

# HEMOSTATIC PARAMETERS IN WOMEN WITH OBSTETRIC HEMORRHAGE

## ABSTRACT

**Aims:** To find out the relationships observed in hemostatic parameters during the course of obstetric hemorrhage for an effective hemostatic management and goal directed therapy.

**Study design:** Analytical, Case-Control Study.

**Place and Duration of Study:** Department of Obstetric and Gynaecology, Department of Pathology, Govt Medical College, Kottayam, Duration: 12 months.

**Methodology:** We included 68 cases (18-45 yrs) with obstetric hemorrhage and 68 controls (18-45 yrs, Third trimester) without Obstetric hemorrhage. Blood collected for PT, APTT, Fibrinogen and platelet count in sodium citrate and EDTA anticoagulated samples. PT, APTT, Fibrinogen assay done by Automatic methods, platelet count done in 5 part coulter machine.

**Results:** The study, involving 136 participants divided into case and control groups, revealed significant differences in key haemostatic parameters. The case group exhibited elevated Prothrombin Time (PT) and Activated Partial Thromboplastin Time (APTT) values, along with lower Platelet count and Fibrinogen concentration compared to the control group. Notably, Antepartum Hemorrhage (APH) and Postpartum Hemorrhage (PPH) showed a substantial linear relationship with platelet levels in the case group, emphasizing the potential predictive role of platelet counts in hemorrhagic events associated with obstetric conditions.

**Conclusion:** Study sheds light on the critical link between obstetric hemorrhage and hemostatic parameters, highlighting significant variations in prothrombin time (PT), activated partial thromboplastin time (APTT), platelet count, and fibrinogen concentration between women with and without obstetric hemorrhage. The findings underscore the complexity of managing obstetric hemorrhages and the importance of laboratory-based screening in evaluating the coagulation status of affected patients.

*Keywords: Maternal hemorrhage, PT, aPTT, Fibrinogen assay, Antepartum hemorrhage, Post partum Hemorrhage*

## 1. INTRODUCTION

Obstetric haemorrhage (OBH) is widely acknowledged as a prominent factor contributing to maternal morbidity and mortality on a global scale. A significant majority of fatalities occur in countries with limited resources. Nevertheless, Major Obstetric haemorrhage (MOH) leads to profound morbidity across diverse healthcare settings, manifesting as multiorgan failure, postpartum hysterectomy, and enduring psychological trauma. These outcomes contribute substantially to extensive economic and social burdens. Severe maternal morbidity linked to haemorrhage surpasses the morbidity rates associated with other obstetric and medical conditions.

Safety bundles are crafted to offer structured, evidence-based protocols that, when implemented, consistently enhance patient outcomes. Standardized obstetric haemorrhage (OBH) bundles, aiding healthcare teams in methodically preparing for, recognizing, and responding to OBH, have shown a notable decrease in patient morbidity and mortality. The incidence of postpartum haemorrhage (PPH) has risen in numerous countries<sup>9</sup>. Despite alterations in obstetric strategies, the rate of PPH exceeding 2500 ml in Scottish maternal morbidity reports has remained relatively constant at approximately 5.8 per 1000 from 2009 to 2012.

Monitoring major obstetric haemorrhage at the hospital level poses a considerable operational challenge. Criteria for tracking severe morbidity, potentially encompassing haemorrhage-related incidents, involve the examination of obstetric hospitalizations of women receiving 4 or more units of red blood cells (RBCs) or necessitating admission to the intensive care unit.

However, such events are infrequent, occurring in approximately 3 out of 1,000 births.

Utilizing data from biomedical informatics, acquired through the interrogation of electronic health records (EHR), can serve as a method for characterizing and monitoring postpartum haemorrhage. Biomedical informatics, as articulated by the American Medical Informatics Association, is an interdisciplinary field dedicated to the efficient utilization of biomedical data, information, and knowledge for scientific exploration, problem resolution, and clinical decision-making, all aimed at enhancing human health.

