

The Effect of Ondansetron and Dexamethasone on Nausea and Vomiting under Spinal Anesthesia

Navid Kalani^{1,3}, Hasan Zabetian^{2,3*}, Mohammad Sadegh Sanie^{2,3}, Mansour Deylami⁴,
Mohammad Radmehr³, Reza Sahraei³, Hossein Kargar Jahromi², Wesam Kooti⁵

1. Medical Ethic Research Center, Jahrom University of Medical Sciences, Jahrom, Iran;
2. Research Center For Non-Communicable Diseases, Jahrom University of Medical Sciences, Jahrom, Iran;
3. Department of Anesthesiology, Jahrom University of Medical Sciences, Jahrom, Iran;
4. Department of Anesthesiology, Golestan University of Medical Sciences, Golestan, Iran;
5. Student Research Committee, Kurdistan University of Medical Sciences, Sanandaj, Iran

*Corresponding Author:

Hasan Zabetian, MD,
Department of Anesthesiology,
Jahrom University of Medical Sciences,
Jahrom, Iran.

Tel: 98-917-314-4589

E-mail: h_zabetian@yahoo.com

Received: February 27, 2016

Revised: September 2, 2016

Accepted: September 20, 2016

ABSTRACT

BACKGROUND

During abdominal surgery under regional anesthesia, nausea may happen due to several contributing factors. This study compared the effects of ondansetron and dexamethasone on nausea and vomiting under spinal anesthesia.

METHODS

One hundred and twenty patients of 15 to 35 years old with ASA class I and II were enrolled. Before administering either ondansetron or dexamethasone, blood pressure and heart rate of the patients were recorded. The patients received 70 mg of 5% lidocaine for spinal anesthesia. Patients who received 6 mg of ondansetron were considered as group A, while group B received 8 mg of dexamethasone. The level of nausea and vomiting, blood pressure, heart rate and respiratory rate of each patient was measured at 1, 5, 10, 15 and 30 minutes after spinal anesthesia and during recovery (every 5 minutes).

RESULTS

There was a significant difference between nausea and vomiting between the two groups after spinal anesthesia within the first and fifth minutes. There was no significant difference between nausea and vomiting between the two groups within 10, 15 and 30 minutes and during recovery at 5, 10, 15 and 30 minutes.

CONCLUSION

Dexamethasone and ondansetron were shown to equally reduce the incidence of nausea and vomiting under spinal anesthesia and can be recommended as a good choice for prevention of nausea and vomiting during surgeries.

KEYWORDS

Ondansetron; Dexamethasone; Nausea; Vomiting; Surgery; Spinal anesthesia

Please cite this paper as:

Kalani N, Zabetian H, Sanie MS, Deylami M, Radmehr M, Sahraei R, Kargar Jahromi H, Kooti W. The Effect of Ondansetron and Dexamethasone on Nausea and Vomiting under Spinal Anesthesia. A Pilot Study. *World J Plast Surg.* 2017;6(1):88-93.

INTRODUCTION

Worldwide, one of the most common obstetrics surgeries is

caesarean section,¹ while spinal anesthesia is used for caesarean sections as a safe, easy and quick technique.² Nausea and vomiting after caesarean sections have been reported in more than 66% of cases,^{3,4} due to sudden contractions in diaphragm and manipulation and stretching of the abdominal viscera.⁵ Spinal anesthesia in a cesarean delivery can prevent the risk of pulmonary aspiration that may occur under general anesthesia.⁵

Stimulation of pharyngeal reflex can be noticed in abdominal surgeries, physical rupture and manipulation of abdominal viscera due to the release of humoral HT-5 substances, which trigger the HT3-5 receptors on vagal afferent neurons.³ During abdominal surgery under regional anesthesia, nausea may happen due to several contributing factors such as sympathetic blocks followed by parasympathetic dominance which is the most important cause of nausea after spinal anesthesia, hypotension, decreased perfusion of central nervous system, psychological changes (anxiety), and sudden abdominal movements during surgery and prescription of drugs.⁶

