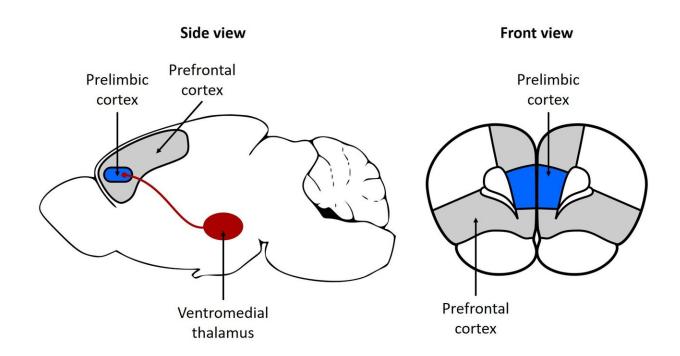


Mapping the decision-making pathways in the brain

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Neurons from the ventromedial thalamus (red area) extend into the prelimbic cortex (blue area) which is a region within the prefrontal cortex (grey area). Credit: OIST

Every day, people make hundreds of decisions. While most are small and inconsequential, like choosing what to eat or wear, others are more complex and involve weighing up potential costs and benefits, like deciding whether to study more for a better grade instead of hanging out



with friends.

Decisions like these are shaped by individual values and preferences, but the <u>brain activity</u> involved in these choices is still not well understood. Now, scientists at the Okinawa Institute of Science and Technology Graduate University (OIST) have identified a new area of the <u>brain</u> that could be involved in cost-benefit decision-making.

"Previously, many neuroscientists believed that each area of the brain carried out a specific function, such as recognizing faces, memory or movement," said first author Dr. Bianca Sieveritz, former Ph.D. student and now junior research fellow in the OIST Brain Mechanism for Behavior Unit. "But in more recent years, we've realized that it's far more complex, and that <u>cognitive processes</u> are carried out by a distributed network across the brain, with many different brain areas communicating."

To understand how cognitive abilities like decision-making work, scientists first need to figure out which parts of the brain are connected to each other. These connections are formed by specialized cells called neurons. Neurons have long, thin projections that can reach out and send signals to other neurons located across regions of the brain.

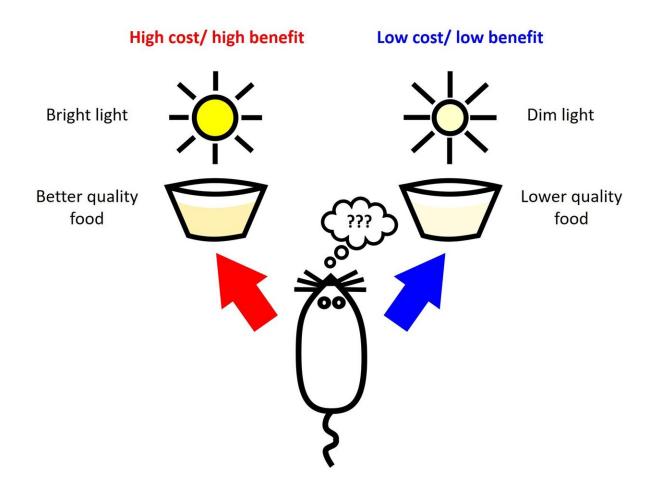
But mapping these connections is not easy. "It's not as simple as identifying that one brain area is connected to another," explained Dr. Sieveritz. "Within each brain region, there are many different types, or sub-classes, of neurons, which you need to be able to identify, and each neuronal sub-class might only connect to one other area of the brain or be involved in one specific function. So it's very messy and complicated."

In this study, published in *Brain Structure and Function*, Dr. Sieveritz examined an area of the brain known as the ventromedial thalamus. This



region of the brain is predominantly involved in movement, with abnormal function in this area associated with symptoms of Parkinson's. But many neuroscientists believe that this area may have other undiscovered functions.

While many neurons from the ventromedial thalamus connect to the motor cortex, other neurons reach further into the prefrontal cortex. This area lies right at the front of our brain and plays a major role in more complex cognitive behaviors, including expression of personality and understanding of language.



In a simple behavioral experiment, rats choose between better quality food that is delivered in conjunction with an uncomfortably bright light, or lower quality



food with a less bright light. Signals from the ventromedial thalamus may affect which choice the rats make. Credit: OIST

But the prefrontal <u>cortex</u> is huge and composed of millions of neurons, so this research focused in on a tiny area within the prefrontal cortex—the prelimbic cortex. Located right where the two brain hemispheres meet, the prelimbic cortex has previously been found to play a role in fear conditioning, working memory and decision-making.

Working with brain slices from rats, Dr. Sieveritz used different chemical markers to stain the ends of the neurons from the ventromedial thalamus as well as the ends of the different neurons within the outermost layer of the prelimbic cortex. She then pinpointed where the ends of the neurons met each other and formed a connection point, or synapse.

She found that most connections formed within the prelimbic cortex were with a specific sub-class of neurons called cortico-striatal neurons.

"Cortico-striatal neurons in the prelimbic cortex are important for costbenefit decision-making," said Dr. Sieveritz. "If the ventromedial thalamus is sending signals to these cortico-striatal neurons, this could mean that the ventromedial thalamus plays a role in cost-benefit decisionmaking too."

Interestingly, Dr. Sieveritz noticed that neurons from the ventromedial thalamus were also sending signals to inhibitory neurons within the prelimbic cortex. Inhibitory neurons slow down or stop other neurons from firing and are essential for keeping the brain's activity under careful control.



"We know that these inhibitory neurons actually extend down into the cortex and connect to the cortico-striatal neurons in deeper layers of the brain and possibly inhibit their activity," said Dr. Sieveritz. "The connections between the neurons from the ventromedial thalamus and the inhibitory neurons therefore may provide a way of finetuning the activity of the cortico-striatal neurons."

Dr. Sieveritz stressed that while her results implicate the ventromedial thalamus in cost-benefit decision-making, further research is needed to uncover its exact role in this cognitive process. The team now plans to conduct behavioral studies in rats to determine if signals from the ventromedial thalamus change whether rats choose higher quality food at a higher cost, or lower quality food at a lower cost.

This research brings scientists one step closer to understanding the circuitry underlying the complex process of cost-benefit decision-making. "Neuroscience is like a huge jigsaw puzzle; everyone can only do a tiny part," said Dr. Sieveritz. "But when all our research is combined, hopefully we will start to see the bigger picture."

More information: Bianca Sieveritz et al, Prelimbic cortical targets of ventromedial thalamic projections include inhibitory interneurons and corticostriatal pyramidal neurons in the rat, *Brain Structure and Function* (2020). DOI: 10.1007/s00429-020-02109-3

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