

# ORIGINAL ARTICLE

## Effect of Intravenous Ketamine on Prevention of Postoperative Shivering: A Comparison with Intravenous Pethedine

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### Abstract

**Introduction:** Postoperative shivering is one of the common problems following general anesthesia and may lead to multiple complications. This study aimed to compare the impact of Ketamine and Pethidine on postoperative shivering.

**Methods:** In a randomized clinical trial, one hundred and thirty-five patients undergoing elective abdominal surgery were recruited for the study. The patients were randomly divided to groups received Normal Saline, Ketamine or Pethidine at the end of surgery. The frequency of shivering was determined immediately after surgery, and at 10, 20, and 30 minutes, postoperatively.

**Results:** In zero time the rate of shivering was more severe than other times in the three groups. Also the rate of postoperative shivering was significantly less different in the pethidine and Ketamine groups than the Normal Saline group. However, no significant difference was seen between the pethidine and Ketamine groups.

**Conclusions:** The result of this study showed that exactly after operation, the rate of postoperative shivering was more severe than other times in 3 groups. Ketamine and Pethidine might reduce postoperative shivering with no significant difference.

**Key Words:** Postoperative shivering; Ketamine; Pethidine; general anesthesia

### Introduction

Shivering is a common complication following surgery [1]. It is the rhythmic contraction of muscles with frequency of 4-8 Hertz [2]. It is the natural response of the body to central hypothermia with release of cytokines from surgical site. It can also occur because of pain

Almost all anesthetics impair autonomic control of body temperature. Its incidence ranges from 5% to 65% following general anesthesia and 30% with epidural anesthesia [3, 4]. However, the incidence appears to be less as more patients are kept normothermic and opioids are administered more frequently and in larger doses than in the past.

Shivering is an unpleasant postoperative feeling

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that might be associated with adverse effects especially in patients with poor myocardial reserve. Shivering decreases mixed venous oxygen saturation as a result of impaired cardiopulmonary function. In addition, postoperative shivering can increase oxygen consumption up to 5 times, might increase carbon dioxide production, minute ventilation, and hence, cardiac output even in healthy adults [5]. It is a potentially serious complication that increases oxygen consumption roughly 100% in proportion to intraoperative heat loss.

It might also be associated with increased blood pressure, intracranial pressure, metabolic rate, lactic acidosis, and postoperative surgical wound pain [6, 7].

Ketanserin, sufentanil, alfentanil, tramadol, physostigmine, clonidine, magnesium sulfate, Pethidine, dexamethasone, and doxaperam have been used for the treatment of postoperative shivering [8].

Among these, Pethidine is widely used as the first line therapy [9, 10]. However, it might result in nausea and vomiting, delayed gastric emptying, and increased length of recovery stay.

N-methyl-D-aspartate agonists including Ketamine can affect temperature regulation and might be effective for the management of shivering. Nevertheless, they are associated with some neuropsychological adverse effects including somnolence and hallucination.

The purpose of this randomized clinical trial was to compare Pethidine and Ketamine in the prevention of postoperative shivering in patients undergoing general anesthesia. The main outcome is to compare the incidence of postoperative shivering.

## Methods

After approval from ethics committee of Mashhad university of medical science (Ethic code: 2083241), 135 females with ASA (American Society of Anesthesiologists) class of I or II and age of 18 to 65 years scheduled for elective abdominal surgery at Qaem Hospital, Mashhad, Iran were recruited for a randomized double blind clinical trial. The patients and the investigators were blind to the study. They were randomly allocated (using randomization table) to receive either Pethidine 20 mg (n=45), Ketamine 0.5 mg/kg (n=46), or placebo (n=44) at the end of the surgery. Exclusion criteria were history of neuropsychological diseases, body mass index (BMI) more than 30, history of seizure, drug allergy, hypertension and any other

cardiovascular diseases, intraoperative blood component transfusion, and surgeries lasting more than 3 hours. After receiving midazolam 0.05mg/kg and fentanyl 2 g/kg, anesthesia was induced with propofol 2 mg/kg and atracurium 0.5mg/kg for endotracheal intubation. Anesthesia was maintained with propofol 70-100µg/kg/min and N<sub>2</sub>O 50% in oxygen. Fentanyl was given 0.02µg/kg/min. The room temperature and the intravenous fluids were kept between 22 to 24 centigrade degrees.