The risk of nausea and vomiting in final stages of pregnancy is higher due to hormonal changes and increased intra-abdominal pressure.⁶ The major complications of spinal anesthesia were shown to be nausea and vomiting, leading to an unpleasant experience in these patients.⁷ In recent decade, spinal anesthesia has been applied as a safe and fast method for cesarean delivery.^{2,7} In many cases in Iran, caesarean section is carried out as emergency without prior preparation, where the patients are usually not fasting during surgery. Hence, spinal anesthesia in such cases is regarded as a standard method to reduce the risk of vomiting and aspiration.^{3,4,8}

One of the most common problems with spinal anesthesia is the incidence of vomiting as the uterus is pushed back into the abdominal area, accounting for 66% of cases^{3,4,8} which is dealt by a variety of medications, among which metoclopramide is the most common.⁹⁻¹¹ However, there are extrapyramidal complications to be associated with the drug, leading to much concern and caution when prescribing metoclopramide.⁹⁻¹¹

Dexamethasone has been introduced as an inexpensive and widely available drug to control nausea and vomiting.^{12,13} Similarly, ondansetron is considered as an effective drug for prevention

and treatment of nausea and vomiting that is well tolerated by the patients.⁶ This drug is applied in surgeries which may be accompanied by nausea and vomiting. The serotonin receptor antagonists (HT3-5) and dexamethasone were reported as ideal medications for controlling nausea and vomiting, without any adverse side effects.¹⁴⁻¹⁷

The exact mechanism of dexamethasone in preventing nausea and vomiting is still unknown, but it may be due to inhibition of prostaglandin synthesis.^{14,15} Moreover, ondansetron can inhibit serotonin receptor leading to the prevention of nausea and vomiting.¹⁵⁻¹⁷ Given the high prevalence of cesarean section in Iran, the harmful effects of nausea and vomiting during and after operation and insufficient studies on the effect of dexamethasone and ondansetron for prevention of nausea and vomiting during spinal anesthesia, this study intended to compare the effect of dexamethasone and ondansetron in controlling of nausea and vomiting during spinal anesthesia.

MATERIALS AND METHODS

In a double-blind, randomized, controlled trial study, 120 patients of 15 to 35 years old with ASA class I and II (American Society of Anesthesia Score) were enrolled. The samples were selected through a simple random sampling method. The study population comprised patients undergoing surgery through spinal anesthesia and were admitted in Motahari Hospital in Jahrom, Iran.

The patients were randomly divided into two equal groups of 60 subjects. Before entering the operation room, the patients did not receive any premedication drugs. Before inclusion in the study, the patients were informed about the purpose of the study and potential complications and an informed written consent was provided from each participant. The study was approved in institution ethics committee.

Inclusion criteria were being pregnant and age of 15 to 35 years old, diagnosis with ASA class I and II, to be candidate to undergo caesarean section, having no history of known physical and mental illnesses, and any history of taking pain killers and anti-depressants, sleeping pills and psychotropics. The exclusion criteria were weighing more than 100 kg, age over 35 years or less than 15 years, history of

drug or alcohol dependence, treatment with antidepressants, sleeping pills and psychotropic, lack of appropriate communication with patients for evaluating postoperative nausea and vomiting, need for hospitalization in the ICU after surgery, any previous history of allergy to ondansetron or dexamethasone, need for additional treatment and sickness during surgery, rising level of anesthesia and reduction or loss of respiration, hemodynamic disorder and patient dissatisfaction.

Both groups of patients were hydrated with 7 ml/kg of Ringer's solution. Before administering either substances (ondansetron or dexamethasone), blood pressure and heart rate of the patients were recorded. The patients received 70 mg of 5% lidocaine for spinal anesthesia. Group A received 6 mg of ondansetron, while group B received 8 mg of dexamethasone (prior to clamping the cord) and in group B, 8 mg dexamethasone was administered at the same time (before clamping the umbilical cord).

