



Massive Transfusion Protocol: A Boon for Salvaging Patients of Obstetric Hemorrhage

Smruti B Vaishnav^{a++}, Rashmita Pal^{a#*}, Sangita Pandey^{a†},
Mayur Shinde^{b‡} and Akshay Padaliya^{a#}

^a Department of Obstetric and Gynaecology, Shree Krishna Hospital, Gujarat, India.

^b Bhartiya Vidyapeeth, Pune, India.

Authors' contributions

This work was carried out in collaboration among all authors. Author SBV contributed to conceiving the idea, analysis, and manuscript preparation. Author RP wrote the project, performed data collection and analysis, and wrote the manuscript. Author SP also contributed to writing the project, analysis, and manuscript preparation. Author MS assisted with statistical data analysis. Author AP contributed to data collection and manuscript writing. All authors read and approved the final manuscript.

Article Information

Open Peer Review History:

This journal follows the Advanced Open Peer Review policy. Identity of the Reviewers, Editor(s) and additional Reviewers, peer review comments, different versions of the manuscript, comments of the editors, etc are available here: <https://www.sdiarticle5.com/review-history/122170>

Original Research Article

Received: 21/06/2024

Accepted: 23/08/2024

Published: 29/08/2024

ABSTRACT

Background: Maternal mortality remains a major global health issue, with obstetric hemorrhage as the primary cause. Timely and effective blood transfusion is crucial for maintaining organ perfusion and oxygenation. This study seeks to evaluate maternal outcomes, focusing on survival rates and life-threatening complications following massive blood transfusion.

⁺⁺ Head;

[#] Resident;

[†] Assistant Professor;

[‡] Statistician;

*Corresponding author: E-mail: palrashmitar@gmail.com;

Cite as: Vaishnav, Smruti B, Rashmita Pal, Sangita Pandey, Mayur Shinde, and Akshay Padaliya. 2024. "Massive Transfusion Protocol: A Boon for Salvaging Patients of Obstetric Hemorrhage". *Asian Research Journal of Gynaecology and Obstetrics* 7 (1):236-48. <https://journalarjgo.com/index.php/ARJGO/article/view/231>.

Materials and Methods: A prospective observational study was conducted between July 2022 to January 2024 in the Department of Obstetrics and Gynecology of Shree Krishna Hospital, Karamsad, Gujarat, India which is a rural tertiary care institute catering to Anand and Kheda districts of Central Gujarat. Data included in the study were age, socioeconomic status, parity, weeks of gestation, underlying comorbidities, cause of hemorrhage, mode of management, number and ratio of blood products transfused, and complications of Massive Transfusion (MT).

Results: The MT utilization rate of our institute was 2.74%. The mean age of the study group was 27.3yrs with 84% belonging to the rural population. Out of all 58% were antenatal cases majority being multipara- 61.7%. Post-Partum Hemorrhage (PPH) was the most common cause of massive obstetric hemorrhage i.e. 42%. At the same time, the average time for issuing the first blood product was 12 min. Most cases could be managed conservatively -31 out of 81 using oxytocics and timely blood transfusion. Amongst the operative interventions, obstetric hysterectomy was done in 23.4 % of cases. The overall ratio of Packed Cell Volume (PCV): Fresh Frozen Plasma (FFP): Platelet Concentrate (PC): Cryoprecipitate (CP) in the study was 1:1.02:0.8:2. In 46% of the cases, patients did not suffer from any MT-related complications; Transfusion Associated Circulatory Overload (TACO) was seen in 16% and Transfusion Related Acute Lung Injury (TRALI) in 7.4%. The mortality rate was 5%. Blood transfusion-related complications are observed more with PC, followed by FFP and RCC.

Conclusions: PPH was the leading cause of obstetric hemorrhage. Maternal morbidity and mortality can be significantly reduced through early referral to a tertiary care center, prompt administration of oxytocics, and the timely initiation of massive transfusion and surgical interventions for uncontrolled bleeding. Maintaining a blood product ratio of approximately 1:1.02:0.8 for packed red blood cells (PRBCs), fresh frozen plasma (FFP), and platelets helps prevent coagulopathy, ensures adequate tissue perfusion, and shields the patient from the detrimental cycle of sepsis, hypothermia, hemodilution, and shock. PRC transfusion was maximally responsible for TRALI.

Keywords: Massive transfusion protocol; post-partum haemorrhage; obstetric haemorrhage; maternal morbidity; maternal mortality.

ABBREVIATIONS

AKI	: Acute Kidney Injury
ANC	: Antenatal Care
APH	: Antepartum Hemorrhage
APTT	: Activated Partial Thromboplastin Time
ART	: Artificial Reproductive Techniques
ATLS	: Advanced Trauma Life Support
CP	: Cryoprecipitate
DIC	: Disseminated Intravascular Coagulation
FFP	: Fresh Frozen Plasma
Hb	: Hemoglobin
HDU	: High Dependency Units
INR	: International Normalized Ratio
IVC	: Inferior Vena Cava
LSCS	: Lower Segment Cesarean Section
MODS	: Multiorgan Dysfunction Syndrome
MOH	: Massive Obstetric Hemorrhage
MT	: Massive Transfusion
MTP	: Massive Transfusion Protocol
OSI	: Obstetrics Shock Index
PC	: Platelet Concentrate
PCV	: Packed Cell Volume
PLT	: Platelet

PNC	: Postnatal Care
PPH	: Postpartum Hemorrhage
PPROM	: Preterm Premature Rupture of Membranes
PROM	: Premature Rupture of Membranes
PT	: Prothrombin Time
RBC	: Red Blood Concentrates
RCC	: Red Cell Concentrate
SI	: Shock Index
SR canula	: Samarth Ram suction canula
TEG	: Thromboelastography
WHO	: World Health Organization

