

Early-life pain may lead to obesity risk, especially in females, study finds

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Inflammatory pain at birth changes how the hippocampus, a part of the brain associated with memory and eating behavior, works later in life, and this pain also causes adult rats to eat more frequently and in larger amounts, according to a study by Georgia State University and the Charlie Norwood VA Medical Center.

The study found early-life <u>inflammatory pain</u> increases sucrose intake in adult male and <u>female rats</u>, and it decreases the expression of a protein that is critical for memory, activity-regulated cytoskeleton-associated protein (Arc), in hippocampal neurons following the consumption of a sweetened solution. In addition, the effects of neonatal <u>pain</u> are more pronounced in female <u>rats</u> and can be reduced in all rats by administering morphine at the time early-life pain is experienced.

The findings demonstrate for the first time that one brief episode of inflammatory pain on the day of birth has a long-lasting, sex-dependent effect on the intake of food into adulthood. The results are published in the journal *Physiology & Behavior*.

Previous studies have found a connection between memory and overeating. In humans, disrupting the encoding of the memory of a meal, such as by watching television or playing computer games, increases the amount of food consumed at the next meal. On the contrary, recalling and enhancing the memory of a meal decreases the amount of food ingested at a future meal.



In this study, the researchers induced neonatal inflammatory pain by injecting an inflammatory agent into a rat pup's paw on the day of birth. Some of the rats received morphine at the time of the inflammatory pain. Then, rats were trained to consume a sucrose solution at a specific time and location daily, and the researchers measured the rats' sucrose intake and sucrose-associated Arc expression in the dorsal hippocampus into adulthood. Sucrose solution was used as the meal because it's pleasant and rewarding to rats, can't be hoarded and overconsumption of sweetened beverages contributes to the development of obesity.

The study found male and female rats that experienced pain on the day of birth ate more of the sucrose solution at each meal than rats that didn't experience pain. However, there were differences between the sexes. Female rats that experienced pain consumed more during each meal and also returned to the sucrose feeding tube sooner after each meal. In addition, female rats exposed to pain at birth showed fewer cells expressing the Arc protein when they consumed sucrose.

The researchers suggest that female rats are more vulnerable to pain at birth than males, and this could be related to hormone changes during the period around birth. Male rats experience a surge of testosterone during the early postnatal period, which could protect them from some of the adverse effects of neonatal pain. Female rats don't experience a similar hormone change.

Rats that received morphine after experiencing early-life pain didn't show the same eating increases as pain-inflicted rats that didn't receive morphine. This suggests changes in eating behavior resulted from the experience of pain.

"Our research may have implications for humans because newborn rats are comparable to third-trimester human infants in terms of brain development and can be used as a model for premature human infants



born into the neonatal intensive care unit (NICU)," said Dr. Marise Parent, professor of neuroscience and psychology and associate director of the Neuroscience Institute at Georgia State. "Premature infants in the NICU can undergo numerous painful and invasive procedures each day, more than half without <u>pain relievers</u>. Our study suggests that inflammatory pain experienced by infants in the NICU may contribute to the increased food consumption and obesity observed in this population, and that when possible, preventing pain in NICU infants could reduce the increased risk for obesity in this vulnerable population."

Provided by Georgia State University

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