The levels of nausea and vomiting, blood pressure, heart rate and respiratory rate of each patient were measured at 1, 5, 10, 15 and 30 minutes after spinal anesthesia and during recovery (every 5 minutes based on the number of vomiting, retching or nausea told by the patient). Ephedrine was applied if hypotension occurred during the operation. At the end of surgery, the patient was transferred to the recovery room. The minimum recovery time was one hour. Neither the patient nor the nurse responsible for postoperative follow-up were aware of the primary drugs prescribed to patients. Data were analyzed using SPSS

software (version 11, Chicago, IL, USA). Fisher and Chi-square tests were performed for comparison of groups. A p value less than 0.05 was considered statistically significant.

RESULTS

There was a significant difference between nausea and vomiting between the two groups of ondansetron and dexamethasone after spinal anesthesia within the first and fifth minutes ($p < 0.05$, Table 1). There was no statistically significant difference between nausea and vomiting between the two groups within 10, 15 and 30 minutes ($p > 0.05$, Table 1).

During 1, 5, 10, 15 and 30 after spinal anesthesia, 6 (10.2%), 8 (13.6%), 2 (3.4%), 9 (15.3%) and 3 patients (5.1%) in the dexamethasone group experienced nausea and vomiting, respectively, while these figures ondansetron group experiencing nausea and vomiting were 6 (10%), 4 (6.7%) and 10 patients (16.7%), respectively. There was no significant difference between nausea and vomiting between the two groups of ondansetron and dexamethasone during recovery at 5, 10, 15 and 30 minutes ($p \text{ value} > 0.05$, Table 2).

In the recovery room at 5, 10, 15 and 30 minutes, 3 (5%), 2 (3.4%), and 1 patient (1.7%) in the dexamethasone group experienced nausea and vomiting, respectively, while the figures for ondansetron group experiencing nausea and vomiting in the recovery room were 4 (6.7%), 4 (6.7%) and 3 patients (6.7%), respectively. From 30 minutes onwards in the recovery, i.e. at 35, 40, 45, 50, 55 and 60 minutes, none of

Table 1: The frequency of nausea and vomiting in two ondansetron and dexamethasone groups at 1, 5, 10, 15 and 30 minutes after spinal anesthesia

Minute	Group	Nausea and vomiting		p value
		Yes	No	
First	Ondansetron	0	60(100%)	0.027*
	Dexamethasone	6 (10.2%)	54 (89.8%)	
5	Ondansetron	0	60 (100%)	0.006*
	Dexamethasone	8 (13.6%)	52 (86.4%)	
10	Ondansetron	6 (10.2%)	54 (89.8%)	0.272*
	Dexamethasone	2 (3.4%)	58 (96.6%)	
15	Ondansetron	4 (6.7%)	56 (93.3%)	0.142**
	Dexamethasone	9 (15.3%)	51 (84.7%)	
30	Ondansetron	10 (16.7%)	50 (83.3%)	0.040**
	Dexamethasone	3 (5.1%)	57 (94.9%)	

*Fisher's Exact Test, **Chi-Square Test

Table 2: The frequency of nausea and vomiting in two ondansetron and dexamethasone groups at 5, 10, 15 and 30 minutes on recovery

Minute	Group	Nausea and vomiting		p value
		Yes	No	
5	Ondansetron	4 (6.7)	56 (93.3%)	0.658**
	Dexamethasone	5 (13.6%)	95 (86.4%)	
10	Ondansetron	4 (10.2%)	56 (89.8%)	0.119*
	Dexamethasone	0	60 (100%)	
15	Ondansetron	3 (5.1%)	57 (94.9%)	0.619*
	Dexamethasone	1 (1.7%)	59 (98.3%)	
30	Ondansetron	4 (10.2%)	56 (89.8%)	0.119*
	Dexamethasone	0	60 (100%)	

*Fisher's Exact Test, **Chi-Square Test

the patients in ondansetron and dexamethasone groups experienced nausea and vomiting.

