

Unique fingerprint: What makes nerve cells unmistakable?

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Protein variants shape wiring of nerve cells in the brain. Credit: Biozentrum, University of Basel

Protein variations that result from the process of alternative splicing control the identity and function of nerve cells in the brain. This allows organisms to build a highly complex neuronal network with only a limited number of genes. The study describing a detailed map of neuronal splicing conducted by a research team at the Biozentrum, University of Basel, has now been published in *Nature Neuroscience*.



Our <u>brain</u> consists of hundreds, if not thousands, of different types of nerve cells that control our brain functions due to their individual characteristics. But how do the different cell types manage to develop their diverse traits? In a genome-wide analysis, the team led by Prof. Peter Scheiffele at the Biozentrum, University of Basel, has now discovered that <u>alternative splicing</u> leads to a broad range of variants of individual proteins, which ultimately allows to distinguish types of nerve cells.

Alternative splicing determines cell types

Alternative splicing can generate multiple different protein variants from a single gene. In the <u>mouse model</u>, Scheiffele's team investigated splice variants in a panel of neuronal cell types. "We have been able to identify hundreds of splice variants that enable us to differentiate between different types of neurons," says Scheiffele. "There are unique repertoires of variants in each nerve cell type."

These repertoires of splice variants significantly shape the identity and function of <u>nerve cells</u>. "Although all neuronal cell types contain the same set of genes, even closely-related cell types produce different splice variants," explains Scheiffele. In particular, proteins located at the neuronal contact points—the synapses, which mediate the transmission and processing of information—are extremely diverse. Thus, the splicing process also controls the function of the neuronal circuits in the brain.

Data platform for scientists

The generation and analysis of the extensive data sets is part of the EU project "SPLICECODE." In collaboration with the "Center for Scientific Computing" (sciCORE), a user-friendly website has been set up which allows scientists worldwide to investigate the role of individual



splice variants in brain function.

More information: Elisabetta Furlanis et al, Landscape of ribosomeengaged transcript isoforms reveals extensive neuronal-cell-classspecific alternative splicing programs, *Nature Neuroscience* (2019). DOI: 10.1038/s41593-019-0465-5

Provided by University of Basel

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