1. INTRODUCTION

The Massive Transfusion Protocol represents a revolutionary approach to early, proactive hemostatic resuscitation for patients at high risk of substantial blood loss over a short timeframe. Massive transfusion is defined as administering more than 4 units of red cell concentrates (RCC) within 1 hour or over 10 units within 24 hours, with massive blood loss defined as exceeding 150 ml/hr or one blood volume within 24 hours [1,2]. Severe obstetric hemorrhage is a major global issue, significantly contributing to maternal

morbidity and mortality. Causes of obstetric hemorrhage include placenta previa, placenta accreta, atonic and traumatic postpartum hemorrhage (PPH), uterine rupture, placental abruption, and uterine inversion. There is a notable increase in the incidence of massive obstetric hemorrhage, often associated with a higher rate of cesarean sections [3].

Haemostatic Resuscitation is a key approach that focuses on achieving local surgical haemostasis and mitigating coagulopathy by preventing hypothermia, acidosis, and ensuring the timely replacement of coagulation factors such as fibrinogen. In the context of massive obstetric hemorrhage (MOH), extensive bleeding leads to reduced blood flow to the uterus. This diminished perfusion causes tissue hypoperfusion, which in turn increases the production of thrombomodulin by vascular endothelial cells. Thrombomodulin activates protein C, setting off a cascade of irreversible events that result in elevated fibrinogen degradation products and impaired uterine contractions [4,5]. These effects are common in both atonic and traumatic postpartum hemorrhage (PPH), leading to rapid deterioration of the patient's condition. Consequently, implementing a Massive Transfusion Protocol (MTP) becomes a crucial strategy for effectively managing MOH.

Using a fixed ratio of red cell concentrate (RCC) to fresh frozen plasma (FFP) at 1:1, and a RCC:FFP:Platelet ratio of 1:1:1, has been shown to provide a survival advantage within the first 6-24 hours of massive hemorrhage, where mortality is primarily associated with hypovolemic shock and its effects [6]. Since the pathophysiology of obstetric bleeding is similar to that of severe trauma, applying a Massive Transfusion Protocol (MTP) with FFP/RCC or RBC ratios similar to those used in trauma care could be beneficial for managing obstetric hemorrhage [7]. To improve non-trauma MTPs, further prospective studies are needed to validate these ratios and to develop screening or prediction tools that can identify which patients will benefit most from MTP activation.

1.1 Aims

The present study to insight regarding maternal outcome, in the form of: life-threatening complications and subsequent survival or mortality following a massive blood transfusion for obstetric hemorrhage.

We secondarily also aimed to acquire information related to causes and management of massive obstetric hemorrhage, optimum blood products ratio and the time interval between requisition and receipt of the blood products.

2. MATERIALS AND METHODS

2.1 Study Design

This is a prospective observational study conducted in the Department of Obstetrics and Gynecology in Pramukh Swami Medical College, Karamsad, Gujarat, India from July 2022 to January 2024.

2.2 Study Subjects

Only Obstetric patients, both antenatal and postpartum, who had undergone massive obstetric hemorrhage necessitating Massive Transfusion Protocol were included in the study. Patients with gynecological hemorrhage were excluded in the study.

2.3 Setting

This prospective, observational study was conducted in the Department of Obstetrics and Gynaecology of Shree Krishna Hospital Pramukh Swami Medical College, Bhaikaka university, Karamsad, Anand, Central Gujarat, India. This is a 750-bedded tertiary care institute that caters to Anand and Kheda districts, over an area of about 50 km radius.

2.4 Data Setting

Data of the patients who had received MT was retrieved from our files and A.D. Gorawala blood bank. The following details about the patient were collected. The patient's detailed performa included patient ID number, age, profession, socioeconomic status, gravida, parity, details of current pregnancy, mode and time of delivery, cause of obstetric haemorrhage, coagulation profile, the timing of onset of haemorrhage to the onset of MT interval, number of blood products transfused, its relative ratio of RBC:FFP:Platelets and cryoprecipitates, hospital stay, ICU stay, the occurrence of complications like TRALI (transfusion-associated acute lung injury), TACO (transfusion-associated circulatory overload), Acute Renal Shutdown, Blood transfusion reaction, the need for mechanical ventilation, the

need of dialysis and vital organ failure and the final outcome as survival or mortality.

2.5 Ethical Clearance

The data collection was started after approval of the institutional ethics committee-IEC/BU/136/Faculty/1/293/2022, as it was a prospective and descriptive study, a waiver of consent was requested.

2.6 Statistical Analysis

Descriptive and multivariate logistic regression were used to analyse data by using the SPSS 29 software. In descriptive statistics mean [SD] and frequency [%] were used to depict the baseline profile of the study participants. A p value <0.05 was considered as statistically significant.

3. RESULTS

The study was conducted between July 2022 to January 2024. There were total 2956 obstetric admissions out of which 81 patients required

massive blood transfusion which accounts for 2.74% of the MTP utilization rates. Highlighting the demographical data, the minimum age of the patients requiring MTP was 19yrs and the maximum age was 38yrs. The mean age of the study population was 27.3 years.