There was no statistically significant difference between the mean levels of heart rate, systolic and diastolic blood pressures, respiration and oxygen saturation among both groups of ondansetron and dexamethasone during the recovery and after spinal anesthesia ($p>0.05$, Table 3). None of the patients in both groups had dysrhythmia or bradycardia. A total of 11 (9.1%) patients in both groups had systolic hypotension. Eight (6.6%) patients in the two groups experienced diastolic hypotension. None of the patients in the two groups experienced increase in respiratory rate, oxygen saturation drop and oxygen desaturation drop after spinal anesthesia.

DISCUSSION

Our findings demonstrated that 6 mg of ondansetron and 8 mg of dexamethasone could equally curtail the incidence of nausea and vomiting in patients undergoing surgery through spinal anesthesia. In some countries including the United States, spinal anesthesia is the method of choice for elective emergency caesarean surgeries, whereas certain hospitals

adopt this method in 41% of patients.¹⁸

The effects of spinal anesthesia in pregnant and non-pregnant women vary. Distribution of anesthetic drug into the cerebrospinal fluid in pregnant women is less predictable, which is associated not only to increased pressure on the spinal canal,¹⁹ but also to a series of successive changes in the balance of acids and bases²⁰ and cerebrospinal fluid protein contents²¹ due to physiological changes during pregnancy. In addition, the side effects of spinal anesthesia, such as hypotension, nausea and vomiting, hypersensitivity to intrathecal opioids is more common in pregnant women compared to non-pregnant women.²²

In this study, a dose of 6 mg of ondansetron was chosen because is as effective in the prevention and treatment of nausea and vomiting after surgery similar to the higher dose. Moreover, there will not be any side effects at this dosage.²³ Pearman *et al.* suggested that the effect of 6 mg ondansetron might be more effective than 4 mg ondansetron in pregnant women who are more prone to nausea and vomiting.²⁴ Borgeat *et al.* found out the direct therapeutic and anti-nausea effect of sub-hypnotic doses of propofol in gynecological, gastroenterological and orthopedic surgeries. Nevertheless, subsequent

Table 3: Ondansetron and dexamethasone vital signs in both groups during recovery and after spinal anesthesia

Group	Group		p value
	Ondansetron	Dexamethasone	
Heart rate	71.06±4.91	71.13±4.86	0.062
Systolic blood pressure	125.86±4.98	125.60±5.07	0.064
Diastolic blood pressure	79.28±3.26	78.93±3.54	0.06
Respiration	12±2.7	12±2.2	0.19
Oxygen saturation	94%±2	96%±2.9	0.28

studies revealed that propofol can prevent nausea and vomiting in elective cesarean section surgery under spinal anesthesia.²⁵

In a study by Szarvas *et al.*, the incidence of nausea and vomiting in the first 24 hours after injection of intrathecal morphine was 70, 73 and 72 percent, respectively.²⁶ Pirat *et al.* showed that 8 mg of oral ondansetron and 4 mg of intravenous ondansetron did not prevent nausea and vomiting caused by intrathecal meperidine during surgery.²⁷ In another study (2009), the HT3-5 receptor antagonists were effective in prevention of nausea and vomiting caused by intrathecal morphine in women undergoing cesarean section.²⁸ In a study by Nortcliffe *et al.*, dexamethasone was not effective in preventing nausea and vomiting caused by intrathecal morphine.²⁹

In addition, the study by Wu *et al.* indicated that dexamethasone was not effective in the prevention of nausea and vomiting caused by intrathecal morphine.³⁰ The study by Tzeng *et al.* suggested that dexamethasone could curtail nausea and vomiting caused by epidural morphine in cesarean section.³¹ Movafegh *et al.* showed that 8 mg of dexamethasone could effectively reduce nausea and vomiting caused by intrathecal meperidine.³² The reason for such inconsistency of results in the prevention of nausea and vomiting can be hormonal changes, gender, age, weight, pain, type of surgery, duration of surgery, history of nausea and vomiting after surgery, intrathecal drug dose and type and dose of HT3-5 antagonists and dexamethasone.

This study demonstrated that 8 mg of dexamethasone and 6 mg of ondansetron were equally able to prevent intrathecal nausea and vomiting after surgery. Considering our findings, dexamethasone can be regarded as a safe alternative to ondansetron for prevention of nausea and vomiting after surgery.

