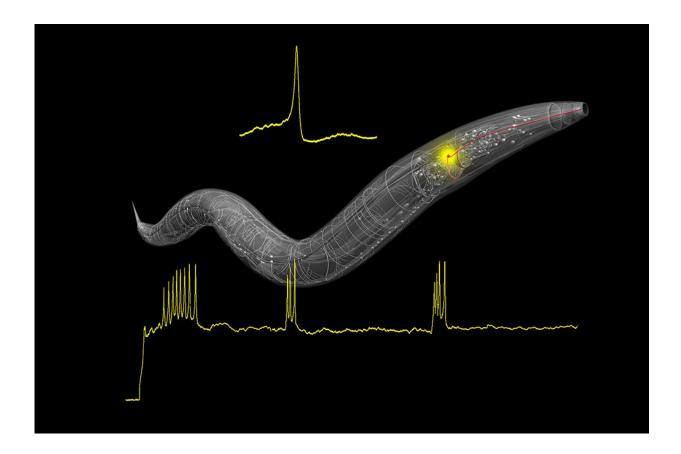


## In tiny worms, researchers find spiking neurons—and clues about brain computation

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Researchers have mapped all 302 neurons that make up the *C. elegans* nervous system. However, until now, they had never observed action potentials in these cells. Credit: Qiang Liu/The Rockefeller University

Contrary to popular belief, the brain is not a computer. However brains



do, in their own way, compute. They integrate informational inputs to generate outputs, including behaviors, thoughts, and feelings.

To process vast amounts of data, the brain uses a kind of digital code. Its cells produce discrete bursts of electric current, known as action potentials, that function as the zeros and ones of the nervous system. This code is assumed to be a vital aspect of computation in animals—that is, in most animals. The tiny roundworm *C. elegans* has long been considered a curious exception; until now, action potentials had never been observed in the organism.

But in a recent study, Rockefeller scientist Cori Bargmann and her colleagues, Qiang Liu, Phil Kidd, and May Dobosiewicz, discovered, among other things, a *C. elegans* olfactory neuron that produces action potentials. The finding, published in *Cell*, overturns decades of dogma and could help scientists understand fundamental principles of brain computation.

## Trial by fire

Neurons communicate with one another by exchanging chemical messages. Each message alters the state of the receiving cell; and as a neuron collects more and more chemical input, it approaches a threshold of activation. An action potential occurs when the cell reaches this threshold, at which point the neuron is said to "fire" or "spike" as an electrical impulse ripples through its extremity. In producing this spike, the cell translates analog chemical messages into digital electric code.

Despite the apparent importance of action potentials, for years researchers believed that *C. elegans* and other nematodes simply didn't use this information processing strategy.

"There's this whole class of animals where the neurons didn't seem to



spike," says Bargmann, the Torsten N. Wiesel Professor. "So our question was: Well, what *do* these neurons do?" Seeking an answer, her team set out to measure the electrical behavior of *C. elegans* neurons—every single one of them, if necessary.

"The *C. elegans* has just 302 neurons, so it's one of the few animals where you can look at each individual neuron," says Liu, a research assistant professor in Bargmann's lab who set out to measure how all of these neurons respond to stimulation.

Almost immediately, Liu was met with a surprise. While stimulating AWA, a neuron that processes smell signals, he observed that the cell's electrical voltage rose very rapidly before dramatically plummeting. Though unexpected, this dynamic was also very familiar: it looked like an action potential.

## A neuron with potential

Additional experiments confirmed that AWA neurons indeed spike. The researchers suspect that other *C. elegans* cells also produce action potentials; yet they note that this is not the norm for this animal's neurons. In fact, their experiments revealed that even AWA fires rather infrequently. Typically, the neuron responds to odors in a more subtle, graded manner. Liu observed action potentials only during experiments in which the stimulus grew stronger over time, suggesting that in nature, AWA fires when the animal is approaching the source of an important smell.

"Here we have a neuron that encodes information in two ways: one way that is slower and graded, and one way that's very nonlinear and sharply tuned to particular circumstances," says Bargmann. "And this lets us see what a spike might be uniquely important for."



While this study initiates *C. elegans* into the ranks of spike-producing animals, the <u>action potentials</u> observed in this organism were not identical to those seen elsewhere. To define the characteristics of wormspecific spikes, postdoctoral associate Phil Kidd created a mathematical model of AWA's electrical dynamics—a step that, the scientists hope, will allow their research to enter into conversation with other advances in computational neuroscience.

"There's a huge field of people working on the coding and computational principles of nervous systems," says Kidd. "And our work with *C*. *elegans* is likely to uncover principles that were unfamiliar to scientists who have been working in these areas for a long time."

This line of research indeed has the potential to both expand scientific understanding of *C. elegans*, and of nervous systems at large.

"Computation in the brain is a deep and important problem," says Bargmann. "With this study, we've shown that *C. elegans* can help solve this puzzle—and in fact, we've already exposed a whole new piece of it."

**More information:** Qiang Liu et al, C. elegans AWA Olfactory Neurons Fire Calcium-Mediated All-or-None Action Potentials, *Cell* (2018). DOI: 10.1016/j.cell.2018.08.018

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