

Potential biomarker for post-traumatic stress disorder

August 12 2014



(Medical Xpress)—Blood expression levels of genes targeted by the stress hormones called glucocorticoids could be a physical measure, or biomarker, of risk for developing Post-Traumatic Stress Disorder (PTSD), according to a study conducted in rats by researchers at the Icahn School of Medicine at Mount Sinai and published August 11 in *Proceedings of the National Academy of Sciences (PNAS)*. That also makes the steroid hormones' receptor, the glucocorticoid receptor, a potential target for new drugs.

Post-Traumatic Stress Disorder (PTSD) is triggered by a terrifying event, either witnessed or experienced. Symptoms may include flashbacks, nightmares and severe anxiety, as well as uncontrollable



thoughts about the event. Not everyone who experiences trauma develops PTSD, which is why the study aimed to identify biomarkers that could better measure each person's vulnerability to the disorder.

"Our aim was to determine which genes are differentially expressed in relation to PTSD," said lead investigator Rachel Yehuda, PhD, Professor of Psychiatry and Neuroscience and Director of the Traumatic Stress Studies Division at the Icahn School of Medicine at Mount Sinai. "We found that most of the genes and pathways that are different in PTSD-like animals compared to resilient animals are related to the glucocorticoid receptor, which suggests we might have identified a therapeutic target for treatment of PTSD," said Dr. Yehuda, who also heads the Mental Health Patient Care Center and PTSD Research Program at the James J. Peters Veterans Affairs Medical Center in the Bronx.

The research team exposed a group of male and <u>female rats</u> to litter soiled by cat urine, a predatory scent that mimics a life-threatening situation. Most PTSD studies until now have used only male rats. Mount Sinai researchers included female rats in this study since women are more vulnerable than men to developing PTSD. The rats were then categorized based on their behavior one week after exposure to the scent. The authors also examined patterns of gene expression in the blood and in stress-responsive brain regions.

After one week of being exposed to soiled cat litter for 10 minutes, vulnerable rats exhibited higher anxiety and hyperarousal, and showed altered glucocorticoid receptor signaling in all tissues compared with resilient rats. Moreover, some rats were treated with a hormone that activates the glucocorticoid receptor called corticosterone one hour after exposure to the cat urine scent. These rats showed lower levels of anxiety and arousal one week later compared with untreated, trauma-exposed rats.



"PTSD is not just a disorder that affects the brain," said co-investigator Nikolaos Daskalakis, MD, PhD, Associate Research Scientist in the Department of Psychiatry at the Icahn School of Medicine at Mount Sinai. "It involves the entire body, which is why identifying common regulators is key. The <u>glucocorticoid receptor</u> is the one common regulator that consistently stood out."

More information: "Expression profiling associates blood and brain glucocorticoid receptor signaling with trauma-related individual differences in both sexes," by Nikolaos P. Daskalakis, Hagit Cohen, Guiqing Cai, Joseph D. Buxbaum, and Rachel Yehuda., *PNAS*, www.pnas.org/cgi/doi/10.1073/pnas.1401660111

Provided by The Mount Sinai Hospital

Citation: Potential biomarker for post-traumatic stress disorder (2014, August 12) retrieved 19 April 2024 from

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