Detailed clinical data, obtained directly from the EHR, encompass granular information such as laboratory results, details on drug and device usage, administration of blood products, recorded procedures, and various clinical particulars. This highly detailed data is well-suited for comprehensive and multifaceted measurement of postpartum haemorrhage, enabling an in-depth assessment of clinical management and associated outcomes at both the hospital and hospital-system levels.

Postpartum hemorrhage (PPH) arises due to diverse obstetric complications, including uterine atony, placenta previa, genital tract trauma, or abruption. It is more frequently observed in conjunction with instrumental delivery, induced labor, and cesarean delivery.

The most recent obstetric management guideline in the UK prioritizes surgical intervention with an early return to the operating theatre and a systematic approach to using uterotonics when uterine atony is identified as a cause of PPH. The guideline underscores the importance of early cross-matching of blood, collaboration with hematology services, and suggests maintaining a hemoglobin level of about 8 g/dL through red blood cell (RBC) transfusions. It recommends using fresh frozen plasma (FFP) to keep the prothrombin time (PT) and activated partial thromboplastin time (aPTT) within 1.5 times the normal range, with fibrinogen levels above 1 g/L and platelet count above  $50 \times 10^9/L$ . These recommendations align with transfusion policies for routine coagulation tests and fibrinogen levels in the non-pregnant population. Research endeavors are essential to establish the appropriate reference values in this context. Enhancing awareness of the accurate application and interpretation of coagulation monitoring is crucial for effective emergency diagnostics.

### **Review of Literature:**

Pregnancy is linked to alterations in hemostasis, characterized by an elevation in most clotting factors, a decrease in the levels of natural anticoagulants, and a reduction in fibrinolytic activity. During a normal pregnancy, the platelet count experiences a decrease,

potentially attributed to heightened destruction and hemodilution, with the most significant reduction observed in the third trimester.

With the increase in most coagulation factors during normal pregnancy, the prothrombin time (PT) and activated partial thromboplastin time (APTT) may exhibit a shortened duration. The PT, along with its derived measurement, the international normalized ratio (INR), assesses factors such as coagulation factors II, V, VII, X, and fibrinogen. Meanwhile, the APTT serves as a reliable screening test for deficiencies in coagulation factors VIII, IX, XI, and XII. Laboratory-based screening, including platelet count, PT, APTT, D-Dimer, and plasma fibrinogen levels, is routinely employed to evaluate the coagulation status in obstetric patients. The platelet count offers a measurement of platelet concentration, focusing on quantity rather than function. Prothrombin time (PT) gauges the extrinsic and common coagulation pathways, responding to the levels of coagulation factors II, V, VII, and X. On the other hand, activated partial thromboplastin time (APTT) evaluates coagulation through the intrinsic and common pathways, being sensitive to all coagulation factors except for FVII and FXIII.

### **Hemostatic changes in normal pregnancy:**

A typical pregnancy is characterized by significant alterations in various aspects of hemostasis. These changes collectively contribute to sustaining placental function throughout pregnancy and preventing excessive bleeding during delivery. The majority of shifts in blood coagulation and fibrinolysis result in a heightened state of coagulability. This phenomenon serves as a protective mechanism, shielding the woman from hemorrhage during delivery. However, it also renders her susceptible to thromboembolism, both during pregnancy and in the postpartum period. The alterations in the coagulation system observed in a typical pregnancy align with an ongoing low-grade process of intravascular coagulation.

**a. Platelets:** "Thrombocytopenia is a prevalent haemostatic abnormality in pregnancy, affecting approximately 10% of healthy women in late pregnancy. This condition is, at least in part, attributed to haemodilution. However, the rise in mean platelet volume indicates a compensated state involving progressive platelet destruction. Further evidence of in vivo platelet activation during late pregnancy is evident through the increased concentrations of  $\beta$ -thromboglobulin and thromboxane A<sub>2</sub> derivatives". [108]

**b. Coagulation system:** Throughout pregnancy, there is a substantial increase in the concentrations of coagulation factors VII, VIII, IX, X, XII, and von Willebrand factor, along with a notable rise in plasma fibrinogen concentration. In late pregnancy, plasma fibrinogen levels often exceed 600 mg/dL. Factor VII may experience up to a tenfold increase during pregnancy. Von Willebrand factor and factor VIII are elevated in late pregnancy, resulting in approximately double the coagulation activity observed in the non-pregnant state. Conversely, factor IX undergoes a small increase, while factor XI experiences a minor decrease. After an initial rise, factor XIII gradually decreases, reaching 50% of the normal non-pregnant value at term. Factors II and V, however, do not exhibit significant changes during pregnancy.

