ORIGINAL RESEARCH

Study Comparing Bolus Intravenous Ephedrine and Phenylephrine for Maintenance of Arterial Blood Pressure in Cesarean Section During Spinal Anesthesia

T. Mohan Singh¹, J. Rajkumar²

ABSTRACT

Introduction: Presently the mainstay in management of hypotension is the use of Vasopressor agents, but those currently available are not ideal and there has been an ongoing search for ideal vasopressor. Ephedrine is supposed to be most widely used agent for this purpose, but causes maternal tachycardia due to its non selective action on both alpha and beta adrenergic receptors. It is difficult to titrate and also exhibits the phenomena of tachyphylaxis due to its indirectly acting nature. Hence there is a need to find an alternative drug for the treatment of hypotension, which lacks the forementioned maternal side effects and also does not cause any detrimental effect on the fetus. Objective of the study was to compare the efficacy of intravenous bolus dose of Phenylephrine (40 mcg) with Ephedrine (6mg) to maintain arterial blood pressure during spinal anaesthesia for Caesarean section.

Material and Methods: The present randomized controlled study of 100 patients undergoing surgery under spinal anaesthesia for elective and emergency caesarean section was conducted in the Department of Anaesthesiology, MediCiti Institute of Medical Sciences, Ghanpur village, Medchal Mandal, Ranga Reddy District, Telangana state during the period of Dec 2012 – June 2014. 100 patients were randomly divided into two groups of 50 each, Group P (Phenylephrine group) and Group E (Ephedrine group). APGAR score of every neonate was noted at 1 min and 5 mins after delivery. Results: The data of subarachnoid block to hypotension time and duration of surgery were compared. There was no statistically significant difference found in the two groups. Group P required average of 1.20 boluses whereas Group E required 1.40 as boluses. Mean dose of 48 mcg was required in Group P and 8.4 mg was required in Group E. It was observed that mean value of APGAR score at 1 min in Group P was 9.66± 0.478 and that of Group E was 9.58 ± 0.49 . The APGAR 1 min Score difference between the two groups is not statistically significant as indicated by p value of greater than 0.05. It was observed that mean value of APGAR score at 1 min in Group P was 9.78± 0.418 and that of Group E was 9.78 ± 0.41 . The APGAR 5 min Score difference between the two groups is not statistically significant as indicated by p value of greater than 0.05.

Conclusion: Phenylephrine in a dose of 40 microgram is more efficacious in comparison to Ephedrine 6 milligrams in maintaining arterial blood pressure following hypotension after spinal anaesthesia in Caesarean section.

Keywords: Ephedrine, phenylephrine, spinal anesthesia

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INTRODUCTION

Presently the mainstay in management of hypotension is the use of Vasopressoragents, but those currently available are not ideal and there has been an Ongoingsearch for ideal vasopressor.¹

Ephedrine is supposed to be most widely used agent for this purpose, butcauses maternal tachycardia due to its non selective action on both alpha and betaadrenergic receptors.² It is difficult to titrate and also exhibits the phenomena of tachyphylaxis due to itsindirectly acting nature. Hence there is a need to find an alternative drug for thetreatment of hypotension, which lacks the forementioned maternal side effects and also does not cause any detrimental effect on the fetus.³ Phenylephrine is one such drug, which is a directly acting sympathomimeticagent with selective alpha1 adrenergic activity. It is easy to titrate and maintain maternalblood pressure without producing undue tachycardia. Moreover, the administration of Phenylephrine is reported to be associated with lower incidence of fetal acidosis than is Ephedrine.⁴ Numerous studies⁵⁻⁸ have been carried out to evaluate the various doses of Phenylephrine, from 20 to 100 micrograms and a dose of 20 mcg has been found to beineffective but dose as high as 100 mcg has caused maternal bradycardia.

Hence, an attempt has been made to compare the efficacy of

intravenousPhenylephrine 40 micrograms and Ephedrine 6 milligram as a vasopressor therapy incase of hypotension associated with spinal anaesthesia in patients undergoingCaesarean section. A good control of maternal blood pressure is one of the mainstayrequirements for ensuring safety during spinal anaesthesia in Caesarean section.9

MATERIALS AND METHODS

Study design

A randomized controlled study of patients undergoing surgery underspinal anaesthesia for elective and emergency caesarean section

Sample size

A total sample size of 100 cases

Inclusion Criteria

Uncomplicated pregnancy

- Weight: 0 to 70kg's
- Aged between 24 to 30 years
- ASA class I & II
- Baseline systolic blood pressure between 100 to 140
- Baseline diastolic blood pressure between 70 to 89 mm Hg and developedhypotension during the surgery

Hypotension is defined as fall in systolic pressure greater than 20% from baseline value or value less than 90 mm Hg

Exclusion Criteria

- Patient refusal
- Patient with medical complications like Diabetes mellitus, cardiovasculardiseases, severe anemia
- Patient with obstretical complications like Ante Partum hemorrhage, Pregnancy induced hypertension.
- Contraindications to spinal anaesthesia

