

Computational model reveals how the brain manages short-term memories

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If you've ever forgotten something mere seconds after it was at the forefront of your mind—the name of a dish you were about to order at a restaurant, for instance—then you know how important working memory is. This type of short-term recall is how people retain information for a matter of seconds or minutes to solve a problem or



carry out a task, like the next step in a series of instructions. But, although it's critical in our day-to-day lives, exactly how the brain manages working memory has been a mystery.

Now, Salk scientists have developed a new computational model showing how the brain maintains information short-term using specific types of neurons. Their findings, published in *Nature Neuroscience* on December 7, 2020, could help shed light on why working <u>memory</u> is impaired in a broad range of neuropsychiatric <u>disorders</u>, including schizophrenia, as well as in normal aging.

"Most research on working memory focuses on the excitatory neurons in the cortex, which are numerous and broadly connected, rather than the inhibitory neurons, which are locally connected and more diverse," says Terrence Sejnowski, head of Salk's Computational Neurobiology Laboratory and senior author of the new work. "However, a recurrent neural network model that we taught to perform a working memory task surprised us by using inhibitory neurons to make correct decisions after a delay."

In the new paper, Sejnowski and Robert Kim, a Salk and UC San Diego MD/Ph.D. student, developed a computer model of the prefrontal cortex, an area of the brain known to manage working memory. The researchers used learning algorithms to teach their model to carry out a test typically used to gauge working memory in primates—the animals must determine whether a pattern of colored squares on a screen matches one that was seen several seconds earlier.

Sejnowski and Kim analyzed how their model was able to perform this task with high accuracy, and then compared it to existing data on the patterns of brain activity seen in monkeys carrying out the task. In both tests, the real and simulated neurons involved in working memory operated on a slower timescale than other neurons.



Kim and Sejnowski found that good working memory required both that long-timescale neurons be prevalent, and that connections between inhibitory neurons—which suppress <u>brain</u> activity—be strong. When they altered the strength of connections between these inhibitory neurons in their model, the researchers could change how well the <u>model</u> performed on the working memory test as well as the timescale of the pertinent <u>neurons</u>.

The new observations point toward the importance of <u>inhibitory neurons</u>, and could inspire future research on the role of these cells in working memory, the researchers say. They also could inform studies on why some people with neuropsychiatric disorders, including schizophrenia and autism, struggle with working memory.

"Working memory impairment is common in neuropsychiatric disorders, including schizophrenia and autism spectrum disorders," says Kim. "If we can elucidate the mechanism of working memory, that's a step toward understanding how working memory deficits arise in these disorders."

More information: Robert Kim et al, Strong inhibitory signaling underlies stable temporal dynamics and working memory in spiking neural networks, *Nature Neuroscience* (2020). <u>DOI:</u> 10.1038/s41593-020-00753-w

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