## Cognitive flexibility in Pathological Gambling and Alcohol Dependence: An fMRI study



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## Introduction Results \* A lack of cognitive flexibility is evident in alcohol dependence (AD): Behavioral: The Alcohol Dependent group tended to have slower reaction times than the \* Perseveration and stereotypical behaviour emerge as compulsive healthy controls (HC), p=.07 drinking behaviour. \* Neurocognitive studies indicate cognitive inflexibility in AD, which is fMRI: sometimes attributed to the neurotoxic effects of ethanol Main effects of switching behavior (switch vs. stay) in all three groups: \* Bilateral: putamen, medial, middle, and superior frontal gyrus, Cognitive inflexibility has also been found in pathological gambling (PG). parahippocampal gyrus an addictive behaviour without toxic effects of drugs on the brain with \* Right: anterior cingulate cortex (ACC) (see figure below left). broad neurobiological resemblance to substance use disorders (van Holst et al., 2010). \* Yet, brain functioning underlying diminished cognitive flexibility in pathological gambling and alcohol dependence has not been studied as of yet. Aims: \* Compare the neural correlates of cognitive flexibility in pathological gambling and alcohol dependence to healthy controls \* to better understand perseverative behaviour in addictions \* compare substance dependence to addictive behaviour without neurotoxic effects of substances Methods Group interaction alcohol dependents vs healthy controls and pathological gamblers vs healthy controls (switch vs. stay): Cognitive flexibility paradigm: Switch-task (Sohn et al., 2000) Figure above right: \* presentation of two stimuli: a letter and a digit. AD < HC: \* if color of letter is red: press left for vowel, press right for consonant right anterior cingulate \* If color is blue: press left for even, press right for odd Bilateral parahippocampal gyrus right middle frontal gyrus 5 Time Figure left: 8 PG < AD: right anterior cingulate В 8 left parahippocampal gyrus Trial types: right middle temporal gyrus L 9 same as previous trial (repetition trials: 4-6 trials) \* switch from color previous trial (switch trials) 4 G Event related paradigm. Task duration: 15 minutes. Conclusions \* The task was explained and practiced outside the scanner. fMRI Pathological gamblers and alcohol dependent persons activate error Participants: monitoring areas less than healthy controls during switch trials 20 pathological gamblers (PG), 21 alcohol dependent (AD) persons, 19 Less control over behavior during effortful cognitive proccesing healthy controls. AD show less activity in executive areas (middle frontal gyrus). This may be related to toxic effect of alcohol on the brain, and behavioral performance PG and AD groups recruited from a local outpatient treatment clinic. Exclusion: severe psychopathology, psychoactive drugs, TBI Behavioral: Alcohol dependent persons tend to react slower and need more trials to reach ANOVA Problem Alcohol Healthy gamblers dependents controls (two-tailed) a similar level as HC (n=20) (n=21) (n=19) Age, mean (SD) 42.90 37.47 (11.33) H(2)=4.1536.15 First study to investigate cognitive flexibility and it's neuronal substrates in (12.11)(8.67) p=0.125 pathological gambling and alcohol dependence: Age of onset 27,25 30,38 F(1,39)=1.03 Less activation during switching may result in more difficulty exerting flexibly addiction, mean (11, 43)(8, 13)p=0.32 over addiction related behavior: e.g. disengaging from addictive behavior. South Oaks 11.16 0.14 0.05 $\chi^2(2) = 49.36$ Gambling Screen (3.18)(0.39)(0.22)p<0.001 Alcohol Use χ²(2)=38.89 References 28.75 5.15 4.89 (2.94)(4.55) Disorder (4.50)p<0.001 Identification test Goldstein RZ, Volkow ND, Drug addiction and its underlying neurobiological basis: neuroimaging evidence for the involvement of the frontal cortex, Am J Psychiatry, 159, 1642-52, 2002. Number of trials 1394 167.0 134 4 F(2.50)=1.46during training (68,1) (61,4) (68,1) p=0.24 Sohn M.H., Ursu S., Anderson J.R., Stenger V.A., Carter C.S. (2000), 'Inaugural article: the role of RT during fMRI 1142.9 (200.1) 1187 1040 F(2,57) = 1.96prefrontal cortex and posterior parietal cortex in task switching', Proc Natl Acad Sci U S A. vol. 97, trials (288.9)(213.0)p=.15 pp.13448-13453. an Holst RJ, van den Brink W. Veltman DJ. Goudriaan AE. Why camblers fail to win: a review of Analysis cognitive and neuroimaging findings in pathological gambling. Neurosci Biobehav Rev 2010; 34(1): 87-107. ANOVA (two-tailed) followed by group comparisons for behavioral data Image: Minimizer fMRI data: Reprints

Contrast images for switch versus repetition trials were entered into a second-level (random effects) analysis.

 Main effects across groups for each contrast were analyzed with oneway ANOVA implemented in SPM5 corrected for multiple comparisons according to the Family Wise Error (FWE) method.

- Group interactions are reported at P<0.001, masked with the appropriate main effect at p<0.05.

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