Table 1 shows that out of the total 81 cases, 68 cases i.e. 84 % cases belonged to rural area. Maximum number of patients belonged to middle class socioeconomic status i.e. 51 cases out of the total 81 cases (63%). Amongst the study cases, antenatal patients were in the majority with 47 (58%) of the total cases, while the rest were post-partum patients. Majority of the patients were multipara-50 (61.7%) followed by primipara cases being 21 (26%). Parity of more than 4 were considered to be grand multipara. Atonic postpartum hemorrhage (PPH) was more common among multiparous women. The category of patients under 20 weeks included those with missed or incomplete abortions, ruptured ectopic pregnancies, abdominal pregnancies, and early-diagnosed placenta accreta syndrome.

Table 1. Demographic Variables

Variables	No. of Patients (n=81)	Percentage (%)
AGE		
15-20	6	7.4
21-25	22	27.2
26-30	33	40.7
31-35	14	17.3
36-40	6	7.4
Locality		
Rural	68	84
Urban	13	16
Socioeconomic Class		
Upper middle class	7	8.6
Middle middle class	51	63
Lower class	23	28.4
Pregnancy Status		
ANC	47	58
PNC	34	42
Parity		
Nullipara	10	12.3
Primipara	21	26
Multipara	44	54.3
Grand multipara	6	7.4
Gestational Age at Presentation		
<20	6	13.3
20-28	3	6.6
28-37	23	46.65
37-42	15	33.3
>42	0	0

Table 2. Causes of Hemorrhage

Causes of Hemorrhage	No. Of Cases	Percentage
ATONIC/ TRAUMATIC PPH	29	37
ABRUPTIO PLACENTA	17	21
DIC/ HELLP/ THROMBOCYTOPENIA	17	21
PLACENTA ACCRETA SYNDROME	8	9.8
PLACENTA PREVIA	4	5
RUPTURED ECTOPIC	3	3.7
RUPTURED UTERUS	2	2.5
ABDOMINAL PREGNANCY	1	1.2
TOTAL	81	100

At Shree Krishna Hospital, the largest proportion of patients requiring Massive Transfusion Protocol (MTP)—26 cases, or 32%—had undergone a lower segment cesarean section (LSCS). Most of these cases were complicated by antepartum hemorrhage, including placenta previa, placental abruption, and placenta accreta syndrome, necessitating either emergency or elective LSCS.

Additionally, 9 patients (11.1%) in the laparotomy category underwent procedures such as uterine rupture repair, hemoperitoneum drainage, stepwise devascularization, or obstetric hysterectomy. These cases included uterine rupture, ruptured ectopic pregnancy, abdominal pregnancy, placenta accreta spectrum, and traumatic PPH.

Table 2 shows that the most prevalent cause of massive obstetric hemorrhage was postpartum hemorrhage (PPH), accounting for 37% of cases. This category includes those cases specifically associated with atonic and/or traumatic PPH. Other contributors to PPH, such as thrombin defects and retained placenta, are detailed in separate sections: DIC/HELLP/thrombocytopenia (17 cases, 21%) and placenta accreta spectrum (PAS) (8 cases, 9.8%).

For cases of antepartum hemorrhage (APH), the most common cause was placental abruption, occurring in 17 cases (21%), while placenta previa was observed in 4 cases (5%). Abdominal pregnancy was the least common, with only 1 case out of the total 81 cases, and ruptured ectopic pregnancy was noted in 3 cases.

Many patients presented with multiple overlapping comorbidities. A significant portion, 66.6%, exhibited deranged coagulation profiles, which included conditions such as HELLP syndrome, DIC and thrombocytopenia. Among those with hypertensive disorders of pregnancy,

preeclampsia was the most common condition necessitating massive transfusions, affecting 43.2% of the cases. Additionally, 31% of the patients had a history of previous lower segment cesarean sections (LSCS). Sepsis was observed in 24.7% of the patients upon admission, often associated with atonic, traumatic, or secondary postpartum hemorrhage (PPH), particularly in cases involving multiple per vaginal examinations. Four patients had pre-existing cardiac conditions, including rheumatic heart disease with post-valve replacement, ongoing anticoagulant therapy, as well as, peripartum cardiomyopathy (PPCM). Acute kidney injury (AKI) was present in 14.8% of patients on admission, characterized by elevated serum creatinine levels exceeding 1.5 mg/dL, primarily due to pre-renal factors like acute blood loss. Furthermore, 29.6% of patients were diagnosed with multiorgan dysfunction syndrome (MODS) upon admission, attributable to underlying conditions such as HELLP syndrome, preeclampsia, and eclampsia. This often involved the simultaneous impairment of multiple systems, including hematological, renal, liver function, cardiac, respiratory, and/or central nervous systems (CNS).

In this study, the majority of postpartum hemorrhage (PPH) cases were attributed to uterine atony - 50.6% . Traumatic PPH was observed in 12.3% . Additionally, 7.4% of cases involved both atonic and traumatic PPH. Coagulation disorders were present in 17.3% of the cases, encompassing issues such as clotting factor abnormalities and thrombocytopenia. Secondary PPH, occurring between 24 hours and 6 weeks postpartum, was identified in 5 out of 81 cases. Other causes of PPH included ruptured ectopic pregnancies, placenta accreta spectrum (PAS), and abdominal pregnancies.

The shock index was the most frequently used parameter for assessing overall blood loss in

patients. This index is calculated by dividing the pulse rate by the systolic blood pressure (SBP). In the majority of cases, 52% of patients had a shock index greater than 1. Those presenting with severe shock typically had experienced a loss of more than 30% of their total blood volume.

The average time from the collection of the blood sample to the issuance of the first blood product was 12 minutes, in this desperate requisition O Negative blood was issued.