Postoperative shivering was assessed immediately after surgery, and at 10, 20, and 30 minutes using the following scales:

Grade 0: no shivering

Grade 1: slight fasciculation in the neck and face

Grade 2: visible shivering in more than one muscle group

Grade 3: shivering and movement throughout the body

We considered a difference in shivering frequency of 20% be clinically significant. A sample size of 34 was required to detect such a difference between the groups for a power of 90% at a significance level of 5%.

Using SPSS software for Windows, version 15 (SPSS Inc, Chicago, IL, USA), arithmetic mean and standard deviation values for different variables were calculated and statistical analyses were performed for each group. We used independent Student *t*-test and ANOVA to compare continuous variables exhibiting normal distribution, and Chi-square and Mann-Whitney *U* test for non-continuous variables. *P* value less than 0.05 was considered significant.

## Results

135 patients completed the study. They aged from 18 to 60 years with mean age of 39±11.95 years and weight of 68.90±4.78kg. There was also no significant difference among the groups with respect to ASA class and duration of surgery (*p* values 0.07 and 0.1 respectively).

Shivering was more severe immediately after surgery in the three groups.

Immediately after surgery shivering with grade 0, 1, 2, and 3 in the Normal Saline group was seen in 27, 7, 5, and 5 cases, respectively, in the Pethidine group shivering with grade 0, 1, 2, and 3 was seen in 43, 1, 1, and 0 cases, respectively, and in the Ketamine group no one had shivering (grade 0). Using Post-hoc ANOVA (Tukey test) revealed that the difference was between the first and the second and third groups (*p* value<0.01). However,

**Table 1: Frequency of shivering in groups under study at 10 minutes**

	Normal Saline	Pethidine	Ketamine	P value
Grade 0	25	42	45	<0/001
Grade 1	9	1	1	<0/001
Grade 2	4	2	0	<0/001
Grade 3	6	0	0	<0/001

Data are presented as numbers

**Table 2: Frequency of shivering in groups under study at 20 minutes**

	Normal Saline	Pethidine	Ketamine	P value
Grade 0	29	44	46	<0/001
Grade 1	14	1	0	<0/001
Grade 2	1	0	0	<0/001
Grade 3	0	0	0	<0/001

Data are presented as numbers

**Table 3: Frequency of shivering in groups under study at 30 minutes**

	Normal Saline	Pethidine	Ketamine	P value
Grade 0	37	45	46	<0/001
Grade 1	7	0	0	<0/001
Grade 2	0	0	0	<0/001
Grade 3	0	0	0	<0/001

Data are presented as numbers

comparison of the Pethidine and Ketamine groups did not show significant difference (p value 0.15).

Shivering was seen more commonly in the Normal Saline than the other two groups at 10 minutes (table 1). Patients in the Normal Saline developed more shivering at 20 and 30 minutes, as well (tables 2, 3). In addition, frequency of shivering was similar in the Ketamine and Pethidine groups at all times.

## Discussion

Our study revealed that intravenous Ketamine and Pethidine can decrease postoperative shivering with no significant difference.

Most anesthetics can decrease thermoregulation thresholds resulting in hypothermia in patients exposing to the operating room. Hypothermia may lead to shivering, prolonged drug action, coagulopathy, and decreased resistance to infection. Therefore, maintaining normal body temperature is pivotal for prevention of hypothermia and its complications.

Postoperative shivering increases recovery length of stay and cost and is an additional burden to both the patients and hospitals.

Comparison of Pethidine, granisetron, and Normal Saline in laparoscopic surgery showed that Pethidine and granisetron could dramatically decrease postoperative shivering with no significant difference [10].

A number of studies showed the same result of efficacy of Pethidine and Ketamine in preventing postoperative shivering [11-15].

Similar to our study, Ketamine has been effective in preventing shivering after prostate [16] and general surgeries [17].

Application of Ketamine and Pethidine in children after tonsillectomy showed the same result with no significant difference between the two drugs [18].

In contrast, Pazuki et al revealed that Pethidine could better reduce postoperative shivering after cesarean section [19].

In another survey, it was reported that Pethidine and dexamethasone were similar in reducing shivering after abdominal surgery [20].

In addition to common drugs used, many researchers have used tramadol [21, 22], clonidine [23], dexamethasone (24), dexmedetomidine [25], acupuncture [26], ondansetron [27, 28], and

hydrocortisone [29] successfully for prevention of postoperative shivering.

## Conclusions

In conclusion, we demonstrated that shivering is more common immediately after surgery and intravenous Pethidine and Ketamine can significantly and similarly reduce postoperative shivering.

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