Efficacy of Oxytocin in Reducing Blood Loss During Abdominal Myomectomy

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ABSTRACT

Introduction: The drug of choice for preventing postpartum uterine atony and bleeding is oxytocin. Myomectomy is a procedure that may result in substantial bleeding. In about 20% women, transfusion of blood may be required during abdominal myomectomy. The objective was to compare the frequency of patients who needed blood transfusions with and without oxytocin during abdominal myomectomy.

Place and duration of study: This was a randomized controlled trial carried out at the Department of Obstetrics and Gynecology of Sharif Medical Complex Lahore from 10th May 2018 to 10th November 2018.

Material & Methods: 108 patients having intramural uterine fibroid > 5 cm in size, requiring planned elective open abdominal myomectomy were included and equally divided in both oxytocin and non-oxytocin groups(n=54). 30 IU oxytocin infusion (in 500 ml normal saline) / 120 milliliter per hour were administered during myomectomy, starting after administration of general anesthesia and completing just prior to surgery in patients of oxytocin group. An infusion of pure normal saline was administered at the same amount in the patients of non-oxytocin group. Blood loss was estimated and need for blood transfusion was determined in both the groups. Data was analyzed using SPSS version 24. P-value<0.05 was considered significant.

Results: Patients age ranged from 35 to 45 years with mean age of 40.481 ± 1.84 years in Oxytocin group while 40.425 ± 2.68 years in Non-Oxytocin group. Mean blood loss was 290.463 ± 120.91 cc in Oxytocin group and 424.981 ± 106.21 cc in Non-Oxytocin group. Need for blood transfusion was seen in 7.4% patients in Oxytocin group as compared to 25.9% in Non-Oxytocin group (p=0.009).

Conclusion: Our study concludes that to reduce loss of blood during abdominal myomectomy, infusion of intraoperative oxytocin may be efficient and safe. To confirm these results, more studies with varied dosages and protocols are required.

Keywords: Abdominal myomectomy, Oxytocin, Blood transfusions

INTRODUCTION

Henry Dale found in 1906 that discovered that human posterior pituitary gland extracts have the ability of contracting the pregnant cat uterus¹. In 1953, du Vigneaud sequenced and synthesized the peptide involved, known as oxytocin².

Oxytocin plays a role in a wide range of physiological & pathological activities, particularly reproduction such as parturition, breastfeeding, maternal behavior, sexual dysfunction, and orgasm³. The drug of choice for preventing postpartum uterine atony and bleeding is oxytocin⁴. The drug must be used with caution, since an intravenous injection of oxytocin in the amount of 10 IU might be harmful to hypovolemic women and women with cardiac problems. While receptors for oxytocin are also present in uterus of non-pregnant women, but their concentration is substantially lower as compared to pregnant women⁵. Because of this, oxytocin's non-pregnancy clinical application is

restricted. Myomectomy is a procedure that may result in substantial bleeding⁶. In about 20% women, transfusion of blood may be required during abdominal myomectomy⁷. A study done by Atashkhoei S, et al. reported that in oxytocin group, 7.5% of the women required blood diffusion while in non oxytocin group, 25% of the women required blood transfusion in abdominal myomectomy⁸. Another study carried out by Agostini A, et al. reported no significant difference in perioperative loss of blood in both oxytocin group and the non oxytocin group⁹. A study done by Atashkhoei S, et al reported oxytocin ability in reduction of hemorrhage and need of blood transfusion during myomectomy8 while another randomised doubleblind study reported no significant ability of oxytocin in reducing loss of blood during abdominal myomectomy9. Due to this variability in results and no such study yet done in general population of Pakistan, this study was planned to compare the frequency of patients need blood transfusions with without oxytocin during abdominal myomectomy in our general population. This



research will be useful in determining the role of oxytocin to reduce bleeding and the need for blood transfusions after myomectomy.