Table 3 depicts the severity of anemia the patients presented with. The minimum was 1.8gm/dl and the mean was 6.5gm/dl. One of the patients had a platelet count of 2000/mm³. Whereas one patient had a fibrinogen level of 30mg% on admission. This shows the extreme of the cases catered by the institute. All of the above-mentioned cases survived. In 20 cases the onset of obstetric haemorrhage was within the hospital during vaginal delivery, LSCS or laparotomy so MTP could be initiated immediately. Such cases included both elective and emergency situations. However in maximum number of cases due to delayed referral MTP could be initiated in 2-6 hrs (41 out of 81 cases).The total number of blood products given to 81

patients in the study were 1866, out of which 388 were PCV, 402 were FFP, 317 were PC and 736 were CP. The average number of PCVs, FFPs, PCs and CPs was thereby 4.8, 5, 4 and 9 respectively; this gave us a ratio of 1:1.02:0.8:2 of PCV: FFP: PC: CP transfusion, which means for every 50 units of PCV transfusion, 51 units of FFP, 40 unit of PC and 100 units of CP was required.

In the majority of cases, specifically 31 (38.2%), massive bleeding was successfully managed through the timely administration of oxytocics including oxytocin, methylergometrin, carboprost, carbetocin and misoprostol and the implementation of a massive blood transfusion protocol. This approach was crucial in preventing complications such as hypothermia, acidosis, and disseminated intravascular coagulation (DIC). As per Fig. 1 when surgical interventions were necessary, obstetric hysterectomy was the most commonly performed procedure, with 16 total and 3 subtotal hysterectomies. Additionally, cervicovaginal exploration and intrauterine packing were used in 13.6% of the cases (11 cases). Among the 81 cases reviewed, the SR cannula—a novel technique for managing atonic postpartum hemorrhage (PPH)—was employed in 5 cases (6.2%).

Table 3. On admission laboratory parameters, interval between onset of moh to mtp and total blood products given

Lab Parameters	Range	Mean	Median
HEMOGLOBIN (GM/DL)	1.8-13.9	6.5	6.5
PLATLET (/MM3)	2000-3,98,000	139	136
PT	9.3-90	22.78	13.7
INR	0.79-9	1.93	1.17
APTT	14.1-135	41	28.8
FIBRINOGEN	30-580	212	188
CREATININE	0.24-4.7	0.98	0.71
K+	3-6.8	4.15	4
Interval between Onset of Moh to Decision of Initiation of MTP			
Time Interval (HRS)	No. of Patients	Percentage	
0 HRS	20	24.7	
<2HRS	6	7.4	
2-6HRS	41	50.6	
6-12 HRS	10	12.3	
>12 HRS	4	4.9	
Blood Products			
	Total	Average	
PCV	388	4.8	
FFP	402	5	
PC	317	4	
CP	736	9	

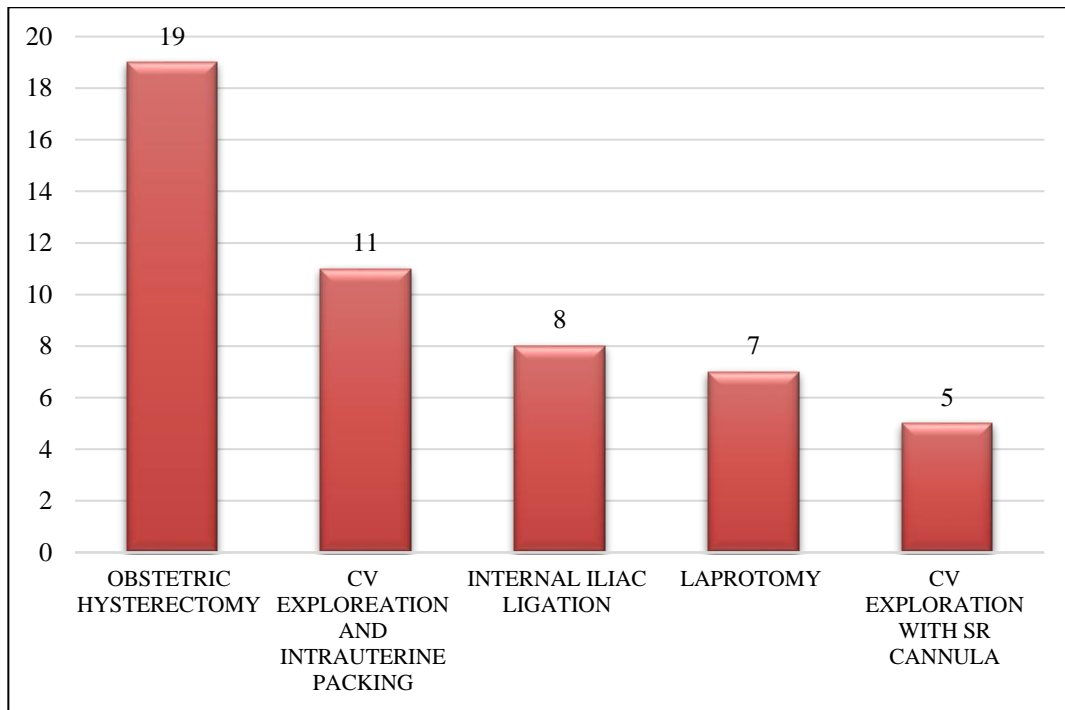


Fig. 1. Surgical interventions

Table 4. Specific complications after massive blood transfusion

Complications	No. of Cases	Percentage
TACO	13	16
TRALI	6	7.4
Electrolyte Imbalance	2	2.4

Many of the complications are overlapping with patients showing more than 2 complications at a time. Table 4 shows among those who developed complications, 13 patients (16%) had transfusion-associated circulatory overload (TACO), and 6 patients (7.4%) had transfusion-related acute lung injury (TRALI). Fluid management was closely monitored using the inferior vena cava (IVC) status via bedside echocardiogram in the ICU. It is noteworthy that 46 patients (56.8%) had no complications.

