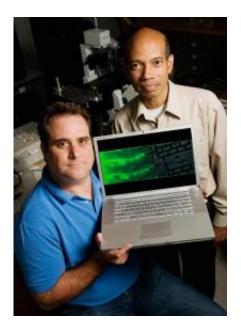


Tension in axons is essential for synaptic signaling, researchers report

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University of Illinois doctoral student Scott Siechen (left), mechanical science and engineering professor Taher Saif and their colleagues found that tension in axons is required for proper neuron signaling. Credit: Photo by L. Brian Stauffer, U. of I. News Bureau.

Every time a neuron sends a signal - to move a muscle or form a memory, for example - tiny membrane-bound compartments, called vesicles, dump neurotransmitters into the synapse between the cells. Researchers report that this process, which is fundamental to the workings of the nervous system, relies on a simple mechanical reality: Tension in the axon of the presynaptic neuron is required.



Without this tension, the researchers found, the vesicles that must haul their chemical cargo to the synapse for neuronal signaling would instead disperse.

The new findings appear this week in the <u>Proceedings of the National</u> <u>Academy of Sciences</u>.

"There is no controversy here," said University of Illinois mechanical science and engineering professor Taher Saif, who conducted the study with biology professor Akira Chiba, now at the University of Miami. Chiba's former doctoral student Scott Siechen and Saif's former doctoral student Shengyuan Yang also contributed significantly to the study. "We're not saying that you don't need chemical or <u>electrical signals</u> for the neurons to fire. All we're saying is that you also need tension in the axons."

The discovery was made almost by accident, Saif said. In a study of fruit fly embryos, Siechen wanted to know whether severing the growing end of an axon would prevent it from reaching its target, a nearby muscle cell.

After severing the axon, he watched the growing tip of the axon, called the growth cone, continue to grow toward and touch the muscle cell. But when he stained it, he noticed that the vesicles in the axon tip were dispersed, not clustered together near the synapse as they normally are.

He then repeated the experiment, but used a micropipette to pull on the severed end of the axon before staining it. This time the vesicles appeared at high density near the synapse.

This indicated that tension in the severed axon was somehow directing the vesicles to collect near the synapse, Saif said.



"The axon is physically cut off, chemically cut off, electrically cut off from the rest of the cell," he said. "So it appears that tension is all that was needed to keep the vesicles in place. But the question is, what keeps the vesicles there?"

Saif hypothesized that the <u>axons</u> in the embryo must be under tension. Otherwise - like a guitar string that is too loose to tune - it would not readily respond to changes in tension.

Using nanoscale probes developed in Saif's laboratory to gently deform an intact axon, the researchers found that the resting tension in a typical axon is about 1 nanonewton. (One newton has been described as the force required to hold a standard-sized apple against the pull of gravity; a nanonewton is one-billionth of that force.) Saif's hypothesis was thus proved true.

The researchers next turned their attention to the structure of the axon terminal, the region that lies closest to the synapse. This region contains many proteins, including actin, which is found in virtually all cell types and is known for maintaining cell shape and generating tension. Under the right conditions, individual actin molecules link together into ropelike filaments. Another protein, myosin, acts as a motor that connects the fibers and causes them to slide in relation to one another. This sliding can increase or decrease tension in the cell.

Other researchers have suggested that actin in axon terminals acts as a kind of scaffold that holds the vesicles near the synapse, Saif said. If that is true, he said, then tension also plays a significant role in the process. It appears that actin cannot properly scaffold the vesicles without sufficient tension in the axon terminal. Further research is needed to identify the exact mechanism that allows this to work, he said.

"This study shows that tension in neurons might be one of the parameters



so far overlooked in the quest for understanding learning and memory," Saif said. "We know from studies done elsewhere that tension in <u>neurons</u> creates folds in the brain, and it may be that a lack of tension in the neuron or a lack of the neuron's ability to generate tension is linked with memory loss or other neurological disorders."

Source: University of Illinois at Urbana-Champaign (<u>news</u> : <u>web</u>)

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