Source: NIH/National Institute of Mental Health

Date: 08 December 2005

## Trust-building Hormone Short-circuits Fear In Humans

A brain chemical recently found to boost trust appears to work by reducing activity and weakening connections in fear-processing circuitry, a brain imaging study at the National Institutes of Health's (NIH) National Institute of Mental Health (NIMH) has discovered.

Functional magnetic resonance imaging data (red) superimposed on structural MRI scans. Frightful faces triggered a dramatic reduction in amygdala activity in subjects who had sniffed oxytocin, suggesting that oxytocin mediates social fear and trust via the amygdala and related circuitry. (Source: NIMH Genes, Cognition and Psychosis Program)

Scans of the hormone oxytocin's effect on human brain function reveal that it quells the brain's fear hub, the amygdala, and its brainstem relay stations in response to fearful stimuli. The work at NIMH and a collaborating site in Germany suggests new approaches to treating diseases thought to involve amygdala dysfunction and social fear, such as social phobia, autism, and possibly schizophrenia, report Andreas Meyer-Lindenberg, M.D., Ph.D., NIMH Genes Cognition and Psychosis Program, and colleagues, in the December 7, 2005 issue of the *Journal of Neuroscience*.

"Studies in animals, pioneered by now NIMH director Dr. Thomas Insel, have shown that oxytocin plays a key role in complex emotional and social behaviors, such as attachment, social recognition and aggression," noted NIH Director Elias Zerhouni, M.D.. "Now, for the first time, we can literally see these same mechanisms at work in the human brain."

"The observed changes in the amygdala are exciting as they suggest that a long-acting analogue of oxytocin could have therapeutic value in disorders characterized by social avoidance," added Insel.

Inspired by Swiss scientists who last summer

reported [1] that oxytocin increased trust in humans, Meyer-Lindenberg and colleagues quickly mounted a brain imaging study that would explore how this works at the level of brain circuitry. British researchers had earlier linked increased amygdala activity to decreased trustworthiness. [2] Having just discovered decreased amygdala activity in response to social stimuli in people with a rare genetic brain disorder that rendered them overly trusting of others, Meyer-Lindenberg hypothesized that oxytocin boosts trust by suppressing the amygdala and its fear-processing networks.

To test this idea, he asked 15 healthy men to sniff oxytocin or a placebo prior to undergoing a functional magnetic resonance imaging (fMRI) scan, which reveals what parts of the brain that are activated by particular activities. While in the scanner, the men performed tasks known to activate the amygdala -- matching angry or fearful faces and threatening scenes.

As expected, the threatening pictures triggered strong activation of the amygdala during the placebo scan, but markedly less activity following oxytocin. The difference was especially pronounced in response to threatening faces, suggesting a pivotal role for oxytocin in regulating social fear. In addition, oxytocin dampened the amygdala's communication with sites in the upper brainstem that telegraph the fear response. The results mirrored findings in rats [3], reported earlier this year by European scientists.

"Because increased amygdala activation has been associated with social fear in social phobia, genetic risk for anxiety and depression, and possibly with social fear in autism assessed during faces processing, this dual mode of action of oxytocin in humans suggests a potentially powerful treatment approach toward socially relevant fear," suggest the researchers.

People with autism characteristically avert their gaze from faces. A fMRI study [4] reported earlier this year by NIMH grantee Richard Davidson, Ph.D., University of Wisconsin, and colleagues, found over-activation of the amygdala in people with autism when they were looking at faces. Meyer-Lindenberg said future studies may test oxytocin as a treatment for such social anxiety symptoms in children with autism.

"Future research may also examine how oxytocin affects the amygdala in women, the mode of action

----

of related hormones such as vasopressin, and how genetic variants in these hormones and their receptors affect brain function," he added.

Also participating in the research were: Peter Kirsch, Christin Esslinger, Daniela Mier, Stefanie Lis, Harald Gruppe, Bernd Galihofer, Justus-Liebig University, Giessen, Germany; Qlang Chen, Sarina Siddhanti, Venkata Mattay, NIMH Genes Cognition and Psychosis Program.

- [1] Kosfeld M, Heinrichs M, Zak PJ, Fischbacher U, Fehr E. Oxytocin increases trust in humans. *Nature*. 2005 Jun 2;435(7042):673-6.
- [2] Winston JS, Strange BA, O'Doherty J, Dolan RJ. Automatic and intentional brain responses during evaluation of trustworthiness of faces. *Nat Neurosci.* 2002 Mar; 5(3):277-83.
- [3] Huber D, Veinante P, Stoop R. Vasopressin and oxytocin excite distinct neuronal populations in the central amygdala. *Science*. 2005 Apr 8;308 (5719):245-8.
- [4] Dalton KM, Nacewicz BM, Johnstone T, Schaefer HS, Gernsbacher MA, Goldsmith HH, Alexander AL, Davidson RJ. Gaze fixation and the neural circuitry of face processing in autism. *Nat Neurosci*. 2005 Apr;8(4):519-26. Epub 2005 Mar 6.

Refs
Breastfeeding
The power of love
Oxytocin and voles
Oxytocin and drugs
Oxytocin: structure
Oxytocin and women
Oxytocin and estradiol
Oxytocin and addiction
Hyper-reactive HPA rats
The evolution of emotion
Oxytocin and the randy rat
Oxytocin and social interaction
Can oxytocin, chemical addiction and the science of love

## HOME

The biology of true love

BLTC Research
The brain in love
Sensualism . com