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# PREGNANCY-INDUCED HYPERTENSION AND ITS EFFECTS ON NEONATAL HEMOGRAM

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#### Abstract

Hypertensive disorders are the most prevalent pregnancy complications. Major pregnancy-related maternal morbidities are caused by preeclampsia toxemia. Preeclampsia increases neonatal morbidity and mortality. Hematological permutation is more common in neonates of hypertensive moms, especially those with preeclampsia, according to many studies.

**Objective:** To determine the frequency of hematological profile of neonates among women with pregnancy induced hypertension visiting to Liaquat National Hospital Karachi.

**Place and Duration:** This Cross-Sectional study was conducted at Department of Obstetrics and Gynecology, Liaquat National Hospital, Karachi, Pakistan from February 8, 2022 to August 7, 2022

**Methods:** All eligible patients who visited LNH, Karachi were studied. After discussing the study's process, drawbacks and benefits, informed consent was obtained. Our study took cord blood from all mothers soon after delivery by double-clamping and severing the cord in between the clamps, then putting the syringe at the placental end of the cord proximally and collecting 4 mL of cord blood. The newborns' blood was transferred to the hospital lab for hematological analysis. All obtained data were entered into the proforma at the end and used electronically for study.

**Results:** Mean maternal age was  $28.4\pm5.5$  years. Mean  $\pm$  SD of neonatal age was  $15.2\pm4.8$  with C.I (14.30.....16.09) days. In distribution of gender of baby, 46 (40.4%) were male while 68 (59.6%) were female. Hematological profile of neonates showed neutropenia in 20 (17.5%) neonates, thrombocytopenia in 45 (39.5%) while polycythemia was noted in 9 (7.9%) neonates.

**Conclusion:** The most prevalent hematological profile was thrombocytopenia, followed by neutropenia and polycythemia. Since the sample originated from different parts of Pakistan, it can be generalized.

**Key words:** Hematological Profile, Neonates, Pregnancy Induced Hypertension, Prevalence

## INTRODUCTION

The definition of pregnancy-induced hypertension (PIH) is systolic blood pressure of 140 mm Hg and diastolic blood pressure of 90 mm Hg. Pregnant women with systolic or diastolic blood pressure increase of 30mm hg or 15mm hg, at least 6 hours apart. One of the leading causes of maternal and newborn death is PIH. PIH complicate 12–22% of pregnancies<sup>1-2</sup>.

According to WHO, every year, about 3.4 million babies have died and about 4.3 million still births have been seen. This is usually seen during first week of life, accounting for 98% of cases in developing countries<sup>3-4</sup>. These mortalities are usually seen in initial few days and weeks of life commonly occurring secondary to events in utero and during childbirth, rather than outside factors. In United States, pregnancy related deaths are more common in blacks in comparison to white females out of which 16% of the deaths are secondary to PIH. Four categories of PIH, has been defined by American College of obstetrics and Gynae including PIH, Pre-eclampsia, eclampsia and HELLP syndrome. Mild PIH described as systolic blood pressure < 160 and diastolic is < 110 mm hg, it is severe when pressures are >160 and 110 respectively. PIH can give rise to many fetal and maternal complications, the maternal complications include thrombocytopenia, DIC, placental abruption, liver failure, intracranial haemorrhage and cardiovascular collapse<sup>5-6</sup>.

Its adverse effects arise due to decrease utero-placental blood supply leading to poor oxygen concentration and acute or chronic utero- placental insufficiency. These complications include 12.6 times higher risk of polycythemia, neutropenia and thrombocytopenia occurring in as many as 22% of the population<sup>7-8</sup>.

In addition, there is increased risk of thrombocytopenia especially in neonates born to hypertensive mothers who develop preeclampsia<sup>9</sup>. It is related with augmented risk of neonatal neutropenia and subsequent neonatal infections<sup>10</sup>. Despite of the fact that low birth weight and prematurity effect the neutrophil count, pregnancy induced hypertension may independently contribute to neutropenia in neonates. Studies done earlier showed that neonates born to mother with PIH, suffer longer duration of decrease neutrophil count and increase susceptibility to hospital acquired infection as compared to normotensive mothers. A study held in Obstetrics/Gynecology Department of National Cheng Kung Medical Centre suggested that pre-eclampsia may adversely affect the fetal hemogram leading to thrombocytopenia, neutropenia and increased red cell mass (polycythemia)<sup>11-12</sup>. A study reported polycythemia 8%, neutropenia 15% and thrombocytopenia 38% in neonates of females with PIH. In another study, mothers with preeclampsia had 50% of their babies with neutropenia and thrombocytopenia<sup>13</sup>.