ACKNOWLEDGMENTS

This paper was an outcome of a doctoral dissertation by Reza Dadar. We appreciate the kind financial support of Jahrom University of Medical Sciences.

CONFLICT OF INTEREST

The authors declare no conflict of interest.

REFERENCES

- 1 Moshiri E, Noruzi A, Pazuki SH, Gazerani N, Choghaei M. The effect of low dose ketamine on postoperative pain after spinal anesthesia in elective cesarean section. *Arak Med Univ J* 2011;**14**:81-8.
- 2 Juhani TP, Hannele H. Complications during spinal anesthesia for cesarean delivery: a clinical report of one year's experience. *Reg Anesth* 2000;**18**:128-31.
- 3 Pan PH, Moore CH. Intraoperative antiemetic efficacy of prophylactic ondansetron versus droperidol for cesarean section patients under epidural anesthesia. *Anesth Analg* 1996;**83**:982-6.
- 4 Kang YG, Abouelish E, Caritis S. Prophylactic intravenous ephedrine infusion during spinal anesthesia for cesarean section. *Anesth Analg* 2002;**61**:839-42.
- 5 Lussos SA, Bader AM, Thornhill ML, Datta S. The antiemetic efficacy and safety of prophylactic metoclopramide for elective cesarean section delivery during spinal anesthesia. *Reg Anesth* 1992;**17**:126-30.
- 6 Kestin IG. Spinal anesthesia in obstetrics. *Br J Anaesth* 2004;**66**:596-607.
- 7 Miller RD, Eriksson LI, Fleisher L, Wiener-Kronish JP, Young WL. Miller's Anesthesia. Volume 1 & 2. 7th ed. New York. Churchill Livingstone. 2009; pp. 6624-8.
- 8 Garcia Migvel FJ, Montao E, Martvicent V, Fuentes AL, Jasan Jose AL. Prophylaxis against intraoperative nausea and vomiting during spinal anesthesia for cesarean section. *Internet J Anesthesiol* 2000;**4**:1-7.
- 9 Apfel, Kranke P, Katz MH, Goepfert C, Papenfuss T, Rauch S. Volatile anaesthetics may be the main cause of easily but not delayed postoperative vomiting. *Br J Anaesth* 200;**88**:659-68.
- 10 Fujii Y, Numazaki M. Dose-Range effects of propofol for reducing emetic symptoms during cesarean delivery. *Obstet Gynecol* 2002;**99**:75-9.
- 11 Watcha MF, White PF. Postoperative nausea and vomiting its etiology, treatment and prevention. *Anesthesiology* 1992;**77**:162-84.
- 12 Elhakim M, Ali NM. Dexamethasone reduce postoperative vomiting and pain after tonsillectomy. *Can J Anaesth* 2003;**50**:392-7.
- 13 Henzi I, Walder B, Tramer MR. Dexamethasone for the prevention of post

