MAGNESIUM SULPHATE INFUSION PREVENTS SHIVERING DURING SPINAL ANAESTHESIA: A RANDOMISED DOUBLE BLINDED CONTROLLED STUDY

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ABSTRACT

Spinal anaesthesia is the most common form of anaesthesia in our hospital. Shivering is a very common complication in patients undergoing surgery under spinal anaesthesia. Shivering increases expenditure of cardiac and systemic energy resulting in increased oxygen consumption and carbon dioxide production, lactic acidosis and raises intraocular and intracranial pressure. It also interferes with haemodynamic monitoring intraoperatively. Regional anaesthesia produces vasodilatation, which facilitates core to peripheral distribution of heat. It also increases sweating threshold and decreases vasoconstriction and shivering threshold. Intra- and postoperative management of shivering is usually done by external heating (Forced air warming, warming blankets), warmed fluids or pharmacological intervention. Prevention and treatment of established shivering should be regarded as clinically relevant intervention in the perioperative period. Therefore, a randomized double-blind control study was done using magnesium sulphate infusion in the prevention of shivering in our hospital. The aim of our study is to assess the efficacy of magnesium sulphate when used for prevention of shivering following spinal anaesthesia.

METHODS

Institutional Ethical Committee approval and informed consent were obtained from all patients.

INCLUSION CRITERIA

80 patients of ASA 1 and 2 belonging to either sex, aged between 18-65 years undergoing lower abdominal and lower limb surgeries were included in the study.

EXCLUSION CRITERIA

Patients suffering from neuromuscular disease, hyperthyroidism, cardiopulmonary disease, psychological disease, refusal to participate or temperature more than 38*C or less than 36.5*C. The patients were randomized into 2 groups namely Group M and Group C. Group M - Each of Group M patients were designated to receive 2 gms (4 mL) of magnesium sulphate in the preloading fluid over a period of 20-30 minutes. Group C – Each of Group C patients were designated to receive 4 mL of normal saline as control in the preloading fluid over a period of 20-30 minutes. Following a detailed pre-anaesthetic checkup along with relevant investigations, patients were brought to the operation theatre and connected to monitors. Then the vital signs of the patients including heart rate, temperature, non-invasive blood pressure, respiratory rate and SpO2 were continuously monitored and recorded. Both group of patients were pre-loaded with 15 mL/kg of Ringer lactate before giving spinal anaesthesia.

GROUP M

The study drug of 4 mL of Inj. Magnesium Sulphate was coded and presented to the anaesthetist not involved in the management of the patient and administered by intravenous infusion by adding it in the preloading fluid before spinal anaesthesia over a period of 20-30 minutes.

GROUP C

The control drug of 4 mL of normal saline was coded and presented to the anaesthetist not involved in the management of the patient and administered by intravenous infusion by adding it in the preloading fluid before spinal anaesthesia over a period of 20-30 minutes. The temperature of the operation theatre was maintained at 22-24*C. After preloading, all patients were administered intrathecal injection of 12.5 mg of 0.5% preservative free heavy Bupivacaine. Intraoperative patients were monitored for shivering along with other vital parameters. Adequacy of subarachnoid block was ensured. Core and skin temperatures were also recorded.

MAIN RESULTS

Total incidence of shivering grade 3/4 was 5% in Group M and 30% in Group C (p < 0.01). Magnesium sulphate significantly reduced the incidence and intensity of shivering. The use of rescue drug Inj. Pethidine was more in Group C (30%) compared to 5% in Group C (p < 0.01).

CONCLUSION

For patients undergoing spinal anaesthesia, prophylactic Inj. Magnesium Sulphate infusion lowered incidence of shivering. When shivering did occur, rescue drug Inj. Pethidine proved to be an effective treatment with minimal side effects.

KEYWORDS

Prophylactic; Magnesium Sulphate; Shivering; Spinal Anaesthesia.

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INTRODUCTION

Shivering is a frequent complication following neuraxial anaesthesia. It is an involuntary, oscillatory muscular activity that augments metabolic heat production. Shivering may occur as a response to hypothermia. However, it may also occur in normothermic patients.^[1] Regional anaesthesia impairs both central and peripheral thermoregulatory control. In patients becoming sufficiently hypothermic, shivering may appear and is often disturbing to both patients and medical staff.^[2] Shivering may interfere with monitoring of heart rate, blood pressure and oxygen saturation.

