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Efficacy of Tranexamic Acid in Decreasing Blood Loss during and after Cesarean Section

Dr. Nasrin Akhter

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6 Abstract

3

Background: Tranexamic acid (TXA) is an anti-fibrinolytic agent. Its use in primary TKR is
supported by many studies that confirmed its efficacy for decreasing blood loss. Cesarean
section rates are increasing all over the world. Per partum hemorrhage is one of the most
common, life-threatening complications of this procedure. The aim of the study to find out

the efficacy of tranexamic acid in decreasing blood loss during and after cesarean section.

¹² Methodology: The present study was performed at the Department of Obstetrics and

¹³ Gynecology of the AL Haramine Hospital, Sylhet, from June 1, 2017, to November 30, 2017.

¹⁴ Subjects were eligible for the trial if the fetus was more than 38 weeks estimated gestational

¹⁵ age and they required elective CS. Elective CS was defined as CS performed before the onset

16 of labor.

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18 Index terms—

¹⁹ 1 Introduction

ranexamic acid (TXA) is an anti-fibrinolytic agent. Its use in primary TKR is supported by many studies that 20 confirmed its efficacy for decreasing blood loss. 1 Cesarean section rates are increasing all over the world. Per 21 partum hemorrhage is one of the most common, life-threatening complications of this procedure. 2 The WHO 22 estimates that PPH accounts for nearly 30% of maternal deaths worldwide with an estimated 20million cases 23 annually. 3 In order to reduce maternal morbidity and mortality caused by bleeding, it is important to reduce 24 25 the amount of bleeding during and after lower segment cesarean section. A popular approach is to minimize peri 26 operative bleeding through the prophylactic use of antifibrinolytic agents such as aprotinin, tranexamic acid and amino caproic acid. 3 Reducing bleeding during and after caesarian directly improve the outcomes of cesarean 27 delivery, especially maternal mortality and morbidity. Tranexamic acid is a fibrinolysis inhibitor that has been 28 used for many years to reduce bleeding in various surgical procedures. 29

30 **2** II.

31 3 Methodology

The present study was performed at the Department of Obstetrics and Gynecology of the AL Haramine Hospital, 32 Sylhet, from June 1, 2017, to November 30, 2017. Subjects were eligible for the trial if the fetus was more than 33 34 38 weeks estimated gestational age and they required elective CS. Elective CS was defined as CS performed 35 before the onset of labor. Women were excluded if they had risk factors associated with an increased risk of 36 postpartum hemorrhage (PPH) such as anemia (hemoglobin < 7 g%), multiple gestation, antepartum hemorrhage (placenta previa or placental abruption), abnormal placentation (accreta, increta, or percreta), uterine fibroids, 37 polyhydramnios, emergency CS, a history of uterine atony and postpartum bleeding, and a current or previous 38 history of significant disease, including heart disease, liver, renal disorders, or a known coagulopathy. Following 39 informed consent, simple randomization using a random number table was performed by the investigational 40 pharmacy staff, who took no further part in the study. Infusion bags were prepared and labeled as bag A (TA 41 group) containing 1 g/10 mL TA T 1 Year 2021 (Transamin, Fako Ilaclar? A.S., Istanbul, Turkey) diluted with 42

20 mL of 5% glucose, and bag B (placebo group) containing 30 mL of 5% glucose. Providers and patients were
blinded to the contents of the bags until the conclusion of the study.

TA or placebo was slowly administered intravenously over a 5-minute period at least 10 minute prior to skin

incision. After delivery, both groups received a 5 IU intravenous bolus of preprepared oxytocin, and then 30 IU

47 oxytocin in 500 mL lactated Ringer's solution was infused at a rate of 125 mL/h. An antibiotic, 1 g cefazolin
48 diluted in 20 mL normal saline, was administered over a 5-minute period. Vital signs (heart rate, blood pressure,

⁴⁹ and respiratory rate) were checked and noted before the operation, immediately after placental delivery, and 1

⁵⁰ and 2 hours after birth. Prothrombin time (PT), prothrombin time (aPTT), and complete blood count (CBC)

⁵¹ were performed before delivery and on the second day after delivery. Estimated blood loss was calculated using

52 the difference in hematocrit values taken prior to and 48 hour after cesarean delivery, according to the following 53 formula:

to contact one of the coauthors (O.A. or O.C.) immediately if any listed symptom occurred. Women who complained of symptoms were examined by one of the coauthors. All participants and babies were examined for thromboembolic events 3 and 6 weeks after surgery. The primary outcome was the estimated blood loss during CS. Other outcomes were excessive bleeding (defined as an estimated blood loss >1000 mL), the need for blood transfusion, the use of additional uterotonic agents indicating atony (such as an oxytocin infusion or

59 prostaglandin F2a), TA side effects (such as nausea, vomiting, or diarrhea), duration of the mother's postnatal 60 hospital stay, and neonatal outcome.

