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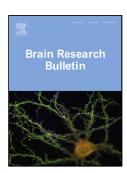
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## ACCEPTED MANUSCRIPT

#### Alterations in nociception and morphine antinociception in mice fed a high-fat diet

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#### Highlights:

- Females are less responsive to the antinociceptive effects of morphine than male littermates.
- HED increases tolerance to the antinociceptive effects of morphine on acute thermal pain.
- HED exposure reduces morphine antinociception in an inflammatory pain model.
- Male mice fed HED are resistant to the hypothermic effects of acute morphine.
- Dietary modifications may be relevant for pain therapy.

#### Abstract

Currently, more than 78.6 million adults in the United States are obese. A majority of the patient population receiving treatment for pain symptoms is derived from this subpopulation. Environmental factors, including the increased availability of food high in fat and sugar, contribute to the continued rise in the rates of obesity. The focus of this study was to investigate whether long-term exposure to a high-fat, energy-dense diet enhances baseline thermal and inflammatory nociception while reducing sensitivity to morphine-induced antinociception. Antinociceptive and hypothermic responses to morphine were determined in male and female

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