"Controversial findings have been reported regarding changes in antithrombin (AT) during pregnancy, but AT often remains within the normal range. Protein C activity seems unaffected by gestation, while protein C antigen levels tend to increase in the second trimester, staying within the normal non-pregnant range. Neutrophil activation triggers endothelial thrombomodulin (TM) proteolysis, increasing TM plasma levels in the third trimester. Some studies report a continuous increase in TM levels during pregnancy, rapidly decreasing post-partum. The physiological decrease in free protein S levels during

pregnancy raises questions about its contribution to the hypercoagulable state and increased thromboembolism risk. Total protein S progressively decreases during gestation, with values going below the normal range early on. The decline in protein S during the first weeks of pregnancy does not allow for a diagnosis of inherited protein S deficiency. Attempts to establish normal protein S levels during pregnancy are not recommended. Heparin cofactor II, another natural coagulation inhibitor, has been reported to increase in plasma during physiological pregnancy". [108]

"Protein Z, a vitamin-K-dependent plasma glycoprotein, acts as a cofactor to a plasma proteinase inhibitor, inhibiting the activation of factor X. Deficiency in Protein Z has been reported in women experiencing unexplained early fetal losses, and the presence of antibodies to protein Z can contribute to adverse pregnancy outcomes. Recent data indicate a gradual rise in protein Z levels with gestational age in normal pregnancies, returning to normal levels approximately 6 to 12 weeks postpartum. This normal increase in protein Z during pregnancy may counterbalance the rise in clotting factors, offering protection against thrombosis in pregnant women. During pregnancy, there is a reduction in activated protein C (APC) sensitivity. At term, approximately 45% of pregnant women exhibit an APC sensitivity ratio below the 5th percentile of the normal range for non-pregnant women of similar age. The decrease in APC ratio is directly correlated to its value in the non-pregnant state, with the most pronounced effect observed in women with the highest APC ratio. Around 50% of healthy women develop APC resistance, reaching its lowest point in the second trimester of pregnancy, with minimal subsequent change". [108]

**c. Fibrinolysis:** Fibrinolytic activity in plasma is diminished during pregnancy, remaining low throughout labor and delivery, and returning to normal shortly after placental delivery. The activity of tissue plasminogen activator (t-PA) decreases during pregnancy, attributed not only to the gradual rise in plasminogen activator inhibitor-1 (PAI-1) but also to increasing levels of plasminogen activator inhibitor-2 (PAI-2). PAI-1 values increase during pregnancy and return to normal levels approximately 5 weeks post-partum.

"Plasminogen activator inhibitor-2 (PAI-2) typically becomes detectable in plasma only in pregnant individuals. Its source, villous cells, suggests that changes in placental tissue quantity may influence plasma levels, establishing a positive correlation between PAI-2 concentrations and placental weights. PAI-2 concentrations vary with birth weights, indicating a dependency on both the quantity and quality of placental tissues and fetal growth. Despite elevated levels of PAI-1 and PAI-2, a highly significant positive correlation has been noted between gestational age and D-dimer concentration. This increase in D-dimer complicates its use in excluding venous thromboembolism in pregnant patients with clinical suspicion. Efforts have been made to establish specific ranges of D-dimer levels in pregnancy". [108]

#### **d. Changes in the postpartum after delivery:**

The surge in clotting activity during delivery is likely linked to the expulsion of the placenta and the release of thromboplastic substances at the separation site. Hemostatic changes during the puerperium mirror those observed after extensive surgery<sup>65,52</sup>. The mean platelet count slightly decreases at placental delivery and begins to rise on days 2–5 postpartum. In high-risk patients requiring postpartum thromboprophylaxis, consideration should be given to the reactive thrombocytosis postpartum following operative delivery.

