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Seminar Series:

## ***Frontiers in Pharmacology***

**Professor Bernd Fakler, M.D.**

Department of Physiology  
University of Freiburg

*“Coupling of Cav and BKCa channels analyzed by functional proteomics”*

*BKCa channels are dually activated by membrane depolarization and elevation of cytosolic Ca<sup>2+</sup>. Under normal cellular conditions BKCa channel activation requires Ca<sup>2+</sup> concentrations that typically occur in close proximity to Ca<sup>2+</sup>-sources. It is shown that BKCa channels affinity-purified from rat brain are assembled into macromolecular complexes with the voltage-gated calcium channels Cav1.2 (L-type), Cav2.1 (P/Q-type) and Cav2.2 (N-type). Heterologously expressed BKCa-Cav complexes reconstitute a functional 'Ca<sup>2+</sup> nano-domain' where Ca<sup>2+</sup> influx through the Cav channel activates BKCa in the physiological voltage-range with sub-millisecond kinetics. Complex formation with distinct Cav channels enables BKCa-mediated membrane hyperpolarization that controls neuronal firing pattern and release of hormones and transmitters in the CNS..*

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