61 **4** III.

62 5 Results

The subject characteristics in the two groups were similar with no statistically significant difference between 63 age, height, weight, gestational age, gravida and duration of surgery (Table I). The heart rate, respiratory rate, 64 systolic BP and diastolic BP in both the Tranexamic acid and Placebo groups, Immediate after placental delivery, 65 1 hour after placental delivery and 2 hours after placental delivery, it was found that there was no significant 66 67 statistical difference (Table-II). Mean hemoglobin after surgery was 8.7 ± 0.7 g/dl and 8.1 ± 0.8 g/dl in tranexamic 68 acid and placebo group respectively (p>0.05). Mean hematocrit after surgery was 34.8 ± 2.9 percent in transxamic acid group and 32.3 ± 3.3 percent in placebo group (p>0.05). Which were statistically significant between two 69 groups (Table III). Compare the incidence of post-partum hemorrhage (PPH) between both study and control 70 groups, it was found that the incidence of post-partum hemorrhage (?500mL) was more in the control group 71 than in the study group (pvalue = 0.032) (Table -IV). where EBV (estimated blood volume) in mL¹/₄ the woman's 72 weight in kg_85.14 Blood loss >1000 mL during the procedure was defined as excessive bleeding. After discharge, 73 women who received TA were instructed about the signs and symptoms of a thromboembolic event, given written 74 instruction sheets with a diary for symptom documentation, and instructed 75

76 6 Discussion

In this present study it was observed that two groups were similar with no statistically significant differencebetween age, height, weight, gestational age, gravida and duration of surgery.

In this current study it was observed that heart rate, respiratory rate, systolic BP and diastolic BP in both the Tranexamic acid and Placebo groups, Immediate after placental delivery, 1 hour after placental delivery and hours after placental delivery, it was found that there was no significant statistical difference. Similar study carried out by Ming-ying Gai et al 4 in China showed that tranexamic acid significantly reduces bleeding from the time of placental delivery to 2 hours post partum. The study showed significant decrease in the incidence of 500 ml blood loss in the study group as compared to control group (P-0.029). Zheng et al , showed similar results after vaginal delivery.

In this study it was observed that mean hemoglobin after surgery was 8.7 ± 0.7 g/dl and 8.1 ± 0.8 g/dl in 86 tranexamic acid and placebo group respectively (p>0.05). Mean hematocrit after surgery was 34.8 ± 2.9 percent 87 in tranexamic acid group and 32.3 ± 3.3 percent in placebo group (p>0.05). Which were statistically significant 88 between two groups. Sekhavat et al. 6 in their prospective study on 90 primi gravidae undergoing CS, found 89 a decrease of blood loss from the end of lower segment CS to 2 h postpartum of only 9 ml (28 versus 37.12 ml 90 in TA and control groups, respectively). However, they found that this reduction was statistically significant. 6 91 This finding has not been described in other studies. 7,8 In this series it was observed that compare the incidence 92 of post-partum hemorrhage (PPH) between both study and control groups, it was found that the incidence of 93 94 post-partum hemorrhage (2500mL) was more in the control group than in the study group (pvalue = 0.032). 95 An estimated blood loss in excess of 500 ml following a vaginal birth or a loss of greater than 1000 ml following 96 caesarean birth has been used for the diagnosis of postpartum hemorrhage. 9 Assessment of blood loss during 97 cesarean delivery is typically underestimated. 10 The most commonly used methods for such assessment are the visual estimation method, the mathematical calculation measuring the hematocrit (Hct) prior to and 1 h 98 postoperatively and the swab weighing method. The visual estimation method is notoriously inaccurate 11; 99 Villeneuve et al. found that obstetricians assessed blood loss inaccurately during CS; an underestimation up to 100 579 ml 12. Brecher et al. found that blood loss calculated using Hct as a variable gave an average 2.1 times 101 overestimation of intraoperative blood loss compared to visual estimation. 10 The drawbacks of using Hct for 102

calculation of blood loss include the use of body weight in the mathematical calculation which is misleading 103 during pregnancy and the possible bias resulting from hemodilution from the intraoperative fluids given during 104 the operation. 13 Similar study carried out in India by Mayur et al. 14 showed comparable results reducing 105 the blood loss in the study group. Another study carried out by Ming-Ying et al., in China showed that TXA 106 significantly reduces bleeding from the time of placental delivery to the end of caesarean section, which was 351 107 mL in the study group while 440 mL in the control group. ??5 Zheng et al. showed similar results after vaginal 108 delivery; there was significantly less blood loss in the TXA group (243 mL) when compared to those who receive 109 no treatment (309 mL). ??6 V. 110

