



Analgesic tolerance of morphine is prevented by modulation of D₄R at the spinal cord level

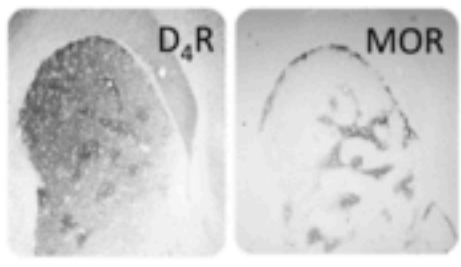
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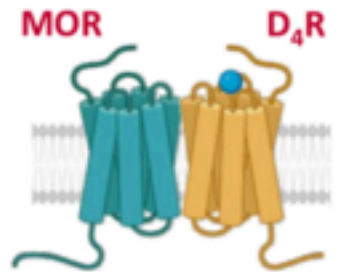
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BACKGROUND



co-localization of D₄R and MOR in the **striosomes*** of the CPU

*involved in drug-habit formation and consolidation



antagonistic allosteric D₄R-MOR interaction

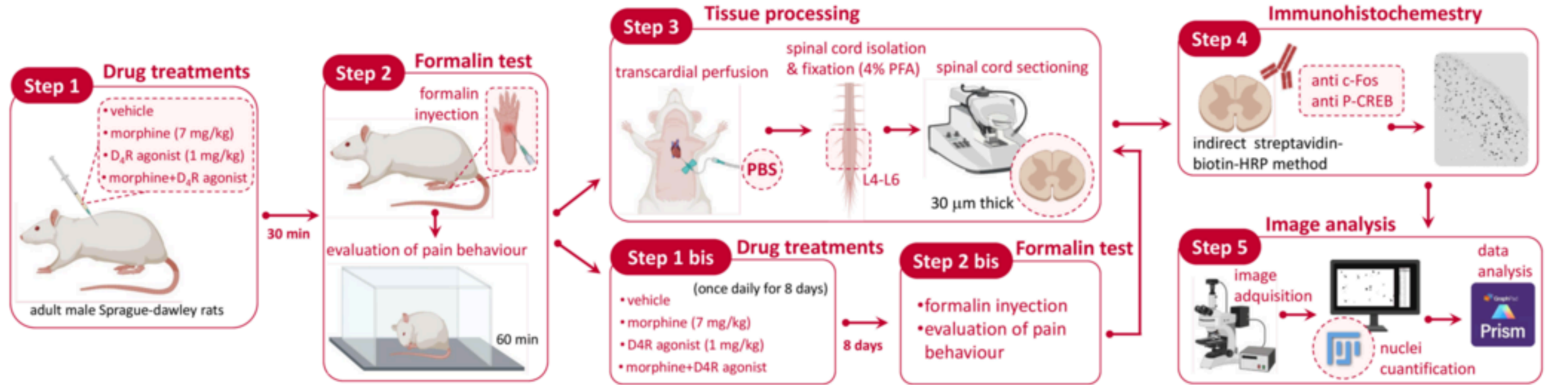
- Dopaminergic signaling
- Cellular & molecular neuroadaptations
- MOR sensitization

D₄R activation prevents morphine addiction without altering morphine-induced analgesia

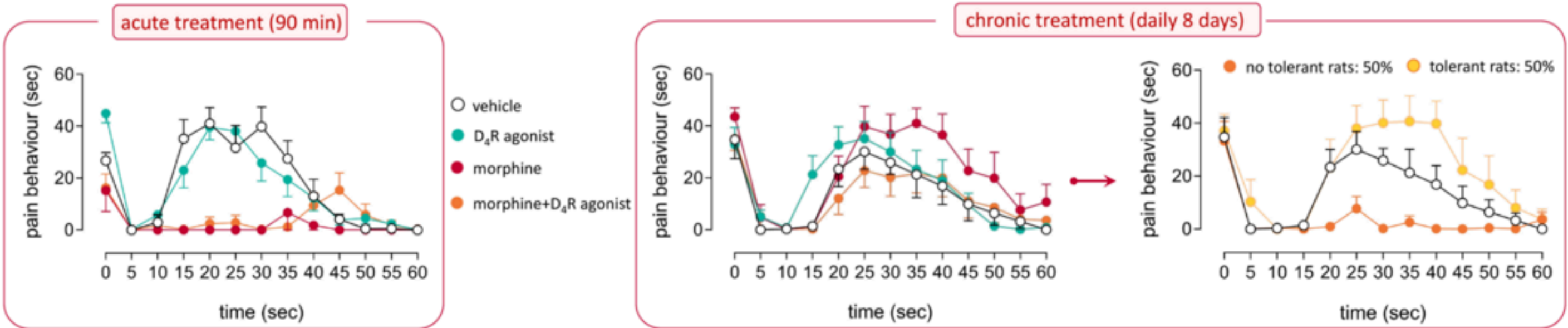
OBJECTIVE

To explore the impact of a chronic treatment of morphine, alone or in combination with a specific D₄R agonist, in the development of analgesic tolerance and alterations in nociceptive circuits within the spinal cord

