                           |   |  |

| 3:20         | Philip Ahring (Sydney)/Rikke S. Møller (Dianalund): Pathogenic <i>GABRA3</i> variants lead to dominant or recessive X-linked disorders depending on functional |
|--------------|--|
|              | outcomes   |
| 3:50-5:00 pm | Poster session (with coffee)   |
| 5:00-7:00    | Session 4: Molecular therapeutic board (case reports: 5-10 min including   |
|              | discussion)  |
|              | Chair: Holger Lerche & Michael Alber (Tübingen)  |
|              | Stephan Lauxmann/Michael Alber   |
|              | Robert Lauerer-Braun   |
|              | Rikke Steensbjerre Møller  |
|              | Further cases from the audience  |
|              |  |
| 8:00 pm      | Dinner Restaurant "Liquid"   |

| Friday, 6 October 2023 |   |  |
|------------------------|---|--|
| 8:30-9:50 am           | Session 5: Gene therapy and human models  |  |
|                        | Chair: Maria-Patapia Zafeiriou & Niklas Schwarz   |  |
| 8:30                   | Gabriele Lignani (London): On demand gene therapy for epilepsy  |  |
| 8:50                   | Moran Rubinstein (Tel Aviv): Viral-mediated expression of Nav1.1 ameliorates Dravet syndrome in mice post seizure onset             |  |
| 9:10                   | Nael Kasri (Nijmegen): Leveraging spontaneous activity in human stem cell derived neurons to model <i>SCN1A</i> -related epilepsies |  |
| 9:30                   | Maria-Patapia Zafeiriou/Hendrik Rosewich (Göttingen): Modeling <i>ATP1A3</i> related disorders in bioengineered neuronal organoids  |  |
| 10:00                  | Sarah Weckhuysen (Antwerpen): KCNQ2-related epilepsies in iPSC-derived neuronal cultures  |  |
| 10:20                  | Filip Rosa (Tübingen): KCNQ2-DEE stem cell model shows a developmental delay along a prologed culture time up to 6 months           |  |
| 10:35-11.00 am         | Coffee Break  |  |
| 11:00-12:30 am         | Session 6: Epigenetics, protein structural modeling & prediction tools  |  |
|                        | Chair: Rikke Steensbjerre Møller & Christopher Reid   |  |
| 11:00                  | Katja Kobow (Erlangen): Epigenetics for diagnosis and mechanistic understanding of drug-resistant focal structural epilepsies       |  |
| 11:20                  | Christian Bosselmann (Cleveland): Predicting the functional effects of ion channel  |  |
|                        | variants: State of the art and future directions  |  |
| 11:40                  | Tobias Brünger (Cologne): Predicting molecular function and pathogenicity in ion channel disorders                                  |  |
| 12:00                  | Giulia Rossetti/Jan-Philipp Machtens (Jülich): Studying functional gating   |  |
|                        | mechanisms in wild-type or mutant Na <sub>V</sub> channels/In-silico variant-specific drug  |  |
|                        | screening   |  |
| 12:30-1:30 pm          | Farewell Lunch with free discussions  |  |









GEFÖRDERT VOM Bundesministerium für Bildung und Forschung