Plasma antithrombin (AT) levels significantly increase for at least 2 weeks postpartum after normal delivery. Protein C levels rise immediately after delivery and remain elevated 3 days postpartum. Total and free protein S levels notably increase after delivery, with total protein

S normalizing in the first week postpartum, while free protein S may not normalize until 5 weeks postpartum. Approximately 15% of women still have levels below the reference range for non-pregnant women at 8 weeks postpartum. The prolonged time for free protein S to reach non-pregnant values should be considered when evaluating thrombotic risk.

Both thrombin-antithrombin complex (TAT) and prothrombin fragment 1+2 levels increase during and immediately after delivery. Three weeks post-delivery, blood coagulation and fibrinolysis generally normalize. Compensated, accelerated intravascular coagulation appears necessary for maintaining the uterine placental interface and preparing for the hemostatic challenge of delivery.

The peak in clotting and platelet activity occurs immediately after placental delivery, while the peak of fibrinolytic activity is observed during the first 3 hours postpartum, reflected by an increase in d-dimer levels.

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## **2. MATERIAL AND METHODS**

### **2.1 HYPOTHESIS STATEMENT:**

It can be hypothesized that there is significant effect of obstetric hemorrhage on the hemostatic parameters.

**2.2 TYPE OF STUDY:** Analytical, Case- Control study

**2.3 PERIOD OF STUDY:** 12 months

**2.4 STUDY SETTING:** Department of Pathology (Hematology division), Govt. Medical College, Kottayam

### **2.5 SAMPLE SIZE**

P0= Proportion of controls

P1=Proportion of cases

$$q = 1-p$$

= 1.96= Value of the standard normal distribution corresponding to a significance level of  $\alpha$ (1.96 for a 2 sided test at the 0.05 level)

= 0.84= Value of the standard normal distribution corresponding to the desired level of power(0.84 for a power of 80 %)

$$n = 68$$

( Considering equal number of cases and controls,  $r = 1$ )

**2.6 SAMPLING METHOD:** Judgemental / Purposive sampling

**2.7 INCLUSION CRITERIA:** All women in the age group of 18-45yrs (primigravida and multigravida) with obstetric hemorrhage reported in Dept. of Obstetrics & Gynaecology, Govt. Medical College Kottayam.

### **2.8 EXCLUSION CRITERIA:**

- Pregnant women with known coagulopathies
- Pregnant women already on hemostatic or antiplatelet therapy.
- Pregnant women with fibroid uterus/ adenomyosis/inflammatory diseases
- Women with congenital hypofibrinogenemia
- Pregnant women with Gestational hypertension

### **2.9 LIMITATIONS EXPECTED:**

Sample size may not be attained

#### **2.10 PERSONNEL RESPONSIBLE FOR DATA COLLECTION:**

DR. JIJ I J S

#### **2.11 PERSONNEL RESPONSIBLE FOR DATA ANALYSIS:**

DR. JIJ I J S

#### **2.12 FUNDING AGENCY:**

NIL

#### **2.13 STUDY TOOLS**

1. Consent form
2. Equipment for blood collection, centrifugation
3. Anticoagulation bottles for sample collection
4. Coagulometer
5. Reagents for coagulation tests
6. Detailed proforma in each case

#### **2.14 STUDY PROCEDURE:**

After identifying the study population based on the inclusion and exclusion criteria and getting informed written consent, relevant blood investigation pertaining the hemostatic parameters is performed including PT, APTT, Platelet count and then more specific fibrinogen concentration is done.

Blood collection for PT, APTT and Fibrinogen concentration done by direct venipuncture in 3.2% buffered sodium citrate in a ration 9:1(9 part of whole blood and one part of anticoagulant)

For platelet count blood is collected in EDTA tube.

