

How the brain hears and fears



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These heatmaps show activity in 756 neurons found in the BLA region of a mouse brain. Spikes in excitement (red) represent responses over time to neutral sounds (c & d), a water reward (e), or a startling puff of air. Maps like these also show how the brain inhibits (blue) an undesired reaction. Black represents moments of non-responsiveness. Credit: Li Lab, CSHL/2018

How is it that a sound can send a chill down your spine? By observing individual brain cells of mice, scientists at Cold Spring Harbor Laboratory (CSHL) are understanding how a sound can incite fear.



Investigator Bo Li focuses on a part of the mouse brain called the amygdala where sights, sounds, and other stimuli take on positive or negative associations through experience. The continuous process of learning and unlearning that occurs in the amygdala appears impaired in people with anxiety disorders or major depression. Understanding brain cell, or neuron activity in the amygdala could result in better treatments.

In the October issue of *Nature Communications*, Li and postdoctoral researcher Xian Zhang describe profound changes in neuron activity when they trained animals to fear a particular sound and associate another sound with a <u>reward</u>. "If you look at the patterns of brain cell activity in the amygdala, you can know whether the animal is expecting a reward or fearing a punishment," Li explains.

Li and Zhang used a microscope with a lens small enough to implant in the brain of a mouse, to track the firing activity of specific neurons before, during, and after an animal's training. They taught the animals to associate particular sounds with reward or punishment and saw the behavior of neurons evolve. The experiment associated one tone with an annoying puff of air—the punishment. The reward tone was a refreshing drop of water to drink.

At first, neurons sensitive to sound responded to each tone by firing randomly. But when one tone was repeatedly accompanied by the puff of air, the neurons fired in a very specific pattern. This pattern closely resembled the firing pattern of another type of <u>brain</u> cell that fires when the mouse actually experienced the punishment. Likewise, when a tone was repeatedly paired with a sip of water, the sound-sensitive neurons fired in a pattern similar to <u>neuron activity</u> when the mouse received the water reward.

As the firing patterns became more specific, the animals licked in response to the reward-associated tone—anticipating water. They



blinked in response to the punishment-associated sound—anticipating an air puff.

The researchers also switched the meaning of each tone. When the "reward" sound was repeatedly accompanied by an air puff, the <u>neurons</u> let go of the established "reward" firing pattern and adopted the "<u>punishment</u>" pattern. "We think this is how <u>sound</u> acquires meaning," Li says.

More information: Xian Zhang et al, Population coding of valence in the basolateral amygdala, *Nature Communications* (2018). DOI: 10.1038/s41467-018-07679-9

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