Hypertensive disorders, coupled with hemorrhage and infection, complicates 5–10% of pregnancies and contribute to maternal morbidity and mortality<sup>14</sup>.

#### **METHODS**

This Cross-Sectional study was conducted at Department of Obstetrics and Gynecology, Liaquat National Hospital, Karachi, Pakistan from February 8, 2022 to August 7, 2022.

Sample size was calculated through W.H.O sample size calculator by using frequency of polycythemia (8%) [6], margin of error (d)=5%, Confidence level (C.I) =95% then the estimated sample size came out to be n=114 and Non-Probability, Consecutive Sampling technique was used for patient selection.

## **INCLUSION CRITERIA**

- Subjects between age group 0 to 28 days.
- Either gender.
- All the neonates born to mother with pregnancy-induced hypertension in accordance with operational definition.

## **EXCLUSION CRITERIA**

- Women with multiple pregnancies.
- Women with diabetes mellitus.
- Women with renal or heart disease.
- History of chronic pulmonary disease.
- Smoking women.
- Neonates with other hematological disorder.
- Neonates with congenital malformations or birth asphyxia.

Data collection was started after approval of synopsis from Research Department of College of Physicians & Surgeons Pakistan and hospital ethical review committee. All the women who delivered the baby at department of Obstetrics and Gynecology, Liaquat National Hospital Karachi, fulfilling the inclusion criteria were included in the study after taken written informed consent. Our study took cord blood from all mothers soon after delivery by double-clamping and severing the cord in between the clamps, then putting the syringe at the placental end of the cord proximally and collecting 4 mL of cord blood. Blood sample of the neonates was also taken and sent to hospital laboratory for hematological profile in accordance with operational definition and results were finalized by consultant hematologists. The study was focused and employed suitable exclusion criteria to control bias and confounders.

Data was entered and analyzed using SPSS-26.0. Shapiro-Wilk tested continuous data normality. Mother's age, parity, gestational age at beginning of pregnancy-induced hypertension, gestational age at delivery, APGAR score, CBC, and birth weight were used to calculate mean  $\pm$  SD/Median. Frequency and percentage were calculated for gender of baby, mode of delivery and hematological profile of neonates i.e., (i.e.: neutropenia, thrombocytopenia and polycythemia). Effect modifiers were controlled through stratification of age of mothers, parity, gestational age at onset of pregnancy induced hypertension, gestational age at delivery, APGAR score of fetuses, CBC (Hb, leukocytes, platelets), birth weight, gender of baby and mode of delivery to see the impact of these on hematological profile of neonates followed by appropriate Chi-Square / Fisher's Exact test consider two-sided P at level of significance  $\alpha$ =5% as significant.

#### **RESULTS**

In this study 114 patients were included to assess the hematological profile of neonates among women with pregnancy induced hypertension visiting to Liaquat National Hospital Karachi and the results were analyzed as:

The distribution of continuous variables was tested by applying Shapiro-Wilk test for maternal age (P=0.129), neonatal age (P=0.117), parity (P=0.092), gestational age at onset of PIH (P=0.071), gestational age at delivery (P=0.109), APGAR score (P=0.355), HB level (P=0.258), leukocyte (P=0.085), platelets (P=0.711) and birth weight (P=0.628) respectively, as shown in TABLE 1.

**TABLE #1:** DESCRIPTIVE STATISTICS OF SHAPIRO-WILK TEST n=114

VARIABLE	MEAN±SD	P-VALUE
Maternal Age (years)	28.7±5.5	0.129
Neonatal Age (days)	15.2±4.8	0.117
Parity	2.2±0.7	0.092
Gestational Age at onset of PIH (weeks)	31.8±5.2	0.071
Gestational Age at delivery (weeks)	37.9±5.9	0.109
APGAR Score	6.7±2.4	0.355
HB level (g/dl)	11.8±1.2	0.258
Leukocytes (cmm)	5646.2±175.6	0.085
Platelets (mcL)	175485.4±618.7	0.711
Birth Weight (kg)	2.9±1.2	0.628

Mean  $\pm$  SD of maternal age was 28.4 $\pm$ 5.5, Mean  $\pm$  SD of neonatal age was 15.2 $\pm$ 4.8, Mean  $\pm$  SD of parity was 2.2 $\pm$ 0.7, Mean  $\pm$  SD of gestational age at onset of PIH was 31.8 $\pm$ 5.2, Mean  $\pm$  SD of gestational age at delivery was 37.9 $\pm$ 5.9, Mean  $\pm$  SD of APGAR score was 6.7 $\pm$ 2.4, Mean  $\pm$  SD of HB level was 11.8 $\pm$ 1.2, Mean  $\pm$  SD of leukocytes was 5646.2 $\pm$ 175.6, Mean  $\pm$  SD of platelets was 175.485.4 $\pm$ 618.7 and Mean  $\pm$  SD of birth weight was 2.9 $\pm$ 1.2 as shown in TABLE 1.