Multiple organ dysfunction syndrome (MODS) was diagnosed in 41 patients likely due to the severity of their preexisting conditions like preeclampsia, eclampsia, HELLP syndrome, hepatic disorders of pregnancy and PPCM; they may act as a confounder so we cannot say it was because of MT. Acute kidney injury (AKI) occurred in 6.2% of cases, primarily due to acute blood loss and delayed referral; 33% of these AKI cases progressed to chronic renal failure (CRF) requiring dialysis. Sepsis was observed in 7 patients (8.6%) by the end of the massive transfusion (MT), resulting from either the

progression of existing sepsis or secondary infections related to prolonged hospital stays and invasive treatments like ventilator-associated pneumonia (VAP). About 20% of the cases involved acute febrile reactions, managed with antipyretics and antihistamines. The incidence of dilutional coagulopathy decreased with the administration of blood products in a ratio of 1:1.02:0.8 for packed cell volume (PCV), fresh frozen plasma (FFP), and platelet concentrates (PC). Most post-MT complications were overlapping and related to the patients' underlying comorbidities.

Table 5 shows the relation to the chances of development of transfusion-related complications after specific blood product transfusion which was adjusted to on-admission maternal high-risk factors like hypertensive disorders of pregnancy, sepsis, placenta accreta spectrum, previous caesarean status, abruption and HELLP syndrome. The maximum chances of blood complications occur due to PC transfusion. It also depicts that the maximum chances of TRALI and TACO are with PC.

Table 5. Correlation of the blood products to complications

Relation of Blood Products to Complications Adjusted with High-Risk Maternal Factors on Admission				
	p-value	OR with 95% CI	95% C.I.for OR	
			Lower	Upper
RCC	0.371	1.235	0.777	1.962
FFP	0.772	1.030	0.829	1.287
PC	0.097	1.264	0.967	1.498
CP	0.539	1.033	0.936	1.134
Direct Relation of Specific Blood Product to Trali (Transfusion-Related Acute Lung Injury) and Taco (Transfusion Associated Circulatory Overload)				
RCC	0.771	1.105	0.565	2.160
FFP	0.170	1.303	0.893	1.901
PC	0.080	1.351	0.964	1.892
CP	0.652	1.047	0.857	1.279

Table 6. Comparison of ratios between different studies

Studies	Year	Cases	Protocol
Bonnet MP et al. [8]	2011	38	FFP/RBC ratio exceeds 1 at 12 h following the onset of obstetric haemorrhage.
Matsunaga S et al. [9]	2012	196	The medically necessary FFP/RCC ratio is 1.3 in obstetric haemorrhage.
Gutierrez MC et al. [10]	2012	26	MTP was defined as a combination of 6 units of O-negative RBC, 4 units of FFP (liquid AB plasma or thawed type-specific plasma), and 1 apheresis platelet (PLT) unit.
Green L et al. [11]	2016	181	FFP/RBC ratio ≥ 1 is required during massive obstetrics haemorrhage.
Tanaka H et al. [12]	2016	52	Transfusion of FFP/RBC ratio ≥ 1 reduces mortality

The average number of days for total hospital stay that the patients requiring massive blood transfusion was 8.9 days. In 95% of the cases patients required intensive care monitoring as well in ICU. Average days for ICU admission were 3.6 days. Many of the patients required mechanical ventilation. Average number of days required for mechanical ventilation was on an average 1.9 days.

Out of the 81 cases, 95% were successfully treated due to the timely administration of oxytocics, higher antibiotics, and the effective support of a well-equipped blood bank and laboratory. This was possible due to multidisciplinary approach and efficient work of obstetricians, anesthesia, intensive care team and blood bank. While maintaining proper aseptic precautions thus minimizing hospital acquired infections. In 4 (5%) cases, patients succumbed to irreversible shock and could not be salvaged. In 60% of the cases baby could be saved due to prompt decision of LSCS. While 31% of the cases were IUFD more commonly due to APH – abruption more than placenta previa. 9% of the abortal

cases include ectopic pregnancy, abdominal pregnancy and incomplete/ missed abortion with PAS.

Table 6 compares the ratio of fresh frozen plasma (FFP) to red blood cells (RBC) across various studies. In present study, the ratio of FFP to packed cell volume (PCV), platelet concentrates (PC), and cryoprecipitate (CP) was 1.02:1.0:8:2. Several retrospective studies have indicated that a higher plasma-to-RBC ratio in massive transfusion (MT) is linked to improved survival rates in patients with traumatic injuries. Since 2007, there has been a growing adoption of higher plasma-to-platelet-to-RBC ratios in MT therapy.

4. DISCUSSION

Trauma and obstetric patients have markedly different physiological profiles, which affect hemorrhage management strategies. During pregnancy, physiological changes such as hemodilution and increased cardiac output can mask significant bleeding, delaying detection until hemoglobin and hematocrit levels drop

significantly. Additionally, pregnancy-related comorbidities can elevate the risk of severe bleeding, consumption coagulopathy, and the early onset of organ failure and multiple organ dysfunction syndrome (MODS). These factors necessitate distinct approaches to managing hemorrhage in obstetric cases compared to trauma patients.