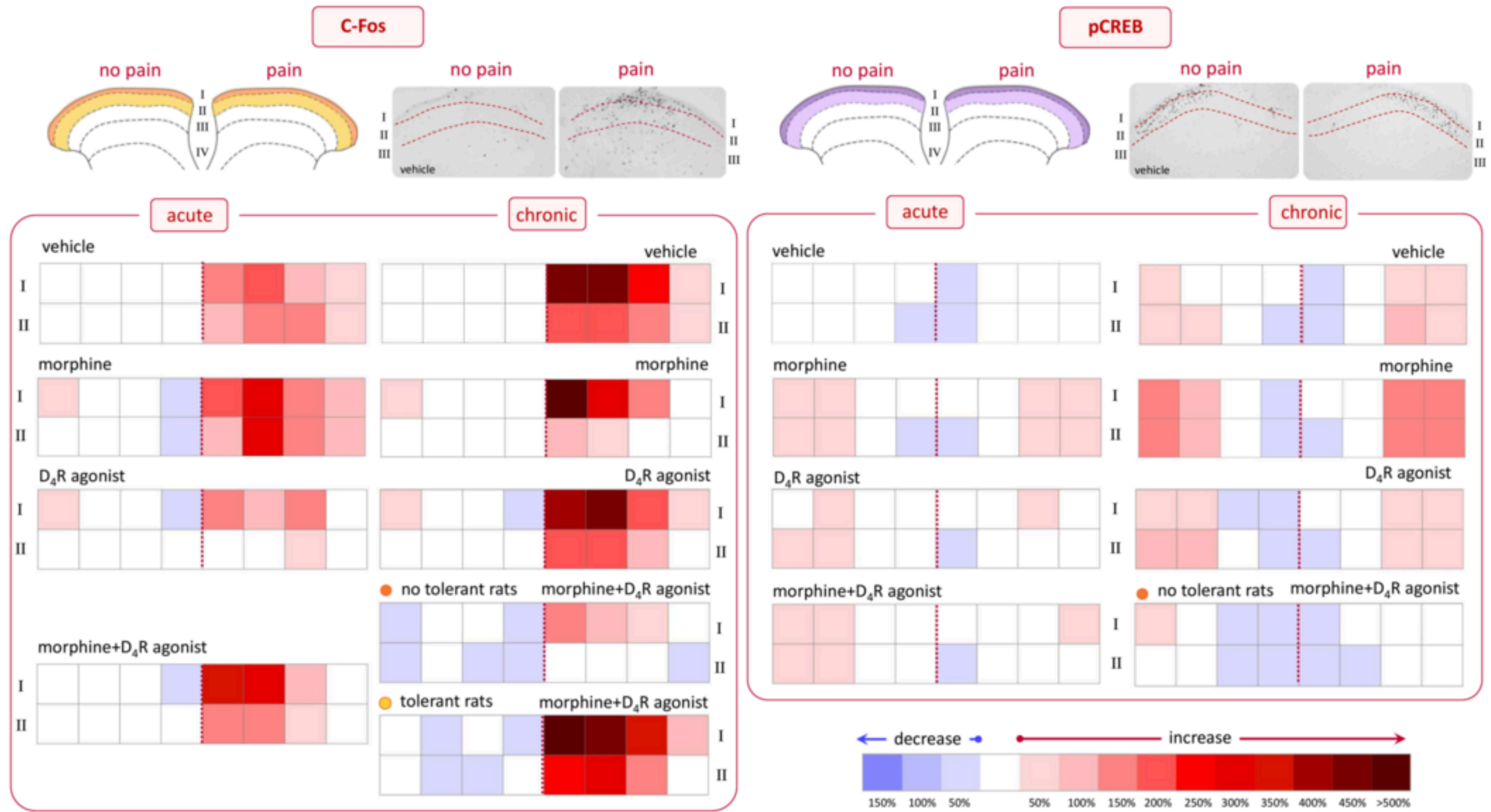
METHODS



BEHAVIORAL RESULTS



IMMUNOHISTOCHEMICAL RESULTS



CONCLUSIONS

- 1 D₄R activation prevents morphine-induced analgesic tolerance
- 2 The counteractive action of D₄R correlates with a specific signature of transcription factor expression in nociceptive pathways in laminae I-II in the dorsal horn

FUNDING

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