The 57th Frontier Brain Science Seminar Sponsored by Research Center for Idling Brain Science (RCIBS)

ADAR2-mediated RNA editing of Cav1.3 channels: Implications in learning and memory



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Date: 14th April, 2023 (FRI.) 17:00~18:30

Venue: Clinical Lecture Room1(附属病院2階 臨床講義室1)

Our lab has a long-standing interest in understanding the diversification of structure and function of voltage-gated calcium channels via post-transcriptional modifications. Here, I will present our discovery of Cav1.3 RNA editing and how editing of the IQ-domain affects Ca2+-dependent inactivation of the Cav1.3 channels. Unexpectedly, we uncovered that RNA editing of the Cav1.3 is neuron-selective and that restricted editing of Cav1.3 channel is due to the alternative splicing of an editing repressor which is a ubiquitously expressed splicing factor, SRSF9. To address the physiological significance of Cav1.3 RNA editing, we generated editing complementary sequence null (Cav1.3 ECS) mice that were genetically targeted to produce unedited Cav1.3 channels. Electrophysiological hippocampal slice recordings showed enhanced late-LTP and the Cav1.3 DECS mice have better spatial learning and memory as demonstrated by behavioural tests using the Morris Water-maze. The next question we will address is whether there are trade-offs to the loss of Cav1.3 RNA editing.

References

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Loss of Ca V 1.3 RNA editing enhances mouse hippocampal plasticity, learning, and memory. Proc Natl Acad Sci USA. 2022 Aug 9;119(32):e2203883119. doi: 10.1073/pnas.2203883119.

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RNA editing of ion channels and receptors in physiology and neurological disorders. Oxford Open Neuroscience 2022, 1, 1–16 doi.org/10.1093/oons/kvac010 Advance access publication date 11 July 2022

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