111 7 Conclusion

¹¹² Mean hematocrit after surgery was not statistically significant in tranexamic acid group than placebo group. ¹¹³ Compare the incidence of post-partum hemorrhage (PPH) between both study and control groups, it was found

- that the incidence of post-partum hemorrhage was more in the control group than in the study group.
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Year 2021				
Volume XXI	Estimated blood loss Preop hema-			
Issue II Ver-	tocrit _Postop hematocrit = EBV			
sion I	\times Preop hematocrit			
D D D D) E				
(
Global Jour-	Age (years) Height (cm) Weight	Tranexamic	(n=50)	p value
nal of Medi-	(kg) Gestational age (weeks) Gra-	acid $(n=50)$	$Mean \pm SD$	0.444 ns
cal Research	vidity	$Mean \pm SD$	$24.7 {\pm} 4.0$	$0.099 \ \mathrm{ns}$
		24.1 ± 3.8	$155.6 {\pm} 4.3$	0.233 ns
		$154.2 {\pm} 4.1$	$64.7 {\pm} 8.8$	$0.564 \ \mathrm{ns}$
		$66.6{\pm}6.9$	$38.4 {\pm} 0.7$	$0.537~\mathrm{ns}$
		38.3 ± 1.0	$2.0{\pm}0.9$	
		$2.1{\pm}0.7$	Placebo	
	Duration of surgery (min)	$44.8 {\pm} 2.6$	$44.7 {\pm} 2.7$	$0.851~\mathrm{ns}$
	ns = not significant			

Figure 1: Table I :

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arean Section

	Tranexamic acid (n=50) Mean±SD	Placebo (n=50) Mean±SD	p value
Immediate after placental delivery			
HR (beat/min)	$90.4 {\pm} 9.0$	$92.9 {\pm} 8.7$	$0.161~\mathrm{ns}$
RR (breaths/min)	$19.1 {\pm} 4.9$	20.5 ± 3.2	$0.093~\mathrm{ns}$
SBP (mmHg)	$120.6{\pm}10.8$	$124.1{\pm}11.8$	$0.125~\mathrm{ns}$
DBP (mmHg)	$76.0{\pm}10.1$	$79.3 {\pm} 9.8$	$0.100~\mathrm{ns}$
One hour after placental delivery			
HR (beat/min)	$92.0 {\pm} 9.1$	$88.7 {\pm} 8.9$	$0.070~\mathrm{ns}$
RR (breaths/min)	$20.2{\pm}6.1$	21.9 ± 5.2	$0.137~\mathrm{ns}$
SBP (mmHg)	127.2 ± 9.2	$124.4{\pm}11.1$	$0.173~\mathrm{ns}$
DBP (mmHg)	$78.5 {\pm} 8.7$	$80.3 {\pm} 10.6$	$0.356~\mathrm{ns}$
Two hours after placental delivery			
HR (beat/min)	$91.4{\pm}10.2$	89.3 ± 8.4	$0.264~\mathrm{ns}$
RR (breaths/min)	$19.8 {\pm} 5.6$	20.1 ± 3.3	$0.745~\mathrm{ns}$
SBP (mmHg)	124.6 ± 11.5	$124.1{\pm}12.0$	$0.832~\mathrm{ns}$
DBP (mmHg)	79.2 ± 8.1	79.4 ± 8.1	$0.902~\mathrm{ns}$
ns = not significant			

 \mathbf{II}

Figure 2: Table II :

\mathbf{III}

	Tranexamic acid	Placebo	
	(n=50)	(n=50)	P value
	$Mean \pm SD$	$Mean \pm SD$	
Hemoglobin (g/dl)			
Before CS	$9.8 {\pm} 0.8$	$9.7{\pm}1.0$	$0.582~\mathrm{ns}$
After CS	$8.7 {\pm} 0.7$	$8.1 {\pm} 0.8$	$0.001~{\rm s}$
Hematocrit (%)			
Before CS	$35.0{\pm}2.6$	34.1 ± 1.9	$0.051~\mathrm{ns}$
After CS	$34.8 {\pm} 2.9$	32.3 ± 3.3	$0.001~{\rm s}$
s = significant, ns = not significant			

Figure 3: Table III :

\mathbf{IV}

Blood loss from placental delivery to 2	Tranexamic acid	(n=50) n %	n Placebo (n=50) %	р
hours postpartum (ml)					value
<500mL ?500mL	43 7	86.0	$34\ 16$	68.0	$0.032~{\rm s}$
		14.0		32.0	

IV.

Figure 4: Table IV :

blood loses:a randomized parative, multicenter trial. Chin J Obstet Gynecol 2001; 36: 590-2.

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Figure 5:

7 CONCLUSION

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