Total 6 ml of blood is needed from one particular case/control. PT, APTT and fibrinogen concentration done by automatic method. Platelet count is done by 5 part/3 part coulter machine

For PT, APTT and fibrinogen concentration a period of 1-2 Hrs is advocated to keep the sample in refrigerator at 4 degree celcius without error in the value and the blood for platelet count can store upto 4 hrs in refrigerator.

Data entry, documentation and significance change observed in maternal hemorrhage is analyses.

#### **2.15 QUALITY ASSURANCE:**

1. The blood sample would be collected under strict aseptic precaution and processed and stored
2. The equipment used for the study would be of standard quality
3. Reading would be obtained as per standardized

#### **2.16 DATA MANAGEMENT AND ANALYSIS**

The data will be entered in Microsoft excel and further statistical analysis will be done using SPSS software. Chi -square test formula would



### 3. RESULTS AND DISCUSSION

The collected data was analysed using IBM SPSS software ver.20

**NOTE:**

- **Kolmogorov-Smirnov test** was used to determine the normality of the data.
- Continuous variables are expressed in terms of **Mean, standard deviation (SD)**.
- Categorical variables are expressed in **frequency (n) and percentage (%)**
- **Chi-square or Fisher's exact test** was used to assess the significance between categorical variables.
- **Independent t test** was used to compare between two variables.
- **Pearson correlation test** was used to assess linear relationship between obstetric haemorrhage and haemostatic parameters like PT, APTT, Platelet count and fibrinogen concentration.
- **P < 0.05** was considered as statistically significant\*
- **There are two groups in this study,**

**a. Case:** Women in the age group 18-45yrs with obstetric haemorrhage

**b. Control:** Women in the age group 18-45yrs in their third trimester without obstetric haemorrhage

- There are totally 136 samples in this study (68 in each group).

**Table 1: Distribution of study participants according to the study group**

Group	FREQUENCY (N)	PERCENT (%)
CASES	68	50.0
CONTROL	68	50.0
TOTAL	136	100.0

**Interpretation:** There are two groups in this study. Case group and control group having 68 participants in each group. Totally there are 136 samples in the study.

**Table 2: Mean age distribution**

Group	MEAN	SD	T	P-VALUE
CASE (N=68)	29.88	4.49	3.2	0.002*
CONTROL (N=68)	27.24	5.13	3.2	

\*p is significant at <0.05

SD = Standard deviation

**Table 3: Comparison of haemostatic parameters between two groups**

Haemostatic	CASE (N=68)	CONTROL	T	P-VALUE
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parameter	(N=68)			
PT (SECONDS)	14.04 ± 0.86	12.3 ± 0.44	14.90	<0.0001*
APTT (SECONDS)	37.6 ± 1.68	29.87 ± 2.96	18.67	< 0.0001*
PLATELET COUNT (10 <sup>9</sup> /L)	1.32 ± 0.28	2.49 ± 0.64	-13.87	< 0.0001*
FIBRINOGEN CONCENTRATION (G/DL)	1.36 ± 0.44	3.71 ± 0.65	-24.65	< 0.0001*

PT – Prothrombin Time, APTT - Activated Partial Thromboplastin Time

\*P is significant at < 0.001

**Interpretation:** Four haemostatic parameters were considered in this study like PT, APTT, Platelet count and Fibrinogen concentration. The mean PT in case group is 14.04 ± 0.86 seconds, and in control group it is 12.3 ± 0.44 seconds. t(df) = 14.90 (134). There was statistically significant difference between two groups (p < 0.05), with mean value higher in case group.

The mean APTT in case group is 37.6 ± 1.68 seconds, and in control group it is 29.87 ± 2.96 seconds. t(df) = 18.67 (134). There was statistically significant difference between two groups (p < 0.05), with mean value higher in case group.

The mean Platelet count in case group is 1.32 ± 0.28 (10<sup>9</sup>/L), and in control group it is 2.49 ± 0.64 (10<sup>9</sup>/L). t(df) = -13.87 (134). There was statistically significant difference between two groups (p < 0.05), with mean value lower in case group.

