

Morphine analgesia and methamphetamine-induced behavioral changes in prenatally buprenorphine-exposed offspring at their adulthood

Yao-Chang Chiang^{1,2}, Tsai-Wei Hung⁴, Ing-Kang Ho^{1,3,4}

¹Center for Drug Abuse and Addiction, China Medical University Hospital, Taichung, Taiwan

²China Medical University, Taichung, Taiwan

³Graduate Institute of Clinical Medical Science, China Medical University, Taichung, Taiwan

⁴Neuropsychiatric Research Center, National Health Research Institutes, Zhunan, Miaoli County, Taiwan

Abstract

Heroin use among young women of reproductive age is an important issue. Buprenorphine is a newer maintenance agent and have been used in pregnant of female addicts. However, there is lack of information on the long-term effects of prenatal exposure to buprenorphine on their offspring. A prenatally buprenorphine-exposed animal model was established to investigate prenatal effects of buprenorphine on the offspring. Results of morphine-induced analgesia and tolerance showed that tolerance development to morphine was quicker in the prenatally buprenorphine-exposed animals. The locomotor activities and conditioned place preference induced by methamphetamine were also significantly increased in prenatally buprenorphine-exposed animals as compared with their control mates. The mRNA expression of dopamine D₁R in the nucleus accumbens was lower in the prenatally buprenorphine-exposed offspring, but no significant changes in μ -, κ -opioid, NOP, D₂R, and D₃R receptors were noted. Furthermore, the basal level of cAMP and the D₁R agonist enhanced cAMP production were significantly altered in

the prenatally buprenorphine-exposed group. Overall, these studies reveal that prenatal exposure to buprenorphine caused long-term effects on the offspring and affected the opioidergic and dopaminergic system-related mechanism. (The work was supported by National Health Research Institutes (NHRI-102A1-PDCO-1312141 and NHRI-EX102-10224NC) and China Medical University Hospital (DMR-101-117) in Taiwan.

Keywords: prenatal exposure, opioids, tolerance, behavioral sensitization

Correspondence to: Dr. Ing-Kang Ho, Center for Drug Abuse and Addiction, China Medical University Hospital, 2 Yude Road, North District, Taichung 404, Taiwan (ROC)

Tel.: +886-4-2205-2121 ext. 7528; fax: +886-4-2205-2121 ext. 7527

E-mail: iho@mail.cmuh.org.tw