MATERIAL AND METHODS

This was randomized controlled trial carried out at the Department of Obstetrics and Gynecology of Sharif Medical Complex Lahore (SMC) from 10th May 2018 to 10th November 2018. Approval for the study was given by Hospital Research and Ethical Committee of SMC pursuant to letter number CPSP/REU/OBG-2015-082-6697. The inclusion criteria for our study was women age 35-45 years, intramural fibroid diagnosed on ultrasound imaging as per operational definition, uterine fibroid > 5 cm in size, planned for elective open abdominal myomectomy of uterine fibroids and ASA score I/II while the exclusion criteria was patients with history of cardiac disease, H/o Diabetes, hemoglobin concentration <10 g/dl on laboratory accompanying adenomyosis and endometriosis and patients with preoperative treatment with GnRH analogues. Totally, 108 patients were included. 54 patients were included in both oxytocin and nonoxytocin group. An informed consent in written was taken from all the patients. Baseline demographic information like age, number, and size of fibroids (by ultrasound) of patients was taken. During myomectomy, a 30 IU oxytocin infusion (in 500 ml normal saline) was given at a speed of 120 milliliter per hour after administration of general anesthesia and just before the surgery in patients of oxytocin group. An infusion of pure normal saline was administered at the same amount in the patients of non-oxytocin group. A pfannenstiel incision was used to expose the abdomen in all the surgeries. All procedures were done by consultant gynecologist of three years post fellowship experience. Blood loss and need for blood transfusion was noted. Data was recorded from both groups by the researcher and documented on predesigned proforma. All the data was analyzed statistically by using SPSS version 24. For qualitative data mean (SD) were calculated while for quantitative data, frequency (percentages) was calculated. Chi-square test was applied to compare need for blood transfusion in both groups, taken p ≤ 0.05 as significant.

RESULTS

A total of 108 patients were included in this research work. In our study, patients ages in Oxytocin group, ranged from 35-45 years with mean age of $40.481\pm~1.84$ years and $40.425\pm$ 2.68 years in the Non-Oxytocin group. Mean duration of complaints was 7.685±2.09 months in Oxytocin group and 6.925±1.72 months in Non-Oxytocin group. Mean size of fibroids was 8.611±1.69 cm in Oxytocin group 8.518±1.94 cm in Non-Oxytocin group. Mean number of fibroids was 3.092 ± 1.10 in Oxytocin group and 3.111 ± 1.17 in Non-Oxytocin group. Mean weight of patients was 79.388 ± 6.84 kg in Oxytocin group and 73.963±9.24 kg in the Non-Oxytocin group. Mean blood loss was 290.463±120.91 cc in Oxytocin group and 424.981±106.21 cc in Non-Oxytocin group.ASA I grade was dominant in both the groups. (Table-1) In Oxytocin group, need for blood transfusion was observed in 7.4% patients while in non-oxytocin group it was observed in 25.9% (p=0.009)patients (Table-2). Stratification of need for blood transfusion in both the groups with respect to age, duration of complaint, size of fibroids, number of fibroids, ASA grade and weight are shown in Table-3.

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Demographic		Oxytocin group	Non-oxytocin group	
Age(years) mean±SD		40.481± 1.84	40.425± 2.68	
Duration of Complain (months) mean±SD		7.685± 2.09	6.925± 1.72	
Size of fibroids(cm) mean±SD		8.611± 1.69	8.518± 1.94	
Number of fibroids mean±SD		3.092± 1.10	3.111± 1.17	
Weight(Kg) mean±SD		79.388± 6.84	73.963± 9.24	
Blood loss (cc)mean±SD		290.463± 120.91	424.981± 106.21	
ASA	I	44(81.5%)	39(72.2%)	
grade n (%)	II	10(18.5%)	15(27.8%)	

Table-1: Demographic features and ASA grade of patients in both the groups.

	n=54	n=54	
Need For Blood Transfusion	Oxytocin group	Non-oxytocin group	P- Value
Yes	4 (7.4%)	14 (25.9%)	
No	50 (92.6%)	40 (74.1%)	0.009
Total	54 (100%)	54 (100%)	

Table-2: Comparison of need for blood transfusion in both groups.