The mean fibrinogen concentration in case group is  $1.36 \pm 0.44$  g/dl, and in control group it is  $3.71 \pm 0.65$  g/dl.  $t(df) = -24.65 (134)$ . There was statistically significant difference between two groups ( $p < 0.05$ ), with mean value lower in case group.

**Table 4: Pearson correlation test between hemostatic parameters and obstetric hemorrhage in case group**

Variables	r-coefficient	p-value
APH and PT	-0.033	0.789
APH and APTT	-0.081	0.511

<b>APH and Platelet count</b>	-0.285	0.018*
<b>APH and fibrinogen concentration</b>	0.231	0.056
<b>PPH and PT</b>	0.033	0.789
<b>PPH and APTT</b>	0.081	0.511
<b>PPH and Platelet count</b>	0.285	0.018*
<b>PPH and fibrinogen concentration</b>	-0.231	0.056

**\*P is significant at < 0.05**

**Interpretation:** There is significant linear relationship between APH and PPH with platelet levels. There was no significant correlation between other haemostatic parameters and APH, PPH. There found to be negative correlation between APH and platelet count and positive correlation between PPH and platelet count.

#### **DISCUSSION:**

Maternal hemorrhage poses a significant threat to maternal health globally, particularly in developing nations, contributing to high rates of mortality and morbidity. Effectively managing obstetric hemorrhages is a complex task, and addressing this challenge requires laboratory-based screening methods to evaluate the coagulation status of obstetric patients. To explore the connection between obstetric hemorrhage and hemostatic parameters, such as platelet count, prothrombin time (PT), activated partial thromboplastin time (APTT), and plasma fibrinogen levels, this study was undertaken.

In this study comprising a total of 136 participants divided into case and control groups (each with 68 participants), significant differences were observed in four key haemostatic parameters. The case group exhibited higher mean values in Prothrombin Time (PT), Activated Partial Thromboplastin Time (APTT), and lower mean values in Platelet count and Fibrinogen concentration compared to the control group, with p-values < 0.05 for all parameters. Notably, PT was  $14.04 \pm 0.86$  seconds in the case group, significantly higher than the control group's mean of  $12.3 \pm 0.44$  seconds. APTT in the case group was  $37.6 \pm 1.68$  seconds, while it was  $29.87 \pm 2.96$  seconds in the control group. Platelet count was markedly lower in the case group ( $1.32 \pm 0.28 \times 10^9/L$ ) compared to the control group ( $2.49 \pm 0.64 \times 10^9/L$ ), and Fibrinogen concentration was  $1.36 \pm 0.44$  g/dl in the case group, significantly lower than the control group's mean of  $3.71 \pm 0.65$  g/dl. Among the case group, 32.4% experienced Antepartum Hemorrhage (APH), and 67.6% had Postpartum Hemorrhage (PPH). Importantly, a significant linear relationship was found between APH and PPH with platelet levels, while no significant correlations were observed between other haemostatic parameters and APH or PPH. These findings underscore the distinct haemostatic profile associated with obstetric conditions, particularly emphasizing the potential role of platelet levels in predicting and understanding hemorrhagic events.

In a study by Erhabor O et al<sup>94</sup>, The study aimed to explore the impact of obstetric hemorrhage on prothrombin time (PT), activated partial thromboplastin time (APTT), and platelet count (PLC) in pregnant women. The participants included 86 pregnant women with a history of obstetric hemorrhage (divided into antepartum hemorrhage (APH) and postpartum hemorrhage (PPH) subgroups) and 43 age-matched nonhemorrhaging pregnant women as controls. Manual methods and commercially prepared Diagen reagent kits were used for PT and APTT determination, while PLC was assessed using a hemocytometer. Results showed significantly elevated PT and APTT values in both APH ( $20.7 \pm 4.226$  seconds and  $46.04 \pm 8.689$  seconds) and PPH ( $23.17 \pm 2.708$  seconds and  $53.78 \pm 4.089$  seconds) groups compared to normal pregnant women ( $15.85 \pm 0.8930$  seconds and  $36.225 \pm 5.010$  seconds). PLC was significantly lower in APH and PPH groups ( $154.83 \pm 47.019 \times 10^9$  and  $136.43 \pm 43.894 \times 10^9$ , respectively) compared to normal pregnant women ( $291.425 \pm 75.980 \times 10^9$ ).