Methodology

Approval from institutional ethical committee was obtained prior to study. After explaining the anaesthetic procedure, written informed consent for Participation inthe study was obtained from the patient. 100 patients were randomly divided into twogroups of 50 each, Group P (Phenylephrine group) and Group E (Ephedrine group)using computer generated randomization table. Study drugs were prepared anddispensed in syringes labelled "Study Vasopressor" by an anaesthesiologist notinvolved in the study. In the operating room Spo2,ECG,NIBP,HR monitored. The baselineheart rate, systolic, diastolic and mean blood pressures were measured. Intravenous preloading was done with 10ml / kg of Ringer lactate solution. Preloading was givenover 15 minutes. Following this, all patients received intrathecal 2 ml of 0.5% hyperbaric Bupivacaine through 23 G Quincke needle at L3-L4 subarachnoid space inleft lateral position. Immediately following the injection, patient was turned to supineposition and received oxygen at the rate of 5 Litre / min by face mask. Level of sensoryblock was assessed by pinprick method from below upwards 5 minutes afterSAB. Heart rate, systolic diastolic and mean arterial pressures were recorded every 2minutes for 20 minutes and thereafter every 5 minutes for 1 hour or till the end of thesurgery.

Whenever Hypotension (fall in BP > 20% of baseline or less than 90 mm of Hg)occurred, Group P received Phenylephrine 40 mcg and Group E received 6 mg

Ephedrine as intravenous bolus. Time taken to develop hypotension and the number ofboluses administered were noted. APGAR score of every neonate was noted at 1 min and 5 mins after delivery.

Statistical analysis

Statistical analysis was done using Student's t test and Mann-Whitney test.'p' value of less than 0.05 was considered to be statistically significant.

RESULTS

The objective of the present study is to compare the efficacy of intravenousbolus dose of Phenylephrine (40 mcg) and Ephedrine (6mg) to treat hypotensionfollowing spinal anaesthesia for Caesarean section.

A total of 100 patients were grouped into two groups, Group P (Phenylephrinegroup, n = 50) and Group E (Ephedrine group, n = 50) by a computer generated randomization table. After collecting data in both the groups the observations and analysis of the dataare presented in tabular form.

The data of subarachnoid block to hypotension time and duration of surgery were compared. There was no statistically significant difference found in the two groups (p > 0.05).

Group P required average of 1.20 boluses whereas Group E required 1.40 asboluses. Mean dose of 48 mcg was required in Group P and 8.4 mg was required in Group E.

It was observed that mean value of APGAR score at 1 min in Group P was 9.66 ± 0.478 and that of Group E was $9.58\pm$ 0.49. The APGAR 1 min Score difference between the two groups is not statistically significant as indicated by p value of greater than 0.05. It was observed that mean value of AP-GAR score at 1 min in Group P was 9.78± 0.418 and that of Group E was 9.78 ± 0.41 . The APGAR 5 min Score difference between the two groups is not statistically significant as indicated by p value of greater than 0.05.

DISCUSSION

Since sympathetic blockade resulting in vasodilatation is the primary cause of fallin arterial pressure, use of vasopressors in conjunction with fluid preloading appears tobe a more logical approach to correct it. It has been shown that the percentagedecrease in placental circulation is related to percent of reduction in maternal arterial pressure but not absolute reduction in pressure.

Incidence of hypotension clearly depends on its definition. Hypotension in most of the studies has been defined as values ranging from 20 to 30 percent reduction from baseline systolic arterial pressure.⁵ In the present study hypotension was defined asdecrease in systolic arterial pressure 20 percent from the baseline systolic pressure.

Bradycardia was defined as fall in heart rate of 20 percent or more from then baselinevalue.

The role of intravenous vassopressors has been well established in case of postspinal hypotension during Caesarean section. Ephedrine acts directly as well asindirectly on alpha and beta adrenergic receptors. Phenylephrine has selective alphaadrenergic activity due to which it was considered to cause reduction in uterine bloodflow, but it has been mentioned that Phenylephrine causes less fetal acidosis thanEphedrine in a review by Ngan et al.¹⁰

The present study is compared only with the studies in which Phenylephrine and Ephedrine was given as IV bolus to treat hypotension following spinal anaesthesia incaesarean section and the studies that have used prophylactic IM and IV doses of thedrug as well as those studies which have given the drug as infusions are not taken into consideration.

