

New study suggests ways to alleviate social withdrawal symptoms in mental illnesses

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Even as people are now being forced to social distance to avoid contracting coronavirus—a situation considered stressful for many—some people avoid social situations even under everyday circumstances. Social avoidance is a hallmark of a variety of psychiatric illnesses, including depression and anxiety disorders as well as autism.



Effective treatments for these symptoms remain elusive, with most therapies working for only a fraction of people that suffer from these illnesses. University of California, Davis, researchers studied the role of oxytocin, a neurotransmitter produced in the hypothalamus, which is known to play an important role in social behavior across species. By analyzing the behavior of mice receiving different treatments, they gained insight into whether oxytocin may be a good target for future pharmacological therapies that could alleviate social withdrawal in mental illness.

Curiously, oxytocin sometimes reduces social anxiety-related behaviors, but other times increases social anxiety, researchers said. In this study, researchers focused on identifying brain pathways that mediate the anxiety-reducing effects of oxytocin. To do this, researchers tested the effects of different pharmaceutical compounds that either activated or blocked the activity of oxytocin in the nucleus accumbens, a part of the brain known to control motivation.

"Unlike SSRIs (selective serotonin reuptake inhibitors, a class of antidepressants), which take at least four weeks to show positive effects, pharmacological compounds changing oxytocin activity show rapidacting effects on behavior," said Alexia Williams, a doctoral student who is the lead author of the study. "This makes oxytocin an exciting neurotransmitter to study. Our goal was to understand more about oxytocin's potential anxiety reducing properties."

The study was published last month in the journal *Neuropsychopharmacology* and highlighted by a commentary.

Researchers found that after California mice experienced negative social interactions, they had less expression of the oxytocin receptor gene. Typically, these negative interactions also create a state of <u>social anxiety</u>, with mice avoiding other mice. In this study, researchers show that when



oxytocin activity was blocked, mice showed <u>social avoidance</u>, even though they had not experienced negative social interactions. When studying mice that had experienced negative social interactions, researchers found that increasing oxytocin activity reduced social avoidance behaviors. Although these mice had been exposed to stress, the pharmacological treatment led to normal social behaviors as if they had not been exposed to stress. Mice, Williams explained, are a social species and normally prefer to interact with other mice.

"This contributes to a clearer picture of how the neural circuits that affect social behaviors are affected by stress. Changes in <u>neural circuits</u> may be leading to some behaviors that have proved difficult to treat for many people struggling with mental disorders," she said.

Findings from basic science studies such as this one may guide the development of pharmacotherapies to effectively treat individuals suffering from <u>anxiety disorders</u>, the researchers said.

"We hope that by elucidating the role of oxytocin in anxiety-like behaviors on a molecular level, that we may begin to understand it's realistic potential as a novel therapeutic" said Williams.

More information: Alexia V. Williams et al, Social approach and social vigilance are differentially regulated by oxytocin receptors in the nucleus accumbens, *Neuropsychopharmacology* (2020). DOI: 10.1038/s41386-020-0657-4

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