Accepted Manuscript

Effects of gender on ketamine-induced conditioned placed preference and urine metabonomics

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PII: S0273-2300(16)30052-6

DOI: 10.1016/j.yrtph.2016.03.007

Reference: YRTPH 3536

To appear in: Regulatory Toxicology and Pharmacology

Received Date: 9 November 2015

Revised Date: 10 March 2016

Accepted Date: 12 March 2016

Please cite this article as: Guo, R., Qunxing, T., Yi, Y., Xiang, L., Fan, C., Xinhua, D., Youyi, Y., Liao, L., Effects of gender on ketamine-induced conditioned placed preference and urine metabonomics, *Regulatory Toxicology and Pharmacology* (2016), doi: 10.1016/j.yrtph.2016.03.007.

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ACCEPTED MANUSCRIPT

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11	Abstract: Objective: The aim of this study was to examine whether or not there was a
12	gender difference in CPP (conditioned placed preference) induced by ketamine and to
13	further explore the effect of sex on metabolic responses to ketamine inducing in SD
14	rats. Methods: We measured ketamine-induced conditioned place preference and
15	ketamine-induced metabolic changes in urine by using ¹ H nuclear magnetic resonance
16	(NMR) coupled with principal component analysis (PCA), partial least squares (PLS)
17	and orthogonal signal correction (OSC) analysis. Results: In the CPP experiment,
18	ketamine served as a positive reinforcing agent in both male and female rats, but, in
19	particularly, the preference score of female rats was significantly higher than that of
20	male rats. Compared with male rats, the metabolic trajectory fluctuation of the female
21	rats was relatively larger. At the same time, different metabolites (1, 3-dimethyluric
22	acid, cysteine-S-sulfate, glyceraldehydes, glycine, ribitol, acetoacetic acid, creatine,

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