

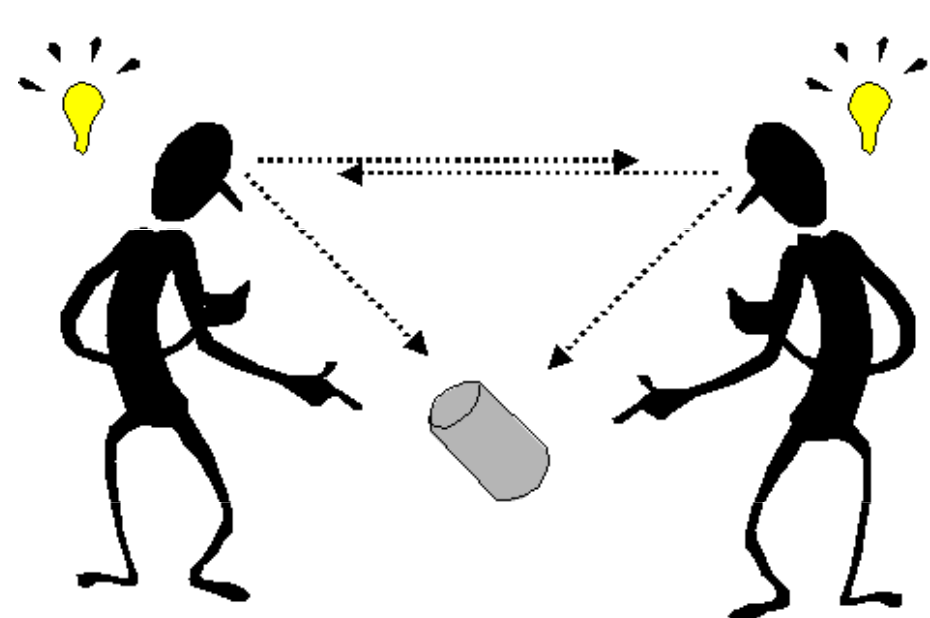
# Social interaction in cocaine users: altered response to joint attention and underlying functional changes of the reward system

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## Introduction

- **Social interaction deficits** in drug users likely have consequences for treatment and contribute to the high costs for the society associated with addiction [1]
- The **neural basis** of altered social interaction in drug users is unknown
- Joint attention is a central element of social interaction (see Figure 1)
- Engagement in joint attention is considered to reflect our understanding of another person's point of view and has been shown to activate the **reward system** [2]



**Fig 1.** Joint attention is established when a person follows the direction of another person's gaze so that both attend to the same object

## Objective

To investigate the nature of basic **social interaction deficits in cocaine users** by applying behavioral, psychophysiological and functional brain imaging methods