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It is associated with substantial adrenergic activation.^[3] It increases oxygen consumption, lactic acidosis, carbon dioxide production and metabolic rate by up to 400%; thus, it may cause problems in patients with low cardiac and pulmonary reserves.^[1] Numerous studies have tested the efficacy of a large variety of interventions to prevent shivering in normothermic or hypothermic surgical patients, but the relative efficacy of these interventions, however, remains unclear.^[4] As the incidence of hypotension is high during regional anaesthesia, hypotensive agents including clonidine and urapidil may not be appropriate in preventing shivering. In addition, Pethidine (the most widely used agent) and tramadol may cause nausea and vomiting and respiratory depression during and after regional anaesthesia. The hypertensive and tachycardiac effects of ketamine limits its use.^[5] The search continues for drugs that sufficiently improve thermoregulatory tolerance without simultaneously producing excessive sedation or respiratory depression. Intravenous magnesium has been shown to suppress postoperative shivering suggesting that the agent reduces the shivering threshold.^[6] Recently, the addition of intravenous magnesium sulphate to a pharmacological antishivering regimen increased the cooling rate in unanaesthetised volunteers.^[7] The drug not only exerts a central effect,^[8] but is also a mild muscle relaxant^[9] and may thus simultaneously reduce the gain of shivering (Incremental shivering intensity with progressing hypothermia).[10]

This study was designed to assess the effectiveness of magnesium sulphate when given prophylactically in the control of shivering.

MATERIALS AND METHODS

Institutional Ethical Committee approval and informed consent were obtained from all patients.

Inclusion Criteria

80 patients of ASA 1 and 2 belonging to either sex, aged between 18-65 years undergoing lower abdominal and lower limb surgeries were included in the study.

Exclusion Criteria

Patients suffering from neuromuscular disease, hyperthyroidism, cardiopulmonary disease, psychological disease, refusal to participate or temperature more than 38*C or less than 36.5*C.

The patients were randomized into 2 groups namely Group M and Group C.

Group M

Each of Group M patients were designated to receive 2 grams (4 mL) of magnesium sulphate in the preloading fluid over a period of 20-30 minutes.

Group C

Each of Group C patients were designated to receive 4 mL of normal saline as control in the preloading fluid over a period of 20-30 minutes.

Following a detailed pre-anaesthetic checkup along with relevant investigations, patients were brought to the operation theatre and connected to monitors. Tympanic (core) temperature was measured by an ear thermometer. The aural probe was inserted until the patient felt it touching the tympanic membrane, the probe was then securely taped in place and the aural canal was occluded with cotton and a gauze bandage was positioned over the external ear. Axillary (Skin) temperature was measured by an axillary thermometer. Temperature difference was measured (DT=core temperature minus skin temperature). Core temperature less than 36 C was considered hypothermia. The vital signs of the patients including heart rate, temperature, non-invasive blood pressure, respiratory rate and SpO2 were continuously monitored and recorded.

Both Groups (Group M and Group C) of patients were preloaded with 15 mL/kg of Ringer lactate before spinal anaesthesia with the inclusion of study drugs in the preloading fluid.

Group M

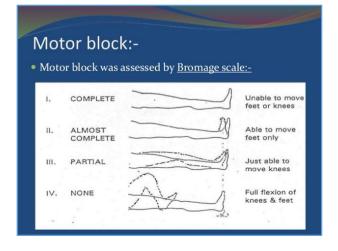
The study drug of 4 mL of Inj. Magnesium Sulphate was coded and presented to the anaesthetist not involved in the management of the patient and administered by intravenous infusion by adding it in the preloading fluid before the spinal anaesthesia over a period of 20-30 minutes. During infusion of Inj. Magnesium sulphate, the vital signs of the patients including heart rate, ECG, temperature, non-invasive blood pressure, respiratory rate and SpO2 were continuously monitored and recorded to pick up early the side effects of magnesium sulphate injection like heart rhythm disturbances, breathing difficulties, poor reflexes confusion, weakness, flushing, sweating, lowered blood pressure or headache.

The infusion was stopped if any adverse effect was observed in the patient, patient treated accordingly and patient excluded from study.