According to the RCOG's "Green-top Guideline: Blood Transfusion in Obstetrics" (October 2006), the recommended dosage for fresh frozen plasma (FFP) is 12-15 ml/kg for every 6 units of red blood cells (RBC). The guideline emphasizes using prothrombin time (PT) and activated partial thromboplastin time (APTT) from coagulation tests as the primary indicators for determining FFP requirements, with target levels set at 1.5 times the normal range for PT and APTT, and a fibrinogen level of 150 mg/dl or higher. It also advises regular monitoring of these tests and blood counts in cases of prolonged bleeding. Additionally, the guideline recommends administering cryoprecipitate in two sets of five units to maintain fibrinogen levels at or above 150 mg/dl.

The ACOG (May 2015) guidelines recommend an early and aggressive transfusion strategy with a 1:1:1 ratio of red blood cells (RBC), fresh frozen plasma (FFP), and platelets (PC) during massive transfusions. This approach aims to address coagulopathy, hypothermia, and acidosis, which are critical factors that significantly increase the risk of patient mortality [13].

Poor outcomes following massive obstetric hemorrhage (MOH) and massive transfusion (MT) are often due to delayed treatment, unavailability of blood products, inaccurate blood loss estimation, lack of treatment protocols, and poor communication among team members. In contrast, our study highlights that present institute has a benefit of a well-organized multidisciplinary approach. This includes an equipped trauma care center, dedicated obstetric team, efficient laboratory and blood bank services, anesthesia, operating theaters, and a critical care team with ICU centers and dialysis units, all operating 24/7.

In the current study, 81 out of 2,956 obstetric admissions required massive blood transfusion (MT), resulting in a utilization rate of 2.74%. This is higher than the 0.7% reported by Ochiai D et al. in Japan [14] but lower than the 3% observed

by Paul I. Ramler et al. in the Netherlands [15]. Notably, a 2020 study by S. Anuraga et al. in Puducherry [16], India, reported a much higher MT rate of 20%. The previous study at the same institute by Rumi Bhattacharjee et al. [17] in 2017 had an MT rate of 2.4%. Regarding case distribution, the prior study showed 60.9% antenatal and 39.1% postnatal cases, while the current study reports 58% antenatal and 42% postnatal cases. This slight shift, along with varying MT rates, suggests evolving practices or patient profiles in obstetric care.

In the current study, the mean age of patients requiring massive blood transfusion (MT) was 27.3 years, with a range of 19 to 38 years. This is consistent with the mean age reported in Puducherry, India, but lower than the 36.8 years observed in Japan and the 32 years in the Netherlands. The previous study at the same institute noted that most patients were in the 21-30 years age group. These findings suggest that younger women are more commonly affected in Asian countries, potentially due to earlier marriages and childbearing compared to developed countries. Supporting this, Patricia et al. found that females under 20 years are more susceptible to pregnancy complications, such as poor fetal growth and postpartum hemorrhage (PPH). Additionally, a study on elderly primigravidas indicated that 3% experienced antepartum hemorrhage and 3% had PPH, further highlighting the impact of age on pregnancy outcomes.

As per the Quality Improvement Program survey conducted in American College of surgeons, 2013 during the pre-hospital resuscitation the most common blood products used were RBCs and plasma, while the most common intravenous hemostatic agent is Tranexamic Acid. Hypotension with SBP \leq 100mmhg was the most common MTP trigger. Laboratory values were infrequently used to initiate MT. Amongst the blood products plasma is immediately available in <5 minutes. most common plasma type used is thawed plasma. The most common FFP:RBC ratio in the first cooler is \geq 1. Use of cryoprecipitates is also encouraged in MT. In the present institute the shock status of the patients was assessed by shock index of the patient (Pulse/ SBP), urine output and by calculating the average amount of blood loss of the patient. In the present institute the average time interval between the blood collection and issuing of the first blood product is

12 min. The most common hemostatic agent used is injectable TRANEXAMIC ACID followed by injectable HEMOCOAGULASE (BOTROPACE). The facility of TEG (thromboelastography) is not currently available in present institute.

Le Bas et al. [18] recently highlighted that during pregnancy, the normal shock index (pulse rate divided by systolic blood pressure) is typically higher compared to non-pregnant adults. This increase is due to a higher pulse rate and a decrease in systolic blood pressure. An obstetric shock index (OSI) greater than 1 is associated with a higher likelihood of requiring a blood transfusion following postpartum hemorrhage (PPH). Consequently, the OSI can serve as a useful bedside clinical tool for assessing the degree of blood loss, offering a more reliable measure than visual estimation, which is prone to significant observer variability. The latest Green Top Guideline from the Royal College of Obstetricians and Gynaecologists underscores the importance of the OSI in identifying women at risk of adverse outcomes. In the present study, 26% of patients were categorized with moderate shock (SI between 1 and 1.39), while another 26% were in severe shock (SI ≥ 1.4). On average, patients with severe shock experienced blood loss exceeding 30% of their total blood volume. As compared to Rumi et al which was a retrospective study, blood products transfusion was decided by physician whereas in the current study which is a prospective study the determinants for blood transfusion were the clinical status of the patient and lab parameters including the Shock Index(SI), coagulation profile and obstetric parameters.

As per literature, the proportion of patients with previous caesarean sections varies, with Ramer et al. reporting 23%, S. Anuraga et al. indicating 33.3%, and the current study showing a rate of 31%. A history of caesarean sections, along with prior myomectomy or dilatation and evacuation, is linked to increased risks in subsequent pregnancies. These risks include uterine rupture, dense adhesions, placenta previa, and placenta accreta syndrome. Notably, the risk of placenta accreta syndrome rises with the number of previous caesarean sections, thereby increasing the likelihood that a patient may require an obstetric hysterectomy in future pregnancies.