- operative nausea and vomiting. *Anesth Analg* 2000;**90**:186.
- 14 Szarvas S, Chellapuri RS, Harmon DC, Owens J, Murphy D, Shorten GD. A comparison of dexamethasone, ondansetron and dexamethasone plus ondansetron as prophylactic antiemetic and antipruritic therapy in patients receiving intrathecal morphine for major orthopedic surgery. *Anesth Analg* 2003;**97**:259-63.
 - 15 Pirat A, Tuncay SF, Torgay A, Candan S, Arslan G. Ondansetron orally disintegrating tablets versus intravenous injection for prevention of intrathecal morphine induced nausea, vomiting and pruritus in young males. *Anesth Analg* 2005;**101**:1330-6.
 - 16 George RB, Allon TK, Habib AS. Serotonin receptor antagonists for the prevention and treatment of pruritus, nausea, and vomiting in women undergoing cesarean delivery with intrathecal morphine: a systemic review and meta- analysis. *Anesth Analg* 2009;**109**:174-82.
 - 17 Nortcliffe SA, Shah J, Buggy DJ. Prevention of postoperative nausea and vomiting after spinal morphine for cesarean section. Comparison of cyclizine, dexamethasone and placebo. *Br J Anaesth* 2003;**90**:665-70.
 - 18 Gibbs CP, Krischer J, Peckham BM, Sharp H, Kirschbaum TH. Obstetrics anesthesia: a national survey. *Anesthesiology* 1986;**65**:298-306.
 - 19 Shah JL. Effect of posture on extradural pressure. *Br J Anaesth* 1983;**55**:907.
 - 20 Mandal B, Batra YK, Varma YS. Acid-base changes of cerebrospinal fluid at full term pregnancy. *Indian J Med Res* 1988;**87**:605-8.
 - 21 Eisenach JC, Dobson CE 2nd, Inturrisi CE, Hood DD, Agner PB. Effect of pregnancy and pain on cerebrospinal fluid immunoreactive enkephalins and norepinephrine in healthy humans. *Pain* 1990 Nov;**43**:149-54.
 - 22 Echevarria M, Caba F, Bernal L, Pallarés JA, Rodríguez R. Influence of local anesthetic on visceral pain during cesarean section with intradural anesthesia. *Rev Esp Anesthesiol Reanim* 1996;**43**:2-6.
 - 23 Diemunsch P, Conseiller C, Clyti N, Marnet JP. Ondansetron compared with metoclopramide in the treatment of established postoperative nausea and vomiting. *Br J Anesth* 1997;**79**:322-6.
 - 24 Paech MJ, Pavy TJ, Evans SF. Single-dose prophylaxis for postoperative nausea and vomiting after major abdominal surgery: ondansetron versus droperidol. *Anaesth Intensive Care* 1995;**23**:548-54.
 - 25 Borgeat A, Wilder Smith OHG, Saiah M, Rifat K. Subhypnotic doses of propofol possess direct antiemetic properties. *Anesth Analg* 1992;**74**:539-41.
 - 26 Szarvas S, Chellapuri RS, Harmon DC, Owens J, Murphy D, Shorten GD. A comparison of dexamethasone, ondansetron and dexamethasone plus ondansetron as prophylactic antiemetic and antipruritic therapy in patients receiving intrathecal morphine for major orthopedic surgery. *Anesth Analg* 2003;**97**:259-63.
 - 27 Pirat A, Tuncay SF, Torgay A, Candan S, Arslan G. Ondansetron orally disintegrating tablets versus intravenous injection for prevention of intrathecal morphine induced nausea, vomiting and pruritus in young males. *Anesth Analg* 2005;**101**:1330-6.
 - 28 George RB, Allon TK, Habib AS. Serotonin receptor antagonists for the prevention and treatment of pruritus, nausea, and vomiting in women undergoing cesarean delivery with intrathecal morphine: a systemic review and meta- analysis. *Anesth Analg* 2009;**109**:174-82.
 - 29 Nortcliffe SA, Shah J, Buggy DJ. Prevention of postoperative nausea and vomiting after spinal morphine for cesarean section. Comparison of cyclizine, dexamethasone and placebo. *Br J Anaesth* 2003;**90**:665-70.
 - 30 Wu JI, Lo Y, Chia YY, Liu K, Fong WP, Yang LC, Tan PH. Prevention of postoperative nausea and vomiting after intrathecal morphine for cesarean section: A randomized comparison of dexamethasone, droperidol, and a combination. *Int J Obstet Anesth* 2007;**16**:122-7.
 - 31 Tzeng JI, Wang JJ, Ho ST, Tang CS, Liu YC, Lee SC. Dexamethasone for prophylaxis of nausea and vomiting after epidural Morphine for post caesarean section analgesia: comparison of droperidol and saline. *Br J Anesth* 2000;**85**:865-8.
 - 32 Movafegh A, Soroush AR, Navi A, Esfehiani F, Akbarian Tefaghi N. The effect of intravenous administration of dexamethasone on postoperative pain, nausea, and vomiting after intrathecal injection of mepridine. *Anesth Analg* 2007;**104**:987-9.