

Research aims to unlock secrets of how neuronal variability is controlled by dendrites



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Three operational regimes. **a**, Left: stationary input potential resulting from the Green's function of a ball-and-stick model receiving localized input currents at the soma (x = 0, $I_s = 5$) and in the dendrite (x = 8, $I_d = 5$). Right: schematic illustration of spike-triggered effects. A somatic spike causes a refractory period in the soma and a BAP along the dendrite. A dendritic spike creates a short-lived depolarization in the soma. **b**,**c**, Firing frequency (**b**) and CV (**c**) against somatic input current for three dendritic input strengths. The black dashed line corresponds to the absence of dendritic spikes. The black dots correspond to points obtained using Monte Carlo simulation. **d**, CV against firing frequency for



three dendritic input strengths. **e**, Schematic of the two-compartment model with noisy input injections. **f**, f–I and CV–I curves highlighting the transition between three operational regimes: the Cox regime where the mean somatic input (I_{det}) plus the dendritic spike amplitude is below threshold, the bursting regime where the mean somatic input plus dendritic spike amplitude is above threshold, and the mean-driven regime where the mean somatic input alone is above threshold. For this simulation, the dendritic spike amplitude was set to D = 0.3. **g**, Voltage traces corresponding to the three regimes. **h**, The strength of the somatic input associated with the peak CV (Idet*{I}_{\text{I}}^{tm}{det}}^{t*}) shown against the amplitude of the dendritic spike. Credit: *Nature Computational Science* (2023). DOI: 10.1038/s43588-023-00580-6

The inner workings of the human brain are a gradually unraveling mystery and Dr. Richard Naud of the University of Ottawa's Faculty of Medicine has led a highly compelling new study that brings us closer to answering these big questions.

The study's results have important implications for theories of learning and working memory and could potentially help lead to future developments in <u>artificial intelligence</u> (AI) since AI developers and programmers watch the work of Dr. Naud and other leading neuroscientists.

Published in *Nature Computational Science*, <u>the study</u> tackles the manylayered mystery of the "response variability" of neurons, <u>brain cells</u> that use electric signals and chemicals to process information and green lights all the remarkable aspects of human consciousness.

The findings unveil the nuts and bolts of how neuronal variability is controlled by dendrites, the antenna that reaches out from each neuron to receive synaptic inputs in our own personal neural communication networks. The rigorous study establishes properties of dendrites that



potently control output variability, a property that's been shown to control synaptic plasticity in the brain.

"The intensity of a neuron's response is controlled by inputs to its core, but the variability of a neuron's response is controlled by the inputs to its little antennas—the dendrites," says Dr. Naud, an Associate Professor at the Faculty of Medicine's Department of Cellular and Molecular Medicine and the uOttawa Department of Physics.

"This study establishes more precisely how single neurons can have this crucial property of controlling response <u>variability</u> with their inputs."

Dr. Naud suspected that if a mathematical framework he'd used to describe the cell body of neurons was extended to take their dendrites into account, then they might have luck efficiently simulating networks of neurons with active dendrites.

Cue the contribution of Zachary Friedenberger, a Ph.D. student at the Department of Physics and a member of Dr. Naud's lab, with a background in <u>theoretical physics</u> to solve the theoretical challenges and the math in a record time. Fast forward to the completed study: The model predictions were validated by analysis of in vivo recording data and observed over a wide range of model parameters.

"He managed to solve the math in a record time and solved a number of theoretical challenges I had not foreseen," Dr. Naud says.

Dr. Naud believed that their technique could provide insight into the neuronal response to variable inputs. So, they began working on a technique that would be able to compute statistics from a neuronal model with an active <u>dendrite</u>.

One of the work's reviewers noted that the <u>theoretical analysis</u> "provides



key insight into biological computation and will be of interest to a broad audience of computational and experimental neuroscientists."

More information: Zachary Friedenberger et al, Dendritic excitability controls overdispersion, *Nature Computational Science* (2023). DOI: 10.1038/s43588-023-00580-6

Provided by University of Ottawa

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