

Cell-to-cell diversity is key to protecting brain from neurological diseases, shows study

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A brain's cell makeup may transform the way we understand and treat



neuropsychiatric conditions, potentially opening a new treatment path for diseases like epilepsy, autism, schizophrenia, and depression.

The study from the University of Ottawa's Faculty of Science published in *Proceedings of the National Academy of Sciences* demonstrates how a brain's ecosystem and its <u>diversity</u> of cells can allow it to maintain normal function while better responding to changes, perturbations, or injury. The response to such "insults"—as the study coins them—is amplified by cells that are too connected or too alike, which makes them less resilient and unable to maintain the activity needed to preserve <u>brain</u> <u>function</u>.

Inspired by Charles Darwin's Origin of Species and the idea that biodiversity of natural ecosystems is the key to survival, lead author Jérémie Lefebvre and his team used mathematical models of brain circuitry to analyze how <u>neurons</u> respond to "insults" and how cell-to-cell diversity helps prevent failures.

"Our results demonstrate how diversity is a foundational, essential ingredient in the constitution of brain circuits," says Lefebvre, a member of uOttawa's Brain and Mind Research Institute. "Just like the biodiversity of ecosystems, the diversity of neurons is as important and represents a new way of interpreting certain illnesses, like epilepsy."

Epilepsy is characterized by recurrent seizures during which brain circuits go haywire and cells become too synchronous with one another. Lefebvre and the team of researchers, which includes members from the University Health Network's Krembil Brain Institute, used electrical recordings to explore the <u>brain cells</u> of patients with epilepsy, finding that cells from the region of the brain that create seizures look more alike and less diverse. Low diversity prevents neurons and cells from adapting adequately as brain circuits are unable to maintain healthy activity, leading to seizures.



"Using our interdisciplinary methods, we were able to show that brain cells must be different from each other to ensure the brain maintains its function in the face of time and changes," says Lefebvre, an Associate Professor in the Department of Biology. "This is an important reminder of the critical role diversity plays in <u>natural systems</u> and how it responds to inevitable change. This not only applies to ecosystems, neurons, and <u>neural circuits</u>, but to humans and communities."

The research may also provide guidance for treatment of autism, schizophrenia, and depression, which are also states of the brain marked by over connectivity and synchronicity.

More information: Axel Hutt et al, Intrinsic neural diversity quenches the dynamic volatility of neural networks, *Proceedings of the National Academy of Sciences* (2023). DOI: 10.1073/pnas.2218841120

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