Women with PPH had significantly higher PT and APTT values than those with APH ( $23.17 \pm 2.708$  seconds and  $53.78 \pm 4.089$  seconds versus  $20.7 \pm 4.226$  seconds and  $46.04 \pm 8.689$  seconds, respectively). PLC was higher in APH compared to PPH ( $P = 0.01$ ). Higher PT and APTT values were observed in the third trimester for women with APH, PPH, and controls, while PLC was significantly lower in the third trimester for all groups. Correlation analysis revealed positive associations between PT and APTT values with trimester ( $r = 0.52$  and  $0.65$ , respectively;  $P = 0.01$ ) and negative correlations between PLC and trimester for APH, PPH, and controls ( $r = -0.36$ ,  $-0.54$ , and  $-0.28$ , respectively;  $P = 0.05$ ). Overall, the findings highlight altered coagulation parameters in obstetric hemorrhage, with variations between APH and PPH, trimester-dependent changes, and correlations between hemostatic parameters and pregnancy trimesters.

Our study revealed a significant elevation in prothrombin time (PT) and activated partial thromboplastin time (APTT) values among women experiencing obstetric hemorrhage, including both antepartum hemorrhage (APH) and postpartum hemorrhage (PPH), when compared to nonhemorrhaging pregnant women. These findings underscore the importance of monitoring PT and APTT as essential parameters in the management of coagulopathy associated with obstetric hemorrhage. The observed increases in PT and APTT values suggest potential disruptions in the coagulation process, emphasizing the clinical relevance of these parameters for assessing and addressing coagulation disorders in the context of obstetric complications.

Prothrombin Time (PT) evaluates the extrinsic and common coagulation pathways, primarily influenced by vitamin K-dependent factors (II, V, VII, and X). Activated Partial Thromboplastin Time (APTT) assesses the intrinsic and common coagulation pathways, sensitive to most coagulation factors except FVII and FXIII. Both PT and APTT are deemed

valuable in monitoring hemostasis during obstetric hemorrhage. In a recent UK study of 18,501 deliveries, 456 cases with blood loss exceeding 1500 mL were examined. PT did not exhibit a clear correlation with the volume of hemorrhage, while APTT showed a weak correlation.

Contrary to earlier studies suggesting limited utility in predicting progression of postpartum hemorrhage (PPH), a retrospective multicenter validation study indicated that an International Normalized Ratio (INR) exceeding 1.5 might predict the necessity for advanced interventions to control obstetric hemorrhage. Current guidelines recommend using an INR greater than 1.5 as an indicator for fresh-frozen plasma (FFP) transfusion, highlighting the potential clinical significance of coagulation monitoring in guiding interventions during obstetric hemorrhage.

The current study noted a noteworthy decrease in platelet count among women experiencing obstetric hemorrhage in comparison to nonhemorrhaging pregnant women. This finding aligns with previous research, indicating that, at the time of diagnosing hemorrhage, obstetric patients with bleeding complications exhibited significantly lower platelet counts compared to healthy parturient women. Moreover, earlier studies have suggested a potential association between decreasing platelet count during obstetric bleeding and the progression to severe hemorrhage. These consistent observations underscore the clinical relevance of monitoring platelet counts as an essential parameter in assessing and managing obstetric hemorrhage, emphasizing its potential role as an indicator of the severity and progression of hemorrhagic events.

In a retrospective analysis encompassing 797 pregnancies, a low platelet count emerged as an independent risk factor for obstetric hemorrhage. Notably, a platelet count below  $100 \times 10^9$  per liter upon admission to the labor ward was associated with an increased incidence of obstetric hemorrhage in certain women. Prior research findings also suggest that platelet transfusion or the use of desmopressin may serve as valid hemostatic therapies for managing obstetric hemorrhage. Current guidelines recommend platelet transfusion when the platelet count falls below  $50 \times 10^9$  per liter in hemorrhaging parturient women. These observations underscore the clinical significance of platelet count



assessment in identifying and managing obstetric hemorrhage, providing valuable insights into potential interventions for maintaining hemostasis in affected individuals.