Among the causes of obstetric hemorrhage, the single most common cause is post-partum

hemorrhage – which includes all the 4Ts - tone, thrombin, tissue and trauma. In the present study (year 2022 to 2024) atonic PPH accounts for 50.6% of the cases while it was 33.5% in previous study of the same institute (year 2014 to 2017 published in 2019). It was 58.3 % in the study conducted in Japan and 57% in Netherlands study.

In the present study, peripartum hysterectomy was necessary in 23.4% of cases involving obstetric hemorrhage. The primary causes for this intervention were atonic postpartum hemorrhage (PPH), traumatic PPH, followed by placenta accreta syndrome and uterine rupture. This rate is comparable to the 23% reported by Rumi Bhattacharjee et al. (2017) but higher than the 11% observed in a study conducted in Puducherry. In contrast, rates in developed countries vary significantly, with Ochiai D et al. reporting 4.2% and Paul I. Ramler et al. reporting 30%. Peripartum hysterectomy is generally considered a last-resort surgical intervention, employed when patients do not respond to aggressive medical management and conservative organ-preserving techniques.

Mortality rates following peripartum hysterectomy differ markedly between developed and developing countries. In developed nations, such as the Netherlands, mortality is low at 1.08%. However, in developing countries, rates are significantly higher, with 11.1% in Puducherry, 10% in a previous study from the current institute, and 5% in the present study. Common causes of mortality include MODS, ARDS, sepsis, and acute kidney failure. Research suggests that early activation of massive transfusion protocols (MTP) and improved antenatal care to identify high-risk patients can help prevent severe outcomes and reduce mortality

In the current study, among 81 cases, 46 (56.8%) did not experience complications related to massive blood transfusion. This favorable outcome is attributed to effective fluid management, monitored through the patient's IVC status using a bedside ECHO machine in the ICU. Among the complications, Multiple Organ Dysfunction Syndrome (MODS) was the most common, likely due to the underlying pathophysiology of the patients. Transfusion-related acute lung injury (TRALI) was observed in 7.4% of cases, while transfusion-associated circulatory overload (TACO) occurred in 16%.

The risk of TRALI varies by blood product, with one case per 5,000 PRBCs, one per 2,000 FFP, and one per 400 platelets [19-21]. A recent ICU study found an 8% incidence of TRALI, with platelet or FFP transfusions increasing the risk nearly threefold. In present study, the odds ratios for developing TRALI were 1.351 for platelet transfusions, 1.303 for FFP, 1.105 for PRBCs, and 1.047 for cryoprecipitate. Dilutional coagulopathy was observed in only 16% of cases, as the PCV:FFP ratio is maintained at 1:1:1. Fibrinogen replacement, through cryoprecipitate or fibrinogen concentrate, has proven effective in managing obstetric hemorrhage and other conditions, with several studies supporting its efficacy. As per a study conducted in Department of Obstetrics and Gynecology, Baylor College of Medicine, Houston, Texas between 2014-2020 uterine atony was found to be the commonest etiology for massive hemorrhage (34%), followed by placenta accreta spectrum (32%). A mean of 6.5 units of packed red blood cells, 14.8 units of fresh frozen plasma and cryoprecipitate, and 8.3 units of platelets were transfused per patient [22].

5. CONCLUSION

In the present study, the utilization rate of the massive transfusion protocol (MTP) was 2.74%. Postpartum hemorrhage (PPH) emerged as the primary cause of obstetric hemorrhage. To enhance maternal morbidity and mortality outcomes, early referral to tertiary care centers, timely administration of oxytocics, and the prompt initiation of MTP and surgical interventions are crucial in managing cases of uncontrolled bleeding.

Maintaining a blood product ratio of 1:1.02:0.8 for PCV:FFP:Platelets proved effective in preventing coagulopathy, ensuring adequate tissue perfusion, and protecting patients from the detrimental cycle of sepsis, hypothermia, hemodilution, and shock. MT related complications were managed by assessing fluid status through chest auscultation and bedside ECHO for IVC status in the ICU, which provided valuable guidance in managing these complications.

Additionally, slightly increasing the proportion of FFP transfusion relative to PCV, as determined by interval blood testing, can support early hemostasis. This approach helps in optimizing

the balance of blood products and improving overall patient outcomes during massive transfusions. All the facilities serving obstetric patients must have an established protocol so that prompt treatment of obstetric patients can improve their survival.

DISCLAIMER (ARTIFICIAL INTELLIGENCE)

Author(s) hereby declare that NO generative AI technologies such as Large Language Models (ChatGPT, COPILOT, etc) and text-to-image generators have been used during writing or editing of manuscripts.

CONSENT

This was an observational study. data of the patients were collected from the file of the patients.

ETHICAL APPROVAL

The data collection was started after approval of the institutional ethics committee-IEC/BU/136/Faculty/1/293/2022, as it was a prospective and descriptive study, a waiver of consent was requested.

ACKNOWLEDGEMENTS

We express our sincere thanks to the Dean of Pramukhswami Medical College, Superintendent of Shree Krishna Hospital, Karamsad and Provost of Bhaikaka University, Karamsad for pratonage, permission and provision of all infrastructure facilities to undertake this study and for letting me use patient details.

COMPETING INTERESTS

Authors have declared that no competing interests exist.

