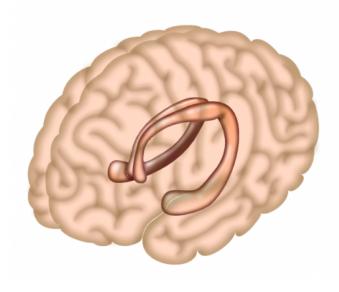


## Study points to critical periods in early-life learning for brain development

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

A new study on infantile memory formation in rats points to the importance of critical periods in early-life learning on functional development of the brain. The research, conducted by scientists at New York University's Center for Neural Science, reveals the significance of learning experiences over the first two to four years of human life; this is when memories are believed to be quickly forgotten—a phenomenon known as infantile amnesia.

"What our findings tell us is that children's brains need to get enough and healthy activation even before they enter pre-school," explains Cristina Alberini, a professor in NYU's Center for Neural Science, who led the study. "Without this, the neurological system runs the risk of not properly developing learning and memory functions."

The other authors of the study, conducted in collaboration with the Icahn School of Medicine at

Mt. Sinai, included: Alessio Travaglia, a post-doctoral researcher at NYU; Reto Bisaz, an NYU research scientist at the time of the study; Eric Sweet, a post-doctoral fellow at the Icahn School of Medicine at Mt. Sinai; and Robert Blitzer, a professor at the Icahn School of Medicine at Mt. Sinai.

In their study, which appears in the journal *Nature Neuroscience*, the researchers examined the mechanisms of infantile memory in <u>rats</u>—i.e., memories created 17 days after birth. This is the equivalent of humans under the age of three and when memories of who, what, when, and where—known as episodic memories—are rapidly forgotten. The phenomenon, referred as to "infantile or childhood amnesia," is in fact the inability of adults to retrieve episodic memories that took place during the first two to four years of life.

In addressing this matter, Alberini and her colleagues compared rats' infantile memory with that when they reached 24 days old—that is, when they are capable of forming and retaining <a href="long-term">long-term</a> memories and at an age that roughly corresponds to humans at six to nine years old.

The episodic memory tested in the rodents was the memory of an aversive experience: a mild foot shock received upon entering in a new place. Adult rats, like humans, remember unpleasant or painful experiences that they had in specific places, and then avoid returning to them.

To do so, rodents were placed in a box divided into two compartments: a "safe" compartment and a "shock" compartment. During the experiment, each rat was placed in the safe compartment with its head facing away from the door. After 10 seconds, the door separating the compartments was automatically opened, allowing the rat access to the shock compartment. If the rat entered the shock compartment, it received a mild foot shock.



The first set of results was not surprising. The authors found infantile amnesia for the 17 day-old rats, which showed avoidance of the "shock" compartment right after the experience, but lost this memory very rapidly: a day later these rats quickly returned to this compartment. In contrast, the rats exposed to the shock compartment at 24 days of life learned and retained the memory for a long time process and store memories long-term. and avoided this place—revealing a memory similar to that of adult rats.

However, remarkably, the younger rats, which had apparently forgotten the initial experience, of the memory. When, later in life, these rats were with recollections of the context and the foot shock—they indicated having a specific memory, which was revealed by their avoidance of the specific context in which they received a shock at day 17 of life. These findings show how early life experience, although not expressed or remembered, can influence adult life behavior.

The findings raised the following question: what is occurring—neurologically—that explains why memories are retained by the younger rats only in a nature.com/articles/doi:10.1038/nn.4348 latent form but are stored and expressed long-term by older ones? Or, more specifically, what occurs during development that enhances the ability to form lasting memories?

To address this, the scientists focused on the brain's hippocampus, which previous scholarship has shown is necessary for encoding new episodic memories. Here, in a series of experiments similar to the box tests, they found that if the hippocampus was inactive, the ability of younger rats to form latent memories and recall them later by reminders as they got older was diminished. They then found that mechanisms of "critical periods" are fundamental for establishing these infantile memories.

A critical period is a developmental stage during which the nervous system is especially sensitive to environmental stimuli. If, during this period, the organism does not receive the appropriate stimuli required to develop a given function, it may be difficult or even impossible to develop that function

later in life. Well-known examples of critical periodbased functions are sensory functions, like vision, and language acquisition.

The study shows that there is a critical period for episodic learning and that during this period the hippocampus learns to become able to efficiently

"Early in life, while the brain cannot efficiently form long-term memories, it is 'learning' how to do so, making it possible to establish the abilities to memorize long-term," explains Alberini. "However, subsequently showed they actually had kept a trace the brain needs stimulation through learning so that it can get in the practice of memory prompted with reminders—i.e., they were presented formation—without these experiences, the ability of the neurological system to learn will be impaired."

> These studies, the researchers observe, suggest that using learning and environmental interventions during a critical period may significantly help to address learning disabilities.

More information: Infantile amnesia reflects a developmental critical period for hippocampal learning, Nature Neuroscience,

Provided by New York University



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