

Brain changes behind pain sensitivity may affect older women more

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A new study has found that the brain system enabling us to inhibit our own pain changes with age, and that gender-based differences in those changes may lead females to be more sensitive to moderate pain than



males as older adults.

Researchers used fMRI scans to examine <u>brain responses</u> in men and women who had rated the intensity and unpleasantness of pain during exposure to increasing levels of heat. The results suggested that established <u>gender differences</u> in pain perception could likely be traced at least in part to this brain network, and offered new evidence that those gender differences may become more disparate with age.

"The most novel part of this study is looking at gender by age," said lead study author Michelle Failla, assistant professor in the College of Nursing at The Ohio State University. "Most of the work characterizing which regions in the brain respond to pain have been done in people aged 18 to 40. We want to understand what's happening between the ages of 30 and 90 years old because that's when people are beginning to experience chronic pain."

The study was published recently in *The Journal of Pain*.

Plenty of previous research has shown that females are more sensitive to pain than males, but the brain regions and functions behind the gender differences in pain perception have mostly remained a mystery. And in later adulthood, when risk for <u>chronic pain</u> is higher and our tolerance for pain drops, even less about the brain's role in pain perception is known.

In this study, the researchers specified that they holistically examined gender-based differences that may relate not just to biological sex, but also to <u>social factors</u> that influence how people respond to pain.

The imaging component of the study zeroed in on the descending pain modulatory system (DPMS), a hub of brain regions that communicate with each other to engage signal transmission—including activation of



opioid receptors—that enables us to reduce our own pain.

The study sample included 27 females and 32 males between ages 30 and 86 who were asked to report when applied heat reached levels of just-noticeable, weak and moderate pain and to rate how unpleasant each level felt. Researchers used the fMRI imaging to observe DPMS activity that corresponded with each participant's individual pain response.

"There are different brain regions involved in those distinctions between perception of pain intensity and unpleasantness, so we thought it was important to look at both and see how those brain regions are recruited during pain," said Failla, also an investigator in the Center for Healthy Aging, Self-Management and Complex Care in Ohio State's College of Nursing.

Results showed that a few regions within the brain's pain modulatory system did indicate a gender-by-age difference: At the moderate pain level, men showed an increased DPMS response with older age, while as women aged, the DPMS response decreased. A decreased response in the brain is presumed to translate into a lower ability to harness our own physiological functions to reduce our pain.

Presumed is a key word: While the DPMS is believed to have a significant role in pain sensitivity and tolerance, researchers are still working toward describing exactly how it works and how an intact versus dysfunctional system shows up in scans.

"We don't know exactly what is an optimum DPMS response," Failla said. "Are we seeing it activated to catch up with your pain, or is it already working, meaning the pain could have been worse?"

The researchers are continuing this work, which includes investigating brain activity in people who may have a difficult time articulating the



pain that they're feeling—such as people with dementia or autism.

The more scientists can learn about the brain's role in <u>pain perception</u>, the better the chances are for more effective pain management, Failla said.

"Pain is such an individual experience. In science we're moving toward individual factors that can influence pain specifically and what makes it different for each person," she said. "This could then identify a mechanism we can target, or even just give us a better understanding that there are different levels of innate abilities to modulate pain."

This study was conducted at Vanderbilt University Medical Center and was supported by a National Institute on Aging grant awarded to coauthors Todd Monroe of Ohio State and Ronald Cowan of the University of Tennessee Health Science Center.

Other co-authors include Paul Beach of Emory University, Sebastian Atalla of the University of North Carolina, and Mary Dietrich and Stephen Bruehl of Vanderbilt University.

More information: Michelle D. Failla et al, Gender Differences in Pain Threshold, Unpleasantness, and Descending Pain Modulatory Activation Across the Adult Life Span: A Cross Sectional Study, *The Journal of Pain* (2023). DOI: 10.1016/j.jpain.2023.10.027

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