

Worm study shows dopamine-regulated brain pathway may be related to anxiety

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Worm (Caenorhabditis elegans) approaching two predators (Pristionchus pacificus) guarding a patch of food. Credit: Salk Institute

The life of the tiny worm called Caenorhabditis elegans consists mostly



of looking for food, eating food, and laying eggs. So, when any of these behaviors are disrupted, there's cause for concern. In a new study, Salk Institute scientists discovered that the "feel good" brain chemical dopamine regulates anxious worm behavior in the presence of nipping predators.

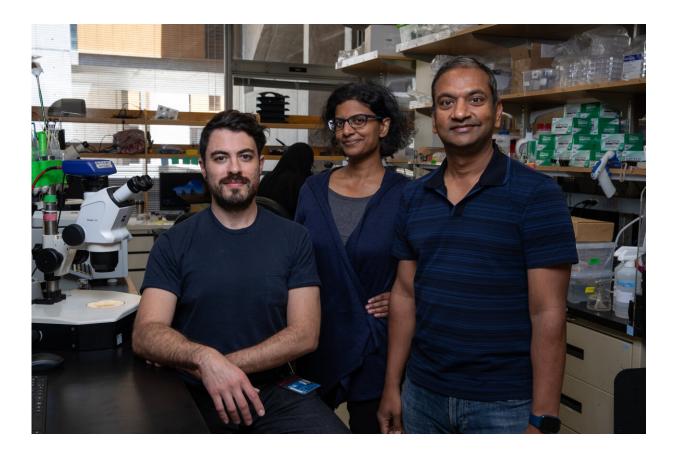
The findings, published in *eLife* on July 11, 2023, illuminate how this dopamine-regulated brain pathway may be related to anxiety and could provide insight into human conditions, such as <u>post-traumatic stress</u> <u>disorder</u> (PTSD).

"Worms are a wonderful model for studying anxiety because the cause of their anxiety is typically known, like predation," says Professor Sreekanth Chalasani, senior author of the new work. "Thus, we can use <u>worms</u> to better understand fundamental neural pathways related to anxiety and stress responses that could be conserved across species."

In this study, the researchers observed what happened when a <u>predator</u> worm species (Pristionchus pacificus) nipped at the worms, discouraging the worms from eating the food source. In response, the worms moved away from the food source, and the predators were able to eat more. Additionally, the worms also laid their eggs away from the old food source and the threat of predation.

The researchers noticed that the worms would stay away even after the predators had left—indicating the worms were learning that it was safer to stay away. Furthermore, the behavior of laying the eggs far away from predators was regulated by dopamine. But when the researchers blocked dopamine pathways in the worms, they did not avoid predators as frequently to lay their eggs.





From left: Michael Rieger, Kirthi Reddy, and Sreekanth Chalasani. Credit: Salk Institute

"To understand how the brain works, it is important to study it in its natural context," says co-first author Amy Pribadi, a former graduate student researcher in Chalasani's lab. "To do that, we can use a simple worm with an easily constructed natural environment, then look at how that worm's brain networks and molecules modulate behavior in that 'natural' environment."

Additionally, the researchers explored how the presence of smaller patches of food away from the main food source impacted wormpredator interactions. While predators were busy monopolizing a main <u>food source</u>, the presence of alternate (albeit smaller) food sources



elsewhere made the worms more likely to avoid predators.

"Worms make decisions based on <u>environmental changes</u> similar to more complex animals and humans," says co-first author Michael Rieger, a postdoctoral researcher in Chalasani's lab. "Difficulty adapting to change—especially due to threat or stress—is common in humans, especially those with neurological disorders. Our research in this <u>simple</u> <u>organism</u> reveals new avenues for exploring the neuroscience of decisionmaking, which we hope can generalize all the way up the food chain to humans."

In the future, the team will examine the role of another brain chemical called serotonin in this evasive behavior, since serotonin is known to regulate other dopamine-dependent behaviors, like searching for <u>food</u>. They also hope their study serves as a springboard for future inquiry into the impact of other brain signaling pathways that influence feeding, reproduction, and other behaviors.

"By looking at worms, you capture so much intricate biology in an animal that has evolved over billions of years," says Chalasani. "Any insight—even in worms—into the mechanisms of predator-prey behaviors enriches our understanding of so many other phenomena that exhibit this biological push-pull relationship, like the co-evolution of animals and their environment or the delicate balance of chemical conversations between cells."

More information: Amy Pribadi et al, Dopamine signaling regulates predator-driven changes in Caenorhabditis elegans' egg laying behavior, *eLife* (2023). <u>DOI: 10.7554/eLife.83957</u>

Provided by Salk Institute



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