

Diversity of cortical neurons captured in comprehensive computer models

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The Allen Institute for Brain Science has produced the first comprehensive, publicly available database of predictive neuron models, along with their corresponding data. The generalized leaky integrate-and-



fire (GLIF) and biophysically detailed models are described in two articles published in the journal *Nature Communications*.

"The publication of these mathematical-physical models of the individual components making up neural networks is an important landmark in our ten-year quest to understand the <u>brain</u>" said Christof Koch, Ph.D., President and Chief Scientist at the Allen Institute for Brain Science. "We now seek to understand how vast assemblies of these elements give rise to behavior, perception and the feeling of life itself consciousness."

The GLIF and biophysical models were built from extensive data in the Allen Cell Types Database: a massive publicly available repository of cortical neurons in both the mouse and human brain. Launched in 2015, this database contains electrophysiological, morphological, and transcriptomic properties gathered from individual cells, and models simulating cell activity, all building toward a "periodic table" of <u>cell</u> types.

"This project is a testament to the Allen Institute's interdisciplinary collaboration and teamwork," said Nathan Gouwens, Ph.D., an Allen Institute scientist. "Five years ago, we all began working together to build the pipeline for the Allen Cell Types Database. Now we have a parallel pipeline for generating models for each cell in the database."

GLIF models reproduce the spike times of neurons and capture the abstract transformations taking place inside of neurons. These models have the benefit of requiring a relatively low level of computer power to simulate, meaning that it's feasible to model the millions and millions of neurons making up the brain of a mouse. The biophysical models are more detailed. As such, they accurately represent the mechanisms reproducing the actual voltage waveform of action potentials (nerve pulses) and other forms of electrical activity across the dendritic tree and



the cell body of neurons. Biophysical models take considerably more computational power than GLIF models to execute.

"We have generated a large library of cell models - ranging from simple to complex - which can be put together like building blocks to construct higher-level circuit models, providing a valuable resource for the neuroscience community," said Anton Arkhipov, Ph.D., an Allen Institute scientist.

"The models can be used as building blocks for larger simulations, but also to understand how some cell types differ from the other. Researchers can classify cell types by only looking at model parameters," said Stefan Mihalas, an Allen Institute scientist.

Perhaps in the future, detailed models of multiple cell types can be used in larger simulations to model neurological or psychiatric disorders, such as epilepsy, autism or Alzheimer's, and applying virtual perturbations to cells or networks of <u>cells</u> may allow us to see how the brain might respond to specific therapies.

"These studies make great use of the Allen's unique single-cell physiology data base by putting all recorded responses into a single model framework," said Adrienne Fairhall, Ph.D., co-director of the Computational Neuroscience program at the University of Washington. "This can play a huge role in allowing researchers to compare and potentially differentiate the computations of different cell types."

"Our models represent a large step in enabling scientists around the world to use standardized data, methods and models for exploring network behavior in the brain," said Corinne Teeter, Ph.D., the lead scientist for the GLIF modeling effort at the Allen Institute.

More information: Nathan W. Gouwens et al, Systematic generation



of biophysically detailed models for diverse cortical neuron types, *Nature Communications* (2018). DOI: 10.1038/s41467-017-02718-3

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