## Methods

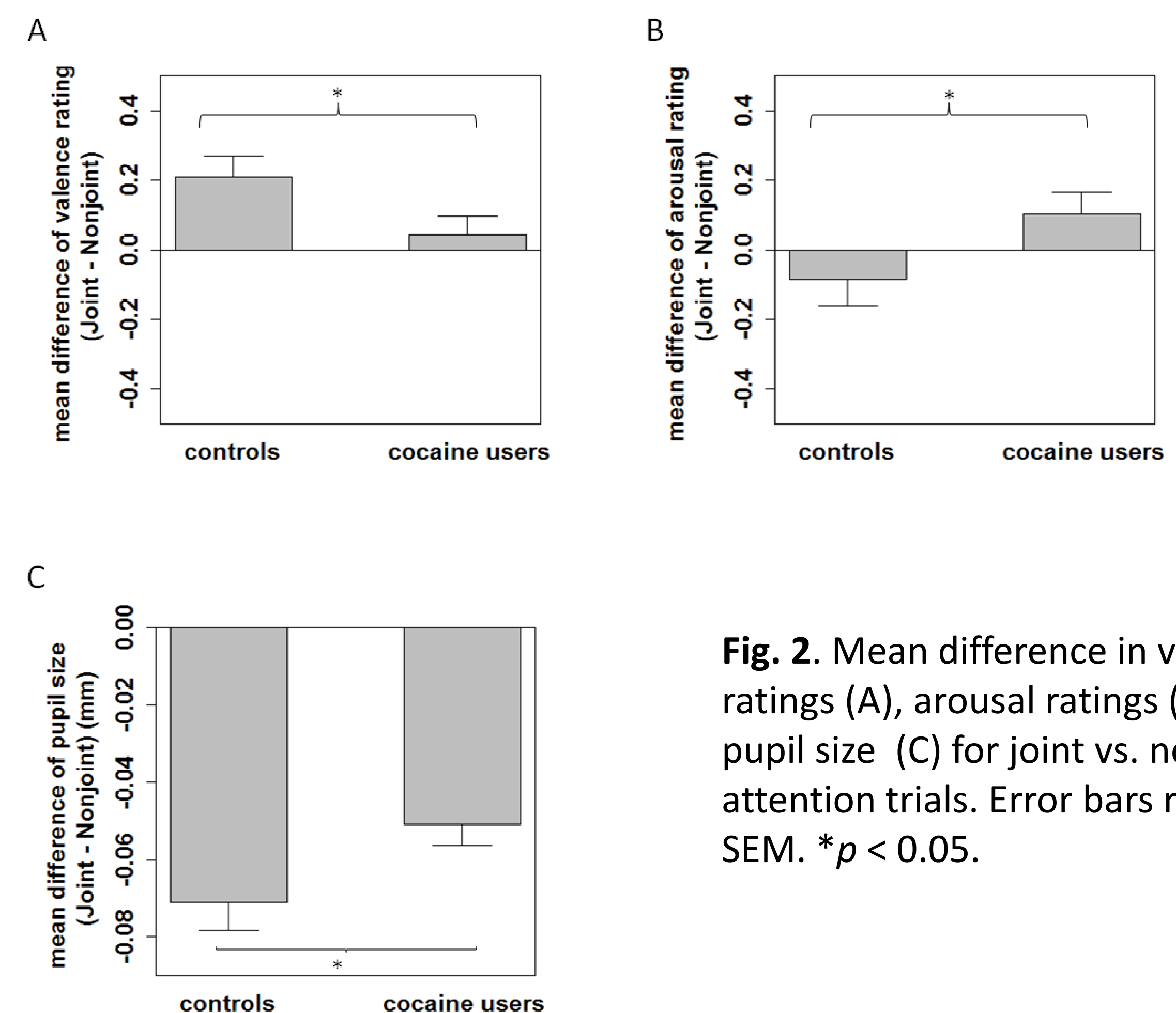
	Study I	Study II
<b>Interactive task</b>	<ul style="list-style-type: none"> <li>• Gaze was recorded by an eye-tracking device and used to control the gaze of an avatar</li> <li>• Participants either looked in the same direction as the avatar (joint attention) or in another direction (nonjoint attention)</li> </ul>	
<b>Participants</b>	<ul style="list-style-type: none"> <li>• 80 regular cocaine users</li> <li>• 63 healthy controls</li> </ul>	<ul style="list-style-type: none"> <li>• 16 regular cocaine users</li> <li>• 16 healthy controls</li> </ul>
<b>Dependent variables</b>	<ul style="list-style-type: none"> <li>• pupil size</li> <li>• valence ratings</li> <li>• arousal ratings</li> </ul>	<ul style="list-style-type: none"> <li>• Functional magnetic resonance imaging: BOLD contrast</li> <li>• network size: Social Network Questionnaire</li> </ul>
<b>Statistics</b>	mixed-effects analyses of variance: between-subjects factor: group; within-subjects factor: joint (joint vs. nonjoint attention)	GLM as implemented in SPM8

## Results Study I

- **Valence ratings** revealed that cocaine users differentiated less between joint attention and nonjoint attention than controls: (interaction group\*joint for valence ratings: (F(1,141)=4.35, p<0.04))
- **Arousal ratings** additionally reflect altered emotional engagement than controls (interaction group\*joint for arousal ratings: (F(1,141)=3.94, p<0.05))
- Subjective ratings were in line with reduced **pupil responses** in cocaine users (interaction group\*joint for pupil size: (F(1,130)=4.91, p<0.03)) (Figure 2a-c)

