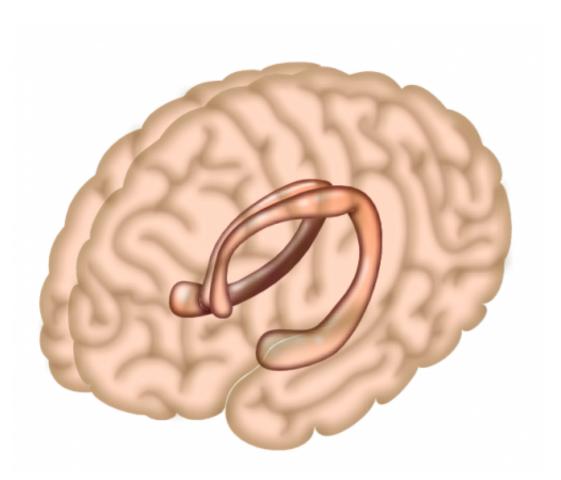


## Bravery-associated cells identified in the hippocampus

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The hippocampus is a region of the brain largely responsible for memory formation. Credit: Salk Institute

Why can some people comfortably walk between skyscrapers on a highwire or fearlessly raft Niagara Falls in a wooden barrel, whereas others



freeze at the mere thought of climbing off escalators in a shopping mall? In a new study, scientists have found that a certain type of cell in the hippocampus plays a key role.

People differ when it comes to trying dangerous or exhilarating activities. Even siblings can show dramatic differences in risk-taking behaviour. The neural mechanisms that drive risk-taking behaviour are largely unknown. However, scientists from the Department of Neuroscience of Uppsala University in Sweden and the Brain Institute of the Federal University of Rio Grande do Norte in Brazil have found that some cells in the hippocampus play a key role in risk-taking behaviour and anxiety.

In an article published in the journal *Nature Communications*, the authors show that neurons known as OLM cells, when stimulated, produce a brain rhythm that is present when animals feel safe in a threatening environment (for example, when they are hiding from a predator but aware of the predator's proximity). The study, produced by Drs. Sanja Mikulovic, Ernesto Restrepo, Klas Kullander and Richardson Leao, among others, showed that anxiety and risk-taking behaviour can be controlled by the manipulation of OLM cells. To find a pathway that quickly and robustly modulates risk-taking behaviour is very important for treatment of pathological anxiety, since reduced risk-taking behaviour is a trait in people with high anxiety levels.

Adaptive (or normal) anxiety is essential for survival because it protects us from harm. Unfortunately, in a large number of people, anxiety can be dysfunctional and severely interfere with daily life. In these cases, doctors often rely on antidepressants to help patients recover from the dysfunctional state. However, these drugs act in the entire brain and not only in the areas where it is needed, and may therefore cause severe sideeffects. Thus, drugs that affect a single brain region or a very specific group of cells may be a major breakthrough in treating anxiety and



associated disorders like depression. Another interesting finding in the study is that OLM cells can be controlled by pharmacological agents. In the past, the same group of scientists found that OLM cells were the gatekeepers of memories in the hippocampus, and that these cells were very sensitive to nicotine.

"This finding may explain why people binge-smoke when they are anxious," says Dr. Richardson Leao, researcher at the Brain Institute of the Federal University of Rio Grande do Norte.

The participation of the hippocampus in emotions is much less studied than its role in memory and cognition. In 2014, for example, the Nobel prize was awarded for the discovery of "place cells" that represent a biological GPS and underlie the memories of where we are located in our surroundings. In the past decade, scientists have also started to appreciate the role of the <u>hippocampus</u> in regulating emotions.

"It is fascinating how different regions of the same brain structure control distinct behaviours and how they interact with each other. Identifying specific circuits that underlie either cognitive or emotional processes is crucial for the general understanding of brain function and for more specific drug development to treat disorders," says Dr. Sanja Mikulovic, Uppsala University.

The discovery of these neurons and their role in <u>anxiety</u> and risk-taking may open a path for the development of highly efficient anxiolytics and antidepressants without common side-effects, such as apathy.

**More information:** Sanja Mikulovic et al, Ventral hippocampal OLM cells control type 2 theta oscillations and response to predator odor, *Nature Communications* (2018). DOI: 10.1038/s41467-018-05907-w



## Provided by Uppsala University

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