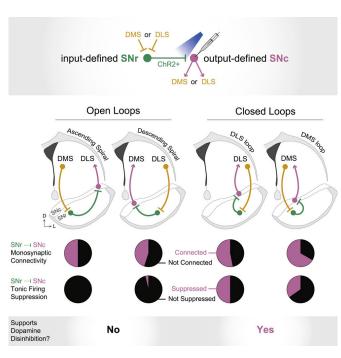


Investigating the role of dopamine circuits in habit formation

25 August 2022, by Melissa Rohman



Graphical abstract. Credit: *Cell Reports* (2022). DOI: 10.1016/j.celrep.2022.111228

Scientists have uncovered how dopamine connects subregions of the striatum essential for habit formation, according to a Northwestern Medicine study published in *Cell Reports*, findings that may change the overall understanding of how habits are formed—and could be broken.

"The <u>circuits</u> are going both ways, and so maybe that gives potential ideas of how to break a bad habit if there's a circuit allowing <u>information flow</u> to go the other way," said Talia Lerner, Ph.D., assistant professor of Neuroscience and senior author of the study.

The <u>basal ganglia</u> are a group of nuclei below the brain's cortex, which operate primarily in closed parallel loops. These loops, or circuits, originate from different regions of the dorsal striatum,

including an associative circuit controlling goaldirected behaviors from the dorsomedial striatum and a somatosensory circuit controlling habit formation from the dorsolateral striatum. In addition to closed loops, it has long been hypothesized that there is an open "ascending spiral" loop that allows goal-directed behaviors to become habits.

Lerner and her team set out to investigate the "ascending spiral hypothesis." In this circuit, signals in the dorsomedial striatum travel to another brain area called the <u>substantia nigra</u>, which contains most of the brain's dopamine neurons. The signals were hypothesized to make dopamine neurons fire and cause release of dopamine in the dorsolateral striatum. Such <u>dopamine</u> release is required for habit formation.

Although the ascending spiral hypothesis was developed 20 years ago, it has been difficult to test. Using new transsynaptic and intersectional genetic tools, the scientists were able to complete a careful investigation.

"The exciting thing in this study is that we were able to use <u>new tools</u> to trace both forward circuits and backward circuits and really test if there are synaptic connections where we think they should be and if they work in the way we think they should," Lerner said.

The evidence they found for the ascending spiral hypothesis was mixed.

"The ascending spiral is there, however, it doesn't work the way that way that it was hypothesized," said Priscilla Ambrosi, a student in Northwestern University's Interdepartmental Neuroscience (NUIN) Ph.D. program and lead author of the study.

In addition to discovering new details about how the ascending spiral circuit may work, the authors also found evidence for a "descending spiral," which would potentially communicate information the



other direction—from habit-related to goal-directed subregions of the <u>striatum</u>.

"People didn't really think this 'descending' spiral existed, but we found that it does and there's actually equivalent evidence for it compared to the ascending spiral," Lerner said. "Now, we have to figure out how these ascending and descending circuits really work. We're interested in the idea that they gate information from other places in the brain and perhaps control when different synapses strengthen or weaken during habit formation," Lerner said.

More information: Priscilla Ambrosi et al, Striatonigrostriatal circuit architecture for disinhibition of dopamine signaling, *Cell Reports* (2022). DOI: 10.1016/j.celrep.2022.111228

Provided by Northwestern University

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