Obstetricians in developing countries face significant challenges in effectively managing obstetric hemorrhage. Key issues include the lack of routine access to laboratory-based screening for assessing coagulation status, especially in rural settings, and the unavailability of safe and sufficient blood and blood products needed for effective hemorrhage management, particularly in African settings. Timely access to the appropriate quantity and quality of blood and blood products is crucial for saving lives across various clinical conditions. The primary goals of blood volume replacement following obstetric hemorrhage involve rapidly and effectively restoring adequate blood volume to prevent hypovolemic shock, facilitating proper hemostasis, maintaining oxygen-carrying capacity and blood biochemistry, correcting coagulopathy promptly, optimizing resuscitation efforts, and ultimately reducing potentially preventable deaths. These objectives emphasize the critical role of timely and appropriate blood volume replacement in the comprehensive and life-saving management of obstetric hemorrhage, especially in resource-limited settings.

Evidence-based, non-blood-product-related measures for controlling hemorrhage include direct pressure/tourniquet application, appropriate stabilization of fractures, and surgical interventions such as damage control surgery, interventional radiology, and the use of endoscopic and obstetric techniques. Blood-sparing measures, such as intraoperative blood salvage (autologous blood salvage), involving the recovery and re-infusion of blood lost during surgery, are indicated in various obstetric and surgical procedures. Hemostatic drugs like vitamin K, tranexamic acid, and prothrombin complex concentrate (PCC) contribute to achieving hemostasis. PCC, containing clotting factors II, VII, IX, X, protein S, and protein C, undergoes viral inactivation through pasteurization and nanofiltration. Novo 7, a vitamin-K-dependent recombinant human coagulation Factor VIIa (rFVIIa), promotes hemostasis by activating the extrinsic pathway of the coagulation cascade. It plays a vital role in the clinical management of hemorrhage, particularly when other measures like blood products and alternative pharmacologic options prove ineffective. These evidenced-based hemostatic treatments can significantly contribute to managing coagulopathy associated with

obstetric hemorrhage, potentially reducing the reliance on blood products. While these hemorrhage control measures are universally available in developed countries, women in developing nations lack access to these life-saving therapies. This disparity underscores the need for improved accessibility to advanced medical interventions in resource-limited settings to enhance maternal healthcare outcomes.

#### **4. CONCLUSION**

In conclusion, this study sheds light on the critical link between obstetric hemorrhage and hemostatic parameters, highlighting significant variations in prothrombin time (PT), activated partial thromboplastin time (APTT), platelet count, and fibrinogen concentration between women with and without obstetric hemorrhage. The findings underscore the complexity of managing obstetric hemorrhages and the importance of laboratory-based screening in evaluating the coagulation status of affected patients.

The distinct hemostatic profile observed in the case group, characterized by elevated PT and APTT values and lower platelet count and fibrinogen concentration, emphasizes the potential utility of these parameters in assessing and addressing coagulation disorders associated with obstetric complications. Particularly noteworthy is the significant linear relationship found between antepartum hemorrhage (APH) and postpartum hemorrhage (PPH) with platelet levels, indicating a potential predictive role for platelet counts in understanding and managing hemorrhagic events in obstetric patients.

These findings contribute valuable insights to the field, emphasizing the need for comprehensive hemostatic assessments and tailored interventions in obstetric hemorrhage cases. As maternal hemorrhage remains a significant global health challenge, this study underscores the importance of ongoing research and the development of targeted strategies to improve outcomes for women experiencing obstetric complications, especially in resource-constrained settings.

#### **CONSENT**

All authors declare that 'written informed consent was obtained from the patient (or other approved parties) for publication of this case report and accompanying images.

## **ETHICAL APPROVAL**

Ethical approval (IRB No: 89/2022) has been taken by the author to carried out the study.

UNDER PEER REVIEW

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