Group C

The control drug of 4 mL of normal saline was coded and presented to the anaesthetist not involved in the management of the patient and administered by intravenous infusion by adding it in the preloading fluid before the spinal anaesthesia over a period of 20-30 minutes.

The ambient temperature was maintained at 22–24 C. Spinal anaesthesia was performed at either L3/4 or L4/5 interspaces. Hyperbaric bupivacaine, 12.5 mg was injected using a 25-gauge spinal needle with the patients in the lateral decubitus position. Adequacy of spinal anaesthesia was ensured. Level of sensory blockade was assessed by pinprick and motor blockade by Bromage scale.^[11]



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Intraoperatively, patients were monitored for shivering along with other vital parameters. Core and skin temperatures were also recorded.

Intensity of Shivering was graded as per Crossley and Mahajan Scale

Scale	Description		
0	No Shivering		
	No visible muscle activity but piloerection,		
1	peripheral vasoconstriction or both are present		
	(other causes excluded)		
2	Muscular activity in only one muscle group		
3	Moderate muscular activity in more than one		
	muscle group, but no generalized shaking		
4	Violent muscular activity that involves the whole		
	body.		

Shivering grade 3/4 was regarded as significant.

Patients were given Inj. Pethidine 25 mg IV when shivering was significant as a rescue drug. At the end of the surgical procedure, the patients were shifted to Post Anaesthesia Care Unit for monitoring purpose.

Rescue Drug

Patients were given Inj. Pethidine Hydrochloride 25 mg IV when shivering was significant as a rescue drug.

Supplemental oxygen was delivered via a nasal cannula (4 L/min) if arterial blood saturation fell below 90%. These parameters were recorded preoperative after induction of anaesthesia, every 10 mins. till the end of surgery, then every 10 mins. Postoperative until 2 segments regression of the sensory blockade. Shivering was assessed after completion of subarachnoid injection until the patient was discharged after 2 segments regression of sensory blockade. Postoperatively, all patients were transferred to postoperative care unit (PACU) and monitored every 10 mins. until discharge.

Statistical analysis Data were analysed by using SPSS. (The Statistical Package for Social Sciences) version 17.0 software. Categorical data including total incidence of shivering and side effects were analysed using chi square. Independent t-test was used for quantitative data for comparison between two groups; paired t-test for quantitative data within each group; one-way ANOVA for quantitative data comparison between the 2 groups; Fisher's exact test for categorical data between the study groups; Mann-Whitney test for quantitative nonparametric data between two groups; Kruskal-Wallis Htest for quantitative nonparametric data between three groups; and Spearman correlation for correlation testing. Pvalue < 0.05 was considered significant.

RESULTS

A total of 80 patients completed this study, 40 in each group. The two groups were similar as regards age, weight, height and ASA physical status (Table 1). They were also comparable in duration of anaesthesia, surgery and amount of crystalloid administered. There was insignificant change as regards to mean heart rate, mean arterial blood pressure, oxygen saturation, and mean respiratory rate in the two groups throughout the study period. There was no correlation between patients who shivered and core temperature. Total incidence of shivering grade 3/4 was 5% in Group M and 30% in Group C (Table 2). Shivering was considered significant

when the patient shivered at least to Grade 3.^[12] Use of rescue Inj. Pethidine was more in Group C (30%) compared to 5% in Group M (p < 0.05).

		Group M	Group C			
Sample size n		n=40	n=40			
Age in years		40.9+21	41.6+21.4			
Weight (kg)		53+20.1	53+19.2			
		Males 18	Males 20			
Sex n (%)		(45%)	(50%)			
Sex II (%)		Females 22	Females 20			
		(55%)	(50%)			
Type of	General surgery	16 (40%)	12 (30%)			
operation n (%)	Orthopaedic	18(45%)	20 (50%)			
11 (%)	Urologic	6(15%)	8 (20%)			
Table 1: Demographic Data and Type of Operation						
in the Study Groups						

Data are presented as Mean ± SD. n = number.