REFERENCES

1. Fabry SM Sheldon GF, et al. Massive transfusion in the surgical patient; American association of blood bank 1994; 135-156.
2. Balasudha K panikar S massive obstetric haemorrhage; Our experiences,

- International Journal of Reproduction, Contraception, Obstetric and gynaecology 2014 March;3(1).
3. Gungar T Smisek A et al. Surgical treatment of intractable postpartum haemorrhage and changing trends in medium Obstetric perspective. Arch Gynaecology obstetric. 2009;280;351-355 DOI:10.1007 / s00404-008-0914%.
 4. Ickx BE, Fluid and blood transfusion management in obstetrics. Eu Anaesthesiology. 2010;27:1031-5.
 5. Brohi K, Cohen MJ, Davenport RA. Acute coagulopathy of trauma: mechanism, identification and effect. Current opinion in critical care. 2007 Dec 1;13(6):680-5.
 6. Lloyed Ketcham, John R Hess Seppo *Hippalen indicacious* for early fresh frozen plasma, cryoprecipitates and Platelet transfusion in trauma; The journal of trauma.01/7 2006;60(6supple):551-8.
 7. Hiroaki Tanaka, Shigetaka Matsunga, Tomoyuki Yamashita et al. A systemic review of MTP in obstetrics. Taiwanese Journal of obstetrics and gynaecology. Sb. 2017;715-718.
 8. Bonnet MP, Deneux-Tharoux C, Bouvier-Colle MH. Critical care and transfusion management in maternal deaths from postpartum haemorrhage, Eur JObstet Gynecol Reprod Biol. 2011;158:183-188
 9. Matsunaga S, Seki H, Ono Y, Matsumura H, Murayama Y, Takai Y, et al. A retrospective analysis of transfusion management for obstetric hemorrhage in a Japanese obstetric center, ISRN Obstet Gynecol. 2012;2012:854064
 10. Gutierrez MC, Goodnough LT, Druzin M, Butwick AJ. Postpartum hemorrhage treated with a massive transfusion protocol at a tertiary obstetric center: a retrospective study, Int J Obstet Anesth, 21 (2012), pp. 230-235
 11. Green L, Knight M, Seeney F, Hopkinson C, Collins PW, Collis RE, et al. The haematological features and transfusion management of women who required massive transfusion for major obstetric haemorrhage in the UK: a population based study, Br J Haematol. 2016;172:616-624
 12. Tanaka H, Katsuragi S, Ikeda T, Osato K, Hasegawa J, Nakata M, et al. Efficacy of transfusion with fresh-frozen plasma:red blood cell concentrate ratio of 1 or more for amniotic fluid embolism with coagulopathy: af case-control study, Transfusion. 2016;56:3042-3046
 13. Cotton BA, Guy JS, Morris Jr JA, Abumrad NN. The cellular, metabolic, and systemic consequences of aggressive fluid resuscitation strategies. Shock. 2006;26:115e21.
 14. Ochiai D, Abe Y, Yamazaki R, Uemura T, Toriumi A, Matsunashi H, Tanaka Y, Ikenoue S, Kasuga Y, Tanosaki R, et al. Clinical Results of a Massive Blood Transfusion Protocol for Postpartum Hemorrhage in a University Hospital in Japan: A Retrospective Study. Medicina 2021;57:983. Available:https://doi.org/10.3390 /medicina57090983
 15. Ramler PI, van den Akker T, Henriquez DDCA, et al. Women receiving massive transfusion due to postpartum hemorrhage: A comparison over time between two nationwide cohort studies. Acta Obstet Gynecol Scand. 2019;98(6):795-804. DOI:10.1111/aogs.13542
 16. Anuragaa S, Chaturvedula L, Basavarajegowda A. Blood component therapy in patients having massive obstetric hemorrhage in a tertiary care center in Puducherry. Asian J Transfus Sci 2023;17:210-6.
 17. Bhattacharjee R, Raithatha N, Sapre S. An Analysis of Large Volume Blood and Blood Product Transfusion in Critically Ill Obstetric Patients: A Retrospective Study. J South Asian Feder Obst Gynae 2019;11(3):148–152
 18. Le Bas A, Chandharan E, Addei A, et al. Use of the “obstetric shock index” as an adjunct in identifying significant blood loss in patients with massive postpartum hemorrhage. Int J Gynecol Obstet 2013;124(3):253–255. DOI: 10.1016/j.ijgo.2013.08.020.
 19. Gazmuri RJ, Shakeri SA. Blood transfusion and the risk of nosocomial infection: an underreported complication? Crit Care Med. 2002;30(10):2389 - 2391.
 20. Silliman CC, Paterson AJ, Dickey WO, et al. The association of biologically active lipids with the development of transfusion-related acute lung injury: a retrospective study. Transfusion. 1997; 37(7):719-726.
 21. Wallis JP, Lubenko A, Wells AW, Chapman CE. Single hospital experience of TRALI. Transfusion. 2003; 43 (8): 1053-1059.

22. Salmanian B, Clark SL, Hui SR, et al. Massive Transfusion Protocols in Obstetric Hemorrhage: Theory versus Reality. Am J Perinatol. 2023;40(1): 95-98.
DOI:10.1055/s-0041-1728833

Disclaimer/Publisher's Note: The statements, opinions and data contained in all publications are solely those of the individual author(s) and contributor(s) and not of the publisher and/or the editor(s). This publisher and/or the editor(s) disclaim responsibility for any injury to people or property resulting from any ideas, methods, instructions or products referred to in the content.

© Copyright (2024): Author(s). The licensee is the journal publisher. This is an Open Access article distributed under the terms of the Creative Commons Attribution License (<http://creativecommons.org/licenses/by/4.0>), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Peer-review history:
The peer review history for this paper can be accessed here:
<https://www.sdiarticle5.com/review-history/122170>