Para- meter	Sub-category	Group	Need for blood transfusion	N (%)	P- value	
	35-40 years	Oxytocin group	yes	2 (7.4%)		
Age	35-40 years	Oxytocin	No	25 (92.6%)		
	25.40	Non-		6	0.155	
	35-40 years	Oxytocin group	yes	(20.7%)		
	35-40 years	Non- Oxytocin group	No	23 (79.3%)		
	41-45 years	Oxytocin group	yes	2 (7.4%)		
	41-45 years	Oxytocin group	No	25 (92.6%)		
	41-45 years	Non- Oxytocin	yes	8 (32%)	0.024	
	41-45 years	Ron- Oxytocin group	No	17 (68%)		
	≤ 6 months	Oxytocin group	yes	0(0%)		
	≤ 6 months	Oxytocin group	No	16 (100%)		
	≤ 6 months	Non- Oxytocin	yes	6 (27.3%)	0.022	
	≤ 6 months	group Non- Oxytocin	No	16 (72.7%)		
Complaint Duration	6 months	Group Oxytocin	yes	4		
	6 months	group Oxytocin	No	(10.5%)		
		Non-		(89.5%) 8	0.109	
	6 months	Oxytocin group Non-	yes	(25%)		
	6 months	Oxytocin group	No	24 (75%)		
	5-10 cm	Oxytocin group	yes	0(0%)		
	5-10 cm	Oxytocin group	No	16 (100%)		
	5-10 cm	Non- Oxytocin group	yes	6 (27.3%)	0.022	
Size of	5-10 cm	Non- Oxytocin group	No	16 (72.7%)		
fibroids	>10cm	Oxytocin group	yes	4(10.5%		
	>10cm	Oxytocin group	No	34(89.5 %)		
	>10cm	Non- Oxytocin	yes	8(25%)	0.109	
	>10cm	group Non- Oxytocin	No	24(75%)		
	1-3 fibroids	Group Oxytocin	yes	0(0%)		
	1-3 fibroids	group Oxytocin	No	34(100%		
	1-3 fibroids	Ron- Oxytocin	yes	6(18.2%	0.009	
	1-3 fibroids	group Non- Oxytocin	No	27(81.8		
Number of	3 fibroids	group Oxytocin		%) 4(20%)		
fibroids		group Oxytocin	yes	4(20%)		
	3 fibroids	group Non-	No	16(80%) 8(38.1%	0.203	
	3 fibroids	Oxytocin group Non-	yes)	0.203	
	3 fibroids	Oxytocin group	No	13(61.9 %)		
	≤ 70 Kg	Oxytocin group	yes	0(0%)		
Weight	≤ 70 Kg	Oxytocin group	No	7(100%)	0.091	
	≤ 70 Kg	Non- Oxytocin group	yes	8(30.8%		
	≤ 70 Kg	Non- Oxytocin	No	18(69.2 %)		
	70 Kg	Oxytocin group	yes	4(8.5%)		
	70 Kg	Group Oxytocin	No	43(91.5	0.111	
	70 Kg	group Non-	yes	%) 6(21.4%		

		Oxytocin group)	
	70 Kg	Non- Oxytocin group	No	22(78.6 %)	
	ASA I	Oxytocin group	yes	3(6.8%)	
	ASA I	Oxytocin group	No	41(93.2 %)	
	ASA I	Non- Oxytocin group	yes	9(23.1%	0.035
ASA	ASA I	Non- Oxytocin group	No	30(76.9 %)	
grade	ASA II	Oxytocin group	yes	1(10%)	
	ASA II	Oxytocin group	No	9(90%)	
	ASA II	Non- Oxytocin group	yes	5(33.3%	0.180
	ASA II	Non- Oxytocin group	No	10(66.7 %)	

Table-3: Stratification of need for blood transfusion in both groups with respect to demographics and ASA grade.

DISCUSSION

In comparison to placebo, oxytocin (30 IU) infusion during abdominal myomectomy led to less intraoperative loss of blood and a lower requirement for blood transfusion, according to this research. Oxytocin increases the production and release of contractile prostaglandins by acting on receptors of oxytocin in the tissues of myometrium and fibroid. As the contractility of the uterus increases, the blood flow to the fibroids and arteries decreases. Because of the reduced volume of blood in uterus and restricted uterine vasculature caused by contraction of uterus and the vasoconstrictive impact of oxytocin, intraoperative loss of blood is reduced. Up to 20% of women having abdominal myomectomy need blood transfusions, according to reports^{7,10,12}. In our study, In Oxytocin group, need for blood transfusion was observed in 7.4% patients while in non-oxytocin group it was observed in 25.9% patients (p=0.009). A study done by Atashkhoei S, et al. reported that in oxytocin group. 7.5% of the women required blood diffusion while in non-oxytocin group, 25% of the women required blood transfusion in abdominal myomectomy⁸. In our study, no side effects were observed in using oxytocin. Tachyarrhythmias, hypotension and hyponatremia are all common oxytocin adverse effects^{13,14}. The absence of such problems in our research is most likely due to use of very low dose of oxytocin^{15,16}. In this research, the oxytocin groups mobilized about 3 hours sooner on average, which might have been a key component in reducing thromboembolic consequences. To reduce the risk of bias and selection a randomized controlled approach was used in this study. Small sample number and lack of follow-up of subjects following

release from the hospital are some of the study's We aimed to standardize drawbacks. computation and enhance its accuracy by utilizing a single, trained, and blind investigator to calculate intraoperative blood loss, but some error was The quantity of irrigation fluid unavoidable. gathered in the sponges could not be calculated. Even though we documented the number of fibroids removed during surgery, we did not weigh the total amount of fibroid mass removed, which, if it differed across groups, might have brought bias into our results. We did not utilize large dosages of oxytocin since there is not enough data to support their safety. In terms of minimizing blood loss, the advantages of giving oxytocin during myomectomy are limited. To confirm these results, more studies with varied dosages and protocols are required.