## References:

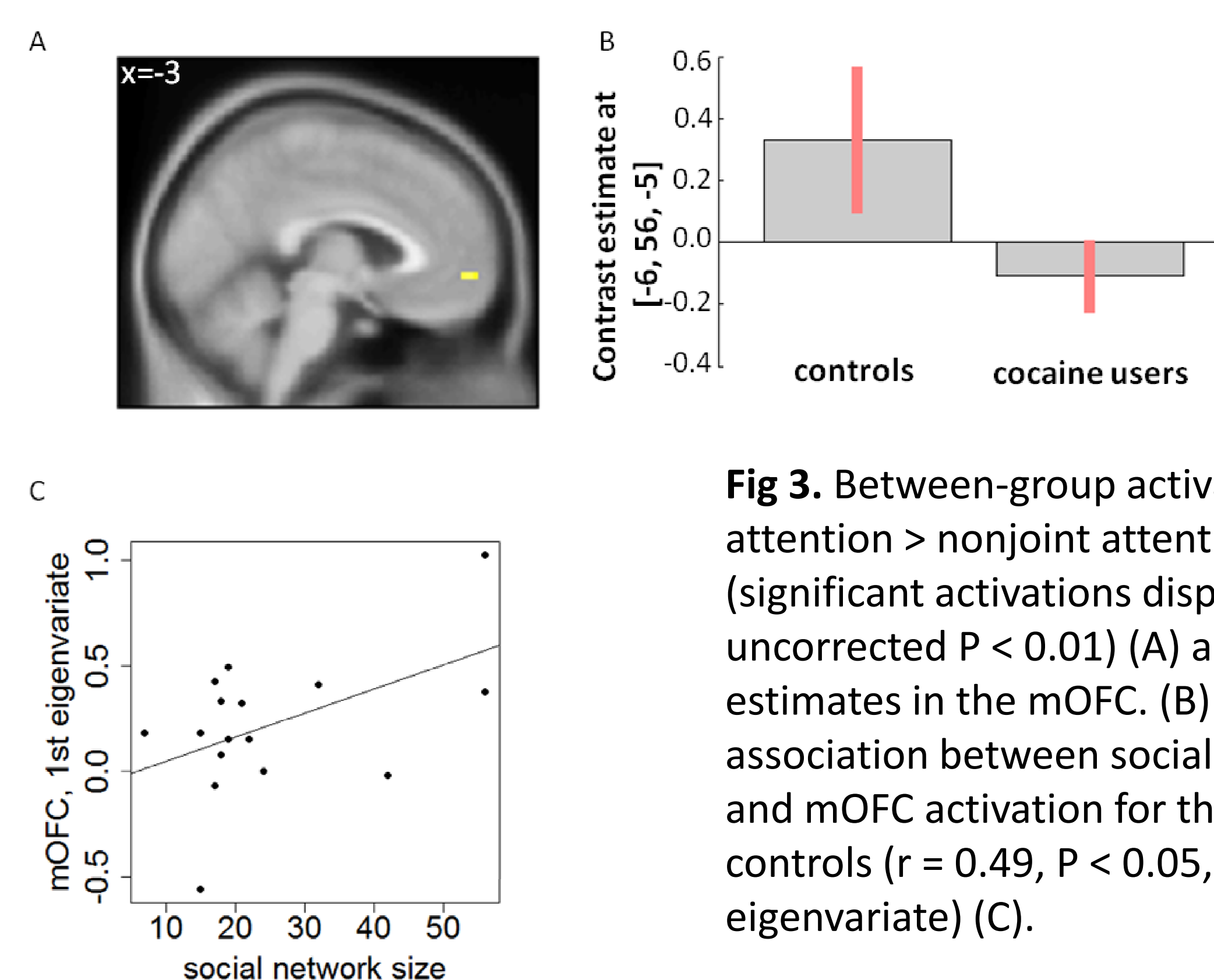
- [1] Volkow, N.D., Baler, R.D., Goldstein, R.Z., 2011. Addiction: pulling at the neural threads of social behaviors. *Neuron* 69, 599-602.
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- These data are published in: Preller K.H, et al., 2014. Functional changes of the reward system underlie blunted response to social gaze in cocaine users. *Proc Natl Acad Sci USA* 111, 2842-2847.



**Fig. 2.** Mean difference in valence ratings (A), arousal ratings (B), and pupil size (C) for joint vs. nonjoint attention trials. Error bars refer to SEM. \*p < 0.05.

## Results Study II

- In response to social interaction cocaine users displayed decreased activation of the **medial orbitofrontal cortex (mOFC)** (peak: x=-6, y=56, z=-5, p<0.05, FWE) – a key region of reward processing
- Blunted activation of the mOFC was significantly correlated with a decreased **social network size** (r=0.35, p<0.05) (Figure 3a-c)



**Fig 3.** Between-group activation for joint attention > nonjoint attention contrast (significant activations displayed at uncorrected P < 0.01) (A) and contrast estimates in the mOFC. (B). Positive association between social network size and mOFC activation for this contrast in controls (r = 0.49, P < 0.05, first eigenvariate) (C).

## Conclusion and implication

- Basic social interaction deficits in cocaine users may arise from **blunted social reward processing**
- Alterations in social gaze processing seem to be related to **impaired real-life social behavior** in cocaine users
- Altered reward sensitivity might reduce the motivation to engage in social interaction and **impair general social competence**
- This might explain why negative social consequences (e.g. family problems) do not lead the addicted person to give up drug use
- Since social reward processing is an important factor in the treatment of substance use disorders, **training of social reward processing** might be beneficial for therapy

## Disclosure

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