

## Maternal anxiety affects the fetal brain

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Anxiety in gestating mothers appears to affect the course of brain development in their fetuses, changing neural connectivity in the womb, a new study by Children's National Hospital researchers suggests. The findings, published Dec. 7, 2020, in *JAMA Open Network*, could help explain longstanding links between maternal anxiety and neurodevelopmental disorders in their children and suggests an urgent need for interventions to diagnose and decrease maternal stress.

Researchers have shown that stress, <u>anxiety</u> or depression in <u>pregnant mothers</u> is associated not only with poor obstetric outcomes but also social, emotional and <u>behavioral problems</u> in their children. Although the care environment after birth complicates the search for causes, postnatal imaging showing significant differences in brain anatomy has suggested that these problems may originate during gestation. However, direct evidence for this phenomenon has been lacking, says Catherine Limperopoulos, Ph.D., director of the Developing Brain Institute at Children's National.

To help determine where these <u>neurological</u> <u>changes</u> might get their start, Dr. Limperopoulos,

along with staff scientist Josepheen De Asis-Cruz, M.D., Ph.D., and their Children's National colleagues used a technique called resting-state functional magnetic resonance imaging (rs-fMRI) to probe developing neural circuitry in fetuses at different stages of development in the late second and third trimester.

The researchers recruited 50 healthy pregnant volunteers from low-risk prenatal clinics in the Washington, D.C. area who were serving as healthy "control" volunteers in a larger study on fetal brain development in complex congenital heart disease. These study participants, spanning between 24 and 39 weeks in their pregnancies, each filled out widely used and validated questionnaires to screen for stress, anxiety and depression. Then, each underwent brain scans of their fetuses that showed connections between discrete areas that form circuits.

After analyzing rs-fMRI results for their fetuses, the researchers found that those with higher scores for either form of anxiety were more likely to carry fetuses with stronger connections between the brainstem and sensorimotor areas, areas important for arousal and sensorimotor skills, than with lower anxiety scores. At the same time, fetuses of pregnant women with higher anxiety were more likely to have weaker connections between the parieto-frontal and occipital association cortices, areas involved in executive and higher cognitive functions.

"These findings are pretty much in keeping with previous studies that show disturbances in connections reported in the years and decades after birth of children born to women with anxiety," says Dr. De Asis-Cruz. "That suggests a form of altered fetal programming, where brain networks are changed by this elevated anxiety even before babies are born."

Whether these effects during gestation themselves linger or are influenced by postnatal care is still unclear, adds Dr. Limperopoulos. Further studies



will be necessary to follow children with these fetal differences in neural connectivity to determine whether these variations in neural circuitry development can predict future problems. In addition, it's unknown whether easing maternal stress and anxiety can avoid or reverse these brain differences. Dr. Limperopoulos and her colleagues are currently studying whether interventions that reduce stress could alter the trajectory of fetal neural development.

In the meantime, she says, these findings emphasize the importance of making sure pregnant women have support for mental health issues, which helps ensure current and future health for both mothers and babies.

"Mental health problems remain taboo, especially in the peripartum period where the expectation is that this is a wonderful time in a woman's life. Many pregnant mothers aren't getting the support they need," Dr. Limperopoulos says. "Changes at the systems level will be necessary to chip away at this critical public health problem and make sure that both mothers and babies thrive in the short and long term."

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