CONCLUSION

Our study concludes that to reduce loss of blood during abdominal myomectomy, infusion of intra-operative oxytocin may be efficient and safe. To confirm these results, more studies with varied dosages and protocols are required.

REFERENCES

- **1.** Dale HH. On some physiological actions of ergot. The Journal of physiology. 1906;34(3):163-206.
- 2. Du Vigneaud V, Ressler C, Trippett S. The sequence of amino acids in oxytocin, with a proposal for the structure of oxytocin. J Biol Chem. 1953;205(2):949-57.
- 3. R Goodin B, J Ness T, T Robbins M. Oxytocin-a multifunctional analgesic for chronic deep tissue pain. Curr Pharm Des. 2015;21(7):906-13.
- 4. Stanton CK, Newton S, Mullany LC, Cofie P, Tawiah Agyemang C, Adiibokah E, et al. Effect on postpartum hemorrhage of prophylactic oxytocin (10 IU) by injection by community health officers in Ghana: a community-based, cluster-randomized trial. PLoS Med. 2013;10(10):e1001524.
- 5. Huang M, Li X, Guo P, Yu Z, Xu Y, Wei Z. The abnormal expression of oxytocin receptors in the uterine junctional zone in women with endometriosis. Reprod Biol Endocrinol. 2017;15(1):1-10.
- **6.** Wan AY-H, Shin JH, Yoon H-K, Ko G-Y, Park S, Seong N-J, et al. Post-operative hemorrhage after myomectomy: safety and efficacy of transcatheter uterine artery embolization. Korean journal of radiology. 2014;15(3):356-63.
- 7. Raga F, Sanz-Cortes M, Bonilla F, Casañ EM, Bonilla-Musoles F. Reducing blood loss at myomectomy with use of a gelatin-thrombin

- matrix hemostatic sealant. Fertil Steril. 2009;92(1):356-60.
- 8. Atashkhoei S, Fakhari S, Pourfathi H, Bilehjani E, Garabaghi PM, Asiaei A. Effect of oxytocin infusion on reducing the blood loss during abdominal myomectomy: a double-blind randomised controlled trial. BJOG. 2017;124(2):292-8.doi:10.1111/1471-0528.14416.
- 9. Agostini A, Ronda I, Franchi F, Bretelle F, Roger V, Cravello L, et al. Oxytocin during myomectomy: a randomized study. European Journal of Obstetrics & Gynecology and Reproductive Biology. 2005;118(2):235-8.
- 10. Kongnyuy EJ, Van Den Broek N, Wiysonge C. A systematic review of randomized controlled trials to reduce hemorrhage during myomectomy for uterine fibroids. International Journal of Gynecology & Obstetrics. 2008;100(1):4-9.
- 11. Helal AS, Abdel-Hady E-S, Refaie E, El Shamy M, Abd El Fattah R. Preliminary uterine artery ligation versus pericervical mechanical tourniquet in reducing hemorrhage during abdominal myomectomy. International Journal of Gynecology & Obstetrics. 2010;108(3):233-5.
- 12. Pourmatroud E, Hormozi L, Hemadi M, Golshahi R. Intravenous ascorbic acid (vitamin C) administration in myomectomy: a prospective, randomized, clinical trial. Arch Gynecol Obstet. 2012;285(1):111-5.
- 13. Langesaeter E, Rosseland L, Stubhaug A. Hemodynamic effects of oxytocin during cesarean delivery. International Journal of Gynecology & Obstetrics. 2006;95(1):46-7.
- 14. Shokeir T, El-Lakkany N, Sadek E, El-Shamy M, Hashim HA. An RCT: use of oxytocin drip during hysteroscopic endometrial resection and its effect on operative blood loss and glycine deficit. J Minim Invasive Gynecol. 2011;18(4):489-93.
- 15. Acharya G, Al-Sammarai MT, Patel N, Al-Habib A, Kiserud T. A randomized, controlled trial comparing effect of oral misoprostol and intravenous syntocinon on intra-operative blood loss during cesarean section. Acta Obstet Gynecol Scand. 2001;80(3):245-.
- **16.** Chang F-W, Yu M-H, Ku C-H, Chen C-H, Wu G-J, Liu J-Y. Effect of uterotonics on intra-operative blood loss during laparoscopy-assisted vaginal hysterectomy: a randomised controlled trial. BJOG. 2006;113(1):47-52.

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