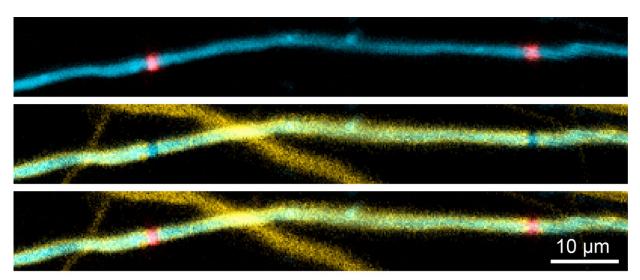


Evidence for myelin sheath remodeling revealed by in vivo imaging

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Myelin, Axon, Node of Ranvier

Using newly developed markers, the scientists were able to visualize axons, myelin, and nodes of Ranvier in vivo. A version of this image is used in the current article in *Current Biology*, DOI: 10.1016/j.cub.2018.01.017 (CC BY-NC-ND 4.0)) Credit: Czopka / TUM

Nerve fibers are surrounded by a myelin sheath. Scientists at the Technical University of Munich (TUM) have now made the first-ever "live" observations of how this protective layer is formed. The team discovered that the characteristic patterns of the myelin layer are determined at an early stage. However, these patterns can be adjusted as



needed in a process apparently controlled by the nerve cells themselves.

The myelin sheath surrounding the axons are comparable to the insulation covering an electric wire. Without it, the rapid propagation of electric signals would not be possible. Damage to this insulating layer, for example from illnesses such as <u>multiple sclerosis</u>, may result in serious impairments.

But myelin does not form a continuous coating along the axon. Instead, it is divided into segments. These can vary in length and are separated by gaps known as nodes of Ranvier. For the complex network of the central nervous system to function properly, the speed of the connections is not the only consideration. The crucial factor is the fine tuning: the signals have to reach the right place at exactly the right time. The transmission speed of information through an axon is partly determined by the number and length of the segments.

The bodies of humans and animals have some ability to repair damaged myelin sheaths. Dr. Tim Czopka, a neuroscientist at TUM, has succeeded in making the first-ever in vivo observations of this process. Czopka and his team used newly developed markers to visualize the formation myelin segments surrounding axons in the spinal cords of zebrafish. They conclude that characteristic patterns made up of myelin segments with different lengths along an axon are defined within a few days after myelin formation begins. Although the segments continue growing from that time onward as the body of the zebrafish grows, the pattern is preserved.

Following this observation, the researchers destroyed selected segments. "What happened next surprised us," says Tim Czopka. "After the destruction of the segments, the <u>myelin sheaths</u> began to dynamically remodel. In the end, the damage was repaired and in most cases the original pattern was restored." The regeneration followed a fixed



sequence. First, the adjacent segments expanded to close the gap. A new segment then formed between them, and they contracted to their original size.

This raises an important question: Who controls the emergence and the restoration of the segment pattern? "Our observations suggest that it is not the oligodendrocytes—the cells that form myelin—that decide where it is formed, but rather the axons," says Tim Czopka. "You could say that they know best which <u>pattern</u> is needed for the signals to be transmitted at optimal speed."

The team is now studying how the segment patterns are affected by the targeted stimulation of nerve cell activity and by the neurotransmitters released as a result. "If we can understand the role of the axons in <u>myelin</u> generation and remodelling, it may yield new approaches to controlling it," explains Czopka. "That would be relevant for the treatment of illnesses such as multiple sclerosis."

More information: F. Auer, S. Vagionitis, T. Czopka, "Evidence for Myelin Sheath Remodeling in the CNS Revealed by In Vivo Imaging", *Current Biology* (2018). <u>DOI: 10.1016/j.cub.2018.01.01</u>

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