In distribution of gender of baby, 46 (40.4%) were male while 68 (59.6%) were female as shown in FIGURE 1.

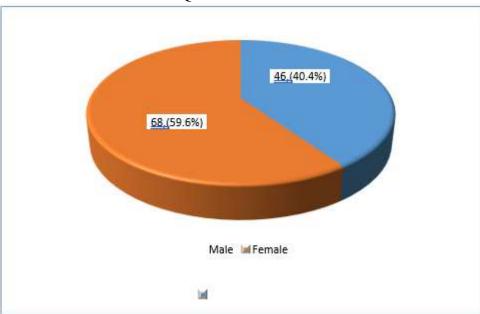


FIGURE # 1 FREQUENCY FOR GENDER OF BABY

Mode of delivery showed vaginal delivery in 50 (43.9%) women while caesarean section was noted in 64 (56.1%) women as shown in FIGURE 2.

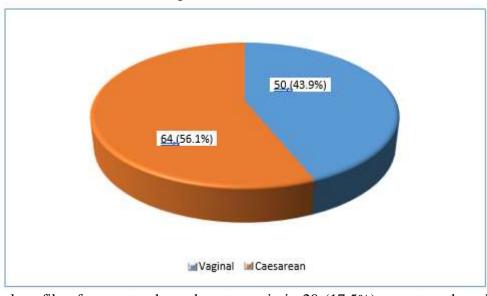


FIGURE # 2 FREQUENCY FOR MODE OF DELIVERY

Hematological profile of neonates showed neutropenia in 20 (17.5%) neonates, thrombocytopenia in 45 (39.5%) while polycythemia was noted in 9 (7.9%) neonates as shown in TABLE 3.

**TABLE # 2:** FREQUENCY FOR HEMATOLOGICAL PROFILE OF NEONATES n=114

HEMATOLOGICAL PROFILE FREQUENCY PERCENTAGE						
	Yes	20	17.5%			
Neutropenia	No	94	82.5%			
	Yes	45	39.5%			
Thrombocytopenia	No	69	60.5%			
	Yes	9	7.9%			
Polycythemia	No	105	92.1%			

Stratification of age of mothers, parity, gestational age at onset of pregnancy induced hypertension, gestational age at delivery, APGAR score of fetuses, CBC (Hb, Leukocytes, Platelets), birth weight, gender of baby and mode of delivery were done with respect to hematological profile of neonates in order to assess statistical difference from TABLE [13-23].

**TABLE # 3** STRATIFICATION OF MATERNAL AGE WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

	AGE GROUP [In years]				
HEMATOLOGICAL PROFILE		20 - 30	>30	P-VALUE	
	Yes	13	7		
		(11.4%)	(6.1%)		
Neutropenia	No	60	34	0.921	
		(52.6%)	(29.8%)		
	Yes	31	14		
		(27.2%)	(12.3%)		
Thrombocytopenia	No	42	27	0.383	
		(36.8%)	(23.7%)		
	Yes	5	4		
		(4.4%)	(3.5%)		
Polycythemia	No	68	37	0.414	
		(59.6%)	(32.5%)		

**TABLE # 4** STRATIFICATION OF PARITY WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

TIEOTITES II-III					
HEMATOLOGICAL		1 - 2	>2	P-VALUE	
PROFILE					
	Yes	11	9		
		(9.6%)	(7.9%)		
Neutropenia	No	39	55	0.269	
		(34.2%)	(48.2%)		
	Yes	31	14		
		(27.2%)	(12.3%)		
Thrombocytopenia	No	19	50	0.0001	
		(16.7%)	(43.9%)		
	•		•		
	Yes	3	6		
		(2.6%)	(5.3%)		
Polycythemia	No	47	58	0.383	
		(41.2%)	(50.9%)		

**TABLE # 5** STRATIFICATION OF GESTATIONAL AGE AT ONSET OF PIH WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

	GESTATIONAL AGE [In weeks]					
HEMATOLOGI PROFILE	HEMATOLOGICAL PROFILE		>30	P-VALUE		
	Yes	15 (13.2%)	5 (4.4%)	0.007		
Neutropenia	No	53 (46.5%)	41 (36.0%)	0.097		
		1	T	_		
	Yes	30 (26.3%)	15 (13.2%)	0.217		
Thrombocytopenia	No	38 (33.3%)	31 (27.2%)	0.217		
Delta dam's	Yes	7 (6.1%)	2 (1.8%)	0.215		
Polycythemia	No	61 (53.5%)	44 (38.6%)	0.215		