Dinesh Sahu¹¹ et al studied 60 patients undergoing elective as well asemergency caesarean section under spinal anaes-

Time (minutes)	Group P	Group E	P value
SAB to hypotension	3.36 <u>+</u> 0.81	3.52 <u>+</u> 0.81	0.3256
Duration of surgery	68.28 <u>+</u> 8.46	69.40 <u>+</u> 8.18	0.5027

Table-1: Sub arachnoid block (SAB) to hypotension time and duration of surgery

	Mean num- ber of bolus	Standard deviation	Mean dose
Group P	1.2	0.45	48 mcg
Group E	1.4	0.64	8.4 mg

Table-2: Number of bolus and mean drug dose required in each group

	Group P	Group E	P value
APGAR at 1 min.	9.66 <u>+</u> 0.478	9.58 <u>+</u> 0.49	0.41
APGAR at 5 min.	0.78 <u>+</u> 0.418	9.78 <u>+</u> 0.41	0.99
Table 3. Comparison of APGAR Scores in both the Groups			

thesia who developed hypotensionafter subarachnoid block. They were randomly allocated to one of three groups toreceive an IV bolus of the following Group P Phenylephrine 100mg (n=20), Group EEphedrine 6mg (n=20) or Group M Mephentermine 6mg (n=20). Hypotension wasdefined as a decrease in systolic arterial pressure > 20% of baseline values or < 90mmHg, whichever was greater. They concluded that elevation of systolic arterial pressurein Phenylephrine group was significantly high for first 6min of bolus dose as compared to Ephedrine & Mephentermine groups. There was significant reduction in heart rate inPhenylephrine group. Neonatal Appar score were >7 in all three groups.

Bhattarai et al¹², study compares three vasopressors-Phenylephrine, Ephedrine, Mephentermine. Study included 90 patients undergoing elective andemergency caesarean section who developed hypotension following sub-arachnoidblock. They were randomly divided into 3 groups, each group of 30 each. Group Preceived bolus of Phenylephrine 25mcg, Group E 5mg of Ephedrine, Group M 6mg of Mephenteramine. It was found that rise of BP was significantly higher in case of Phenylephrine group in first 6 minutes, after the bolus, there was significant reduction inheart rate in Phenylephrine group, but there was tachycardia following administrationof bolus Ephedrine and Mephentermine. Neonatal APGAR scores were similar in allthe groups.

The median value of level of anaesthesia in Sahu et al study in Group P is 8 whereas in Group E is 6. The mean value of level of anaesthesia in present study in Group P is 6.96 ± 0.85 whereas in Group E is 7.16 ± 0.738 . The values of the present study are comparable with Sahu et al study. The mean value of Sub-arachnoid block (SAB) to hypotension time in Sahu et al study in Group P is 4.3 whereas in Group E is 3.4. The mean value of Sub-arachnoid block (SAB) to hypotension time in Bhattarai et al study in Group P is 3.63 \pm 3.33 whereas in Group E is 4.55 \pm 0.42. The mean value of Sub-arachnoid block (SAB) to hypotension time in Present study in Group P is 3.36 ± 0.81 whereas in Group E is 3.52 \pm 0.81. The mean value of the Sub-arachnoid block (SAB) to hypotension time in present study are comparable with the study of Bhattarai et al & Sahu et al.

The mean value of APGAR Score 1 minute in Bhattarai et al study in Group Pis 7.40 ± 0.814 whereas in Group E is 7.31

	SAB to hy	SAB to hypotension		Level of anesthesia	
	Group P	Group E	Group P	Group E	
Sahu et al ¹¹	4.3	3.4	8	6	
Bhattarai et al ¹²	3.63 <u>+</u> 3.33	4.55 <u>+</u> 0.42	-	-	
Present study	3.36 <u>+</u> 0.81	3.52 <u>+</u> 0.81	6.96 <u>+</u> 0.85	7.16 <u>+</u> 0.738	
,	Table-4. Comparison of SAR to	hypotension time & Level	of Anaesthesia withother st	udies	

	APGAR at 1 min.		APGAR at 5 min.	
	Group P	Group E	Group P	Group E
Bhattarai et al ¹²	7.40 <u>+</u> 0.814	7.31 <u>+</u> 0.806	9	9
Present study	9.66 <u>+</u> 0.478	9.58 <u>+</u> 0.49	9.78 <u>+</u> 0.418	9.78 <u>+</u> 0.41
	Table-5: Comp	parison of APGAR Score w	rith other studies	

± 0.806. The median value of APGAR Score 5 minutes in Bhattarai et al study in Group P is 9 whereas in Group E is 9. The mean value of APGAR score 1 minute in Present study in Group P is 9.66 ± 0.478 whereas in Group E is 9.58 ± 0.49 . The mean value of APGAR Score 5 minutes in Present study in Group P is 9.78 ± 0.418 whereas in Group E is 9.78 ± 0.41 . Other studies have compared neonatal outcome based on umbilical cord bloodpH for both the drugs.

In Present study only APGAR score was compared, and there was no significant effect on the APGAR score of neonate at 1 and 5 minutes in both the groups, which correlated well with study by Sahu et al The mean value of APGARScore 5 minutes in Present study is comparable with that of Bhattarai study inboth group.

CONCLUSION

Phenylephrine in a dose of 40 microgram is more efficacious in comparison to Ephedrine 6 milligrams in maintaining arterial blood pressure followinghypotension after spinal anaesthesia in Caesarean section.

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