## Accepted Manuscript

Opioid-Induced Constipation Relief of Fixed-Ratio Combination Prolonged-Release Oxycodone/Naloxone Compared with Oxycodone and Morphine for Chronic Nonmalignant Pain: A Systematic Review and Meta-analysis of Randomized Controlled Trials

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PII: S0885-3924(17)30288-9

DOI: 10.1016/j.jpainsymman.2017.07.025

Reference: JPS 9450

To appear in: Journal of Pain and Symptom Management

Received Date: 3 February 2017

Revised Date: 16 April 2017

Accepted Date: 6 July 2017

Please cite this article as: Huang L, Zhou J-G, Zhang Y, Wang F, Wang Y, Liu D-H, Li X-J, Lv S-P, Jin S-H, Bai Y-J, Ma H, Opioid-Induced Constipation Relief of Fixed-Ratio Combination Prolonged-Release Oxycodone/Naloxone Compared with Oxycodone and Morphine for Chronic Non-malignant Pain: A Systematic Review and Meta-analysis of Randomized Controlled Trials, *Journal of Pain and Symptom Management* (2017), doi: 10.1016/j.jpainsymman.2017.07.025.

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#### ACCEPTED MANUSCRIPT

### Opioid-Induced Constipation Relief of Fixed-Ratio Combination Prolonged-Release Oxycodone/Naloxone Compared with Oxycodone and Morphine for Chronic Non-malignant Pain: A Systematic Review and Meta-analysis of Randomized Controlled Trials

Running title: Opioid-Induced Constipation Relief of OXN PR for Chronic Non-malignant Pain.

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#### Abstract

**Background.** Opioid-induced constipation (OIC) is one of the most frequent and severe adverse events (AEs) following treatment with opioids. Recent studies have indicated that fixed-ratio combination prolonged-release oxycodone/naloxone (OXN PR) could decrease OIC with similar pain relief compared with other opioids.

**Objectives.** We systematically reviewed (PROSPERO registration numbers: CRD42016036244) the constipation relief of OXN PR compared with other opioids regardless of formulation, prolonged-release or extended-release used for the relief of chronic pain.

**Methods.** Relevant studies were identified by searching PubMed, EMBASE, Web of Science, and the Cochrane library from inception to May 2016, with an update to December 2016. We

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