**TABLE # 6:** STRATIFICATION OF GESTATIONAL AGE AT DELIVERY WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

		GESTATIONAL		
HEMATOLOGICAL PROFILE		35 - 37	>37	P-VALUE
	Yes	12	8	
		(10.5%)	(7.0%)	
Neutropenia	No	45	49	0.325
		(39.5%)	(43.0%)	
	Yes	25	20	
		(21.9%)	(17.5%)	
Thrombocytopenia	No	32	37	0.338
		(28.1%)	(32.5%)	
	Yes	5	4	
		(4.4%)	(3.5%)	
Polycythemia	No	52	53	0.500
		(45.6%)	(46.5%)	

**TABLE # 7:** STRATIFICATION OF APGAR SCORE WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

		APGAR	SCORE	
HEMATOLOGICAL P	ROFILE	1 - 5	>5	P-VALUE
	Yes	10	10	
		(8.8%)	(8.8%)	
Neutropenia	No	43	51	0.729
		(37.7%)	(44.7%)	
	Yes	19	26	
		(16.7%)	(22.8%)	
Thrombocytopenia	No	34	35	0.461
		(29.8%)	(30.7%)	
	Yes	3	6	
		(2.6%)	(5.3%)	
Polycythemia		·	· 	0.320

**TABLE #8** STRATIFICATION OF HB LEVEL WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

		HB LEVEL [In g/dl]		
HEMATOLOGICAL PROFILE		10 – 12	>12	P-VALUE
	Yes	9	11	
		(7.9%)	(9.6%)	
Neutropenia	No	34	60	0.459
		(29.8%)	(52.6%)	
	·			
	Yes	14	31	
		(12.3%)	(27.2%)	
Thrombocytopenia	No	29	40	0.240
		(25.4%)	(35.1%)	
	Yes	4	5	
		(3.5%)	(4.4%)	
Polycythemia	No	39	66	0.460
		(34.2%)	(57.9%)	

**TABLE # 9:** STRATIFICATION OF LEUKOCYTE WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

	LEUKOCYTE [In cmm]			
HEMATOLOGICAL PROFILE		4400 - 6000	>6000	P-VALUE
	Yes	14	6	
		(12.3%)	(5.3%)	
Neutropenia	No	64	30	0.867
		(56.1%)	(26.3%)	
	Yes	29	16	
		(25.4%)	(14.0%)	
Thrombocytopenia	No	49	20	0.461
		(43.0%)	(17.5%)	
	Yes	4	5	
		(3.5%)	(4.4%)	
Polycythemia	No	74	31	0.110
		(64.9%)	(27.2%)	

**TABLE # 10** STRATIFICATION OF PLATELETS WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

		PLATELET	S [In mcL]	
HEMATOLOGICAL 1	PROFILE	150,000 - 250,000	>250,000	P-VALUE
	Yes	13	7	
		(11.4%)	(6.1%)	
Neutropenia	No	59	35	0.851
		(51.8%)	(30.7%)	
	Yes	26	19	
		(22.8%)	(16.7%)	
Thrombocytopenia	No	53	16	0.031
		(46.5%)	(14.0%)	
	Yes	6	3	
		(5.3%)	(2.6%)	
Polycythemia	No	73	32	0.561
		(64.0%)	(28.1%)	

**TABLE # 11** STRATIFICATION OF BIRTH WEIGHT WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

HEMATOLO	GICAL	BIRTH	WEIGHT [In kg]	
PROFILE		24 25	2.5	P-VALUE
		2.4 - 3.5	>3.5	
	Yes	9	11	
		(7.9%)	(9.6%)	
Neutropenia	No	47	47	0.685
		(41.2%)	(41.2%)	
	Yes	32	13	
		(28.1%)	(11.4%)	
Thrombocytopeni	No	24	45	0.0001
a		(21.1%)	(39.5%)	
	Yes	3	6	
		(2.6%)	(5.3%)	
Polycythemia	No	53	52	0.263
		(46.5%)	(45.6%)	