	Group M	Group C			
Total Number of patients	n=40	n=40			
Incidence of shivering %	5% (n=2)	30% (n=12)			
Table 2					

	Grou	рM	Group C			
	(n=4	40)	(n=40)			
Anaesthesia Duration (min.)	75.5±17		71.5±14.6			
Surgery Duration (min.)	71±16		69.3±13.9			
Crystalloids (cc) (Intra- operative)	1050±426.1		1025±372.6			
	None	n=33 (82.5%)	n=23 (57.5%)			
Drug Administration	Ephedrine	n=5 (12.5%)	n=5 (12.5%)			
Administration	Atropine	n=0 (0%)	n=0 (0%)			
	Pethidine	n=2(5%)	n=12 (30%)			
Shivering 3/4	n=2 (5%)		n=12 (30%)			
Table 3: Operative Data						

Data are presented as Mean ± SD, n = number. * p< 0.001 Group M vs. Group C.

DISCUSSION

Post-anaesthesia shivering is very common following neuraxial anaesthesia. In our study, shivering was reported in 30% of patients in the control group C and 5% in the prophylactic groups similar to incidence reported on earlier studies.^[13] The exact mechanism of shivering during regional anaesthesia has not been fully established. Post-spinal shivering is probably a combination of both thermoregulatory and non-thermoregulatory shivering.^[1] Thermoregulatory

shivering is a response to hypothermia due to redistribution of body heat from the core to the periphery.^[14] The resultant vasoconstriction and shivering are restricted to the upper body due to the sympathetic and somatic neural blockade below the level of spinal blockade.^[15]

In our study, we found that post-spinal shivering may appear within minutes after injection of local anaesthetic and long before sufficient time has elapsed for significant heat loss to have occurred; therefore, other mechanisms of shivering exist. The same finding led Bromage.^[11] to postulate that spinal cord response is secondary to misinterpretation of afferent thermal clues due to differential blockade of warm and cold sensation, which is also supported by Kaoy et al.^[16]

Also, shivering has been reported in normothermic patients suggesting that inhibited spinal reflexes, apprehension, decreased sympathetic activity, pyrogen release, adrenal gland suppression and respiratory alkalosis are also involved.^[17]

A wide variety of agents with different mechanisms of action including opioids, non-opioid analgesics, 5-HT3 antagonists, a2 adrenoceptor agonists and cholinomimetic agents have been studied in an attempt to prevent or treat shivering, but unfortunately there is no "gold-standard" drug for the prevention of shivering.

The search continues for drugs that sufficiently reduce the shivering threshold and improve thermoregulatory tolerance without simultaneously producing excessive sedation or respiratory depression.^[4] Magnesium is a naturally occurring calcium competitor and a noncompetitive antagonist of Nmethyl-D-aspartate (NMDA) receptors.[18] The exact protective mechanism of MgSO4 remains uncertain; magnesium prevents or controls convulsions by blocking neuromuscular transmission and decreasing the release of acetylcholine at the motor nerve terminals.^[19] Parenteral magnesium sulphate is the drug of choice for the prevention and treatment of eclamptic convulsions.[20] It is an effective treatment for postoperative shivering following general anaesthesia.^[6] Magnesium has been reported to reduce shivering threshold in healthy volunteers,[21] as treatments that reduce the shivering threshold by a few tenths of a degree Celsius may be sufficient to attenuate postoperative shivering.[22]

In our study, core hypothermia occurred in both groups. It is likely that larger doses of MgSO4 would produce greater thermoregulatory effects, but carries a greater risk of complications as the thermoregulatory response to most IV drugs is linear with plasma concentration.^[23]

Patients receiving prophylactic magnesium in our study had a lower incidence of shivering than the control group. This is due to its mild muscle relaxant effect, thereby reducing the gain of shivering (Incremental shivering intensity with progressing hypothermia). Magnesium has been suggested as a possible means of resolving muscle rigidity and spasm in tetanus.^[24]

An earlier study by Gozedemir et al achieved an incidence of shivering 6.7% compared to 5% in our study.^[25]

Kizilirmak et al used 30 mg/kg bolus as the treatment for postoperative shivering following general anaesthesia and found that it was as effective as meperidine, but meperidine stopped shivering slightly faster than magnesium.^[6] None of the differences, however, was statistically significant.^[26] Hypotension is due to the vasodilating effect of MgSO4. This property was used by Ryu et al, to induce controlled hypotension in middle ear surgery.^[27] and also reduce blood loss during endoscopic sinus surgery.^[28]

CONCLUSION

From our study we conclude that for spinal anaesthesia, prophylactic MgSO4 infusion lowered incidence of shivering. Inj. Pethidine also proved to be an effective treatment when postspinal shivering occurred with minimal side effects.

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