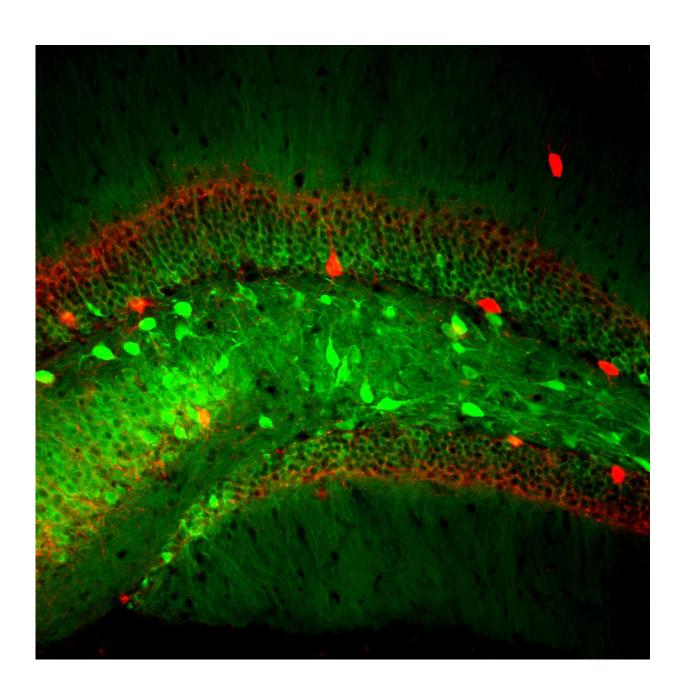


Mice move through the virtual world of a video game and provide insight into the mechanisms of memory formation

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A look into memory: The video recording shows that the neurons begin to flash as soon as they are activated. Credit: Thomas Hainmüller, Marlene Bartos

Dr. Thomas Hainmüller and Prof. Dr. Marlene Bartos of the Institute of Psychology of the University of Freiburg have established a new model to explain how the brain stores memories of tangible events. The model is based on an experiment involving mice seeking a place where they received rewards in a virtual environment. The scientific journal *Nature* has published the study.

Take a few steps forward, stop, and look around. In a video game, the walls that depict a corridor four meters long are made up of green and blue patterned blocks. The floor is marked with turquoise dots. A short distance away, there's a brown disc on the floor that looks like a cookie. That's the symbol for the reward location. The mouse heads for it, arrives, and the symbol disappears. The next cookie promptly appears a bit further down the corridor. The mouse is surrounded by monitors and is standing on a styrofoam ball that is floating on compressed air and turns beneath the mouse when it runs. The ball makes it possible to transfer of the mouse's movements to the virtual environment. If the mouse reaches the reward symbol, a straw is used to give it a drop of soy milk and stimulate it to form memories of its experiences in the virtual world. The mouse learns when, and at which location, it will receive a reward. It also learns how to locate itself and discriminate between different corridors in the video game.

"As the mouse is getting to know its environment, we use a special microscope to look from the outside into its brain and we record the activities of its nerve <u>cells</u> on video," explains Thomas Hainmüller, a



physician and doctoral candidate in the MD/Ph.D. program of the Spemann Graduate School of Biology and Medicine (SGBM) of the University of Freiburg. He says that works because, in reality, the head of the mouse remains relatively still under the microscope as it runs through the virtual world of the video game. On the recordings, the mice's genetically-manipulated nerve cells flash as soon as they become active. Hainmüller and Marlene Bartos, a Professor of Systemic and Cellular Neurobiology are using this method to investigate how memories are sorted and retrieved. "We repeatedly place the mouse in the virtual world on consecutive days," says Hainmüller. "In that way, we can observe and compare the activity of the nerve cells in different stages of memory formation," he explains.

The hippocampus plays a decisive role in the formation of memory episodes, or memories of tangible experiences. In their study, published in the scientific journal *Nature*, they demonstrate that the nerve cells in the hippocampus create a map of the <u>virtual world</u> in which single neurons code for actual places in the video game. Earlier studies done at the Freiburg University Medical Center showed that nerve cells in the human hippocampus code video games in the same way. The cells become activated and flash when the mouse is at the respective place, otherwise they remain dark. "To our surprise, we found very different maps inside the hippocampus," reports Hainmüller. In part, they provide an approximate overview of the position of the mouse in the corridor, yet they also consider time and context factors, and above all, information about in which of the corridors the <u>mouse</u> is located. The maps are also updated during the days of the experiment and as a result can be recognized as a learning process.

The research team says their observations provide a model that explains how activity of the <u>nerve cells</u> in the hippocampus can map the space, time and and context of memory episodes. The findings allow for better understanding of the biological processes that affect the formation of



memory in the brain. Hainmüller says, "In the long term, we would like to use our results to contribute to the development of treatments to help people with neurological and psychiatric illnesses."

More information: Thomas Hainmueller et al, Parallel emergence of stable and dynamic memory engrams in the hippocampus, *Nature* (2018). DOI: 10.1038/s41586-018-0191-2

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