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SCIENTIFIC ARTICLE

# Comparison of droperidol and ondansetron prophylactic effect on subarachnoid morphine-induced pruritus<sup>A</sup>



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KEYWORDS	Abstract
Droperidol; Morphine; Ondansetron; Pruritus;	Background and objectives: The prophylactic effect of ondansetron on subarachnoid morphine- induced pruritus is controversial, while evidence suggests that droperidol prevents pruritus. The aim of this study is to compare the effects of droperidol and ondansetron on subarachnoid morphine-induced pruritus.
Subarachnoid injection	<ul> <li>Methods: 180 ASA   or    patients scheduled to undergo cesarean sections under subarachnoid anesthesia combined with morphine 0.2 mg were randomized to receive, after the child's birth, metoclopramide 10 mg (Group    - control), droperidol 2.5 mg (Group   ) or ondansetron 8 mg (Group   ). Postoperatively, the patients were assessed for pruritus (absent, mild, moderate or severe) or other side effects by blinded investigators. Patients were also blinded to their group allocation. The tendency to present more severe forms of pruritus was compared between groups. NNT was also determined.</li> <li>Results: Patients assigned to receive droperidol [Proportional odds ratio: 0.45 (95% confidence interval 0.23–0.88)] reported less pruritus than those who received metoclopramide. Ondansetron effect was similar to metoclopramide [Proportional odds ratio: 0.95 (95% confidence interval 0.49–1.83)]. The NNT for droperidol and ondansetron was 4.0 and 14.7, respectively.</li> <li>Conclusions: Ondansetron does not inhibit subarachnoid morphine-induced pruritus.</li> <li>© 2014 Sociedade Brasileira de Anestesiologia. Published by Elsevier Editora Ltda. All rights reserved.</li> </ul>

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PALAVRAS-CHAVE Droperidol; Morfina; Ondansetron; Prurido; Injeção subaracnoidea Comparação dos efeitos profiláticos do droperidol e do ondansetron sobre o prurido provocado pela morfina subaracnoidea

#### Resumo

*Justificativa e objetivos:* O efeito profilático do ondansetron sobre prurido provocado pela morfina subaracnoidea é controverso, enquanto evidências sugerem que o droperidol previne o prurido. O objetivo do presente trabalho é comparar o efeito do droperidol com o do ondansetron sobre o prurido provocado pela morfina subaracnoidea.

Métodos: 180 pacientes ASA I OU III programadas para serem submetidas a cesarianas sob anestesia subaracnoidea à qual foram acrescentados 0,2 mg de morfina foram divididas aleatoriamente para receber, logo após o nascimento da criança, 10 mg de metoclopramida (grupo I – controle), 2,5 mg de droperidol (grupo II), ou 8 mg de ondansetron (grupo III). No período pós-operatório as pacientes foram avaliadas quanto ao prurido (ausente, leve, moderado ou intenso) ou outros efeitos colaterais por observadores que não sabiam a alocação das pacientes. As pacientes também não sabiam da sua alocação. Os grupos foram comparados pela sua tendência a apresentar formas mais severas de prurido. Também determinamos o NNT.

*Resultados*: As pacientes alocadas para receber droperidol [*Odds Ratio* Proporcional: 0,45 (Intervalo de Confiança de 95% 0,23 – 0,88)] relataram menos prurido do que as que receberam metoclopramida. O efeito do ondansetron foi semelhante ao da metoclopramida [*Odds Ratio* Proporcional: 0,95 (Intervalo de Confiança de 95% 0,49 – 1,83)]. O NNT do droperidol foi 4,0 e o do ondansetron foi 14,7.

Conclusões: O ondansetron não inibiu o prurido provocado pela morfina subaracnoidea.

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### Introduction

In a previous work,<sup>1</sup> we compare the prophylactic effect of droperidol, alizapride, propofol, and promethazine on subarachnoid morphine-induced pruritus. Droperidol was the most effective agent; propofol and alizapride were less efficient; and promethazine, as other antihistamines,<sup>2</sup> was ineffective. Kjelberg and Tramér,<sup>3</sup> in a review study of pharmacological treatment of morphine-induced pruritus, concluded that droperidol was more effective than any other drug, other than morphine antagonists. But their review only included one study in which ondansetron was used to antagonize the alfentanil-induced pruritus in patients undergoing general surgery.

Evidences of ondansetron effectiveness are contradictory. Some studies have reported ondansetron effectiveness for treating<sup>4</sup> or preventing pruritus.<sup>5,6</sup> It has also been suggested that ondansetron reduces pruritus severity without reducing its incidence.<sup>7</sup> On the other hand, other studies have reported the ineffectiveness of ondansetron or its lower efficacy compared to other drugs.<sup>8-10</sup>

Given this contradiction and lack of comparison between droperidol and ondansetron, we decided to compare the prophylactic effect of the two drugs in patients undergoing cesarean section (C-section).

#### Methods

This study was approved by the Research Ethics Committee of the Universidade Católica de Pelotas (Ref: 2011/18), and written informed consent was obtained from all patients. This randomized double-blind trial enrolled 180 patients ASA I or II scheduled for C-section, regardless of the cause of obstetric indication. In addition to the refusal to participate in this research, patients were excluded in the following cases: inadequate anesthesia, any itchy skin disease, recent use of opioids or any other drug that causes respiratory depression, hyperemesis, or inability to answer questions clearly.

Upon arrival at the operating room, patients received an infusion of Ringer's lactate and 50 mcg of fentanyl were intravenously (IV) administered. The total volume of fluid infused during surgery was recorded in three moments: at lumbar puncture; at the child's birth, and at the end of surgery. Standard monitoring (non-invasive blood pressure, SpO2, and ECG) was established.

Subarachnoid anesthesia was induced via the lateral approach<sup>11</sup> with Quincke needle at L2-L3 or L3-L4, using 2 mL of 5% lidocaine hyperbaric solution (100 mg) or 4 mL of 5% bupivacaine hyperbaric solution (20 mg). Two hundred micrograms of morphine was added to the injected anesthetic. The manual displacement of the uterus to the left was established prophylactically and, in case of hypotension (systolic blood pressure 70% of baseline values or below 90 mmHg), improvement in displacement was attempted and/or fractionated doses of metaraminol (0.5 mg each) were given. As the leading cause of hypotension before birth is cava compression, we made a distinction between its incidence before birth (initial hypotension), treated with metaraminol only if persisting after improving the manual displacement, and hypotension after child birth (final hypotension), which has the same pathophysiology of hypotension from any spinal anesthesia and was treated

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