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Authors: Robert N. Hughes, Jennifer J. Hamilton

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ACCEPTED MANUSCRIPT

Sex-dependent modification by chronic caffeine of acute

methamphetamine effects on anxiety-related behavior in rats

Robert N. Hughes*, Jennifer J. Hamilton

Department of Psychology, University of Canterbury, Christchurch 8140, New Zealand

*Corresponding author at: Department of Psychology, University of

Canterbury, Private Bag 4800, Christchurch 8140, New Zealand. Tel.: +64 3 369 4382;

fax: +64 3 364 2189.

E-mail address: <u>rob.hughes@canterbury.ac.nz</u> (R.N. Hughes).

(Highlights)

- Acute methamphetamine (MA) is anxiolytic for males irrespective of caffeine intake.
- MA may only be anxiolytic for females when accompanied by 31 mg/kg/day caffeine.
- Possible MA effects on female memory with caffeine may be due to modified anxiety.
- 31 mg/kg/day caffeine on its own does not affect either sex.

(Abstract)

For fourteen days, male and female PVG/c hooded rats were provided continuously with either pure drinking water, or water containing caffeine in a quantity approximating a daily dose of 31.1 mg/kg. Then at intervals of 3 days, they were administered 1, 2 mg/kg methamphetamine (MA) or saline before being tested for anxiety-related behavior in a zero maze or a light/dark box, or their short-term spatial memory was assessed in a Y maze following introduction of a novel brightness change in one of the arms. Each rat experienced each type of apparatus with the same acute MA or saline treatment while still exposed to chronic caffeine or

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