**TABLE # 12** STRATIFICATION FOR GENDER OF BABY WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

		GENDER	OF BABY	
HEMATOLOGICAL PROFILE		Male	Female	P-VALUE
	Yes	16	4	
		(14.0%)	(3.5%)	
Neutropenia	No	30	64	0.0001
		(26.3%)	(56.1%)	
	Yes	17	28	
		(14.9%)	(24.6%)	
Thrombocytopenia	No	29	40	0.651
• •		(25.4%)	(35.1%)	
		,		•
	Yes	4	5	
		(3.5%)	(4.4%)	
Polycythemia	No	42	63	0.529
		(36.8%)	(55.3%)	

**TABLE # 13** STRATIFICATION FOR MODE OF DELIVERY WITH HEMATOLOGICAL PROFILE OF NEONATES n=114

TROTIEE OF TRESTATIES II-111				
		MODE OF DELIVERY		
HEMATOLOGICAL PROFILE		Vaginal	Caesarean	P-VALUE
	Yes	14	6	
		(12.3%)	(5.3%)	
Neutropenia	No	36	58	0.009
		(31.6%)	(50.9%)	
	•	,		•
	Yes	21	24	
		(18.4%)	(21.1%)	
Thrombocytopenia	No	29	40	0.626
		(25.4%)	(35.1%)	
		•	, ,	•
	Yes	2	7	
		(1.7%)	(5.9%)	
Polycythemia	No	48	61	0.180
		(40.7%)	(51.7%)	

Applied Chi-Square & Fisher's Exact test

## **DISCUSSION**

Hypertension during pregnancy is a leading cause of maternal and newborn morbidity. Hypertensive moms may deliver prematurely due to intrauterine growth limitation <sup>15</sup>.

Hypertension in pregnancy (HIP) is a major cause of maternal and perinatal morbidity and mortality in the obstetric population. HIP is defined as diastolic blood pressure of at least 90 mmHg, systolic blood pressure of at least 140 mmHg, a 15-mmHg spike in diastolic blood pressure, or a 30-mmHg rise in systolic blood pressure in a pregnant woman on two occasions at least 6 hours apart. HIP problems complicate 12–22% of pregnancies. Blacks account for 16% of HIP pregnancy-related deaths in the US, 2 to 3 times more than white women. United Nations and American College of Obstetricians and Gynecologists recognize four HIP categories. Chronic, gestational, preeclampsia, and superimposed preeclampsia/eclampsia are examples. HELLP syndrome is a severe preeclampsia consequence<sup>16</sup>.

Each pregnancy aims for a spontaneous, normal vaginal delivery. Hypertension can complicate pregnancy. Hypertensive pregnancies have a greater rate of newborn morbidity. Hypertensive pregnancies are more likely to result in premature deliveries and low birth weight (LBW)<sup>17-18</sup>. One of the leading causes of maternal and newborn morbidity is pregnancy-induced hypertension (PIH). Pregnancy hypertension reduces blood flow to the uteroplacental unit and causes placental malfunction, which can be deadly. Fatal consequences rise with hypertension severity. Hypertension during pregnancy increases newborn morbidity<sup>19-20</sup>. These children may be born early due to intra-uterine growth limitation. High-rate surgical deliveries and maternal medication side effects may also affect them. These newborns may have a variety of hematological abnormalities that increase morbidity<sup>21-22</sup>. Neutropenia in PIH moms' infants is well reported. Notably, it is the most prevalent congenital Neutropenia. Due to relative intrauterine hypoxia, these babies risk birth asphyxia and meconium aspiration. Drugs used to cure the mother harm the infant. Due to reduced blood supply in pregnant hypertensive disorders, it is lighter than usual<sup>23</sup>.

Our findings are consistent with others. In this study, the average maternal age was  $28.4\pm5.5$  years. Another study reported a mother age of  $31.7\pm4.1$  years, while Al-bahadily AK, et al reported  $30.86\pm6.02$  years<sup>24</sup>.

The baby gender distribution was 46 (40.4%) male and 68 (59.6%) female. Okoye HC, et al. found 53% male and 47% female newborns, while Al-bahadily AK, et al. found 42% male and 58% female<sup>25</sup>.

The neonates had neutropenia in 20 (17.5%), thrombocytopenia in 45 (39.5%), and polycythemia in 9 (7.9%). A study found polycythemia 8%, neutropenia 15%, and thrombocytopenia 38% in neonates of pregnant women with hypertension. In another study, mothers with preeclampsia had 50% of their babies with neutropenia and thrombocytopenia<sup>26</sup>.

Preeclampsia affects 3–10% of pregnancies and causes significant fetal and maternal mortality. Preeclampsia kills 60,000 mothers annually. During pregnancy, pregnant women develop hypertension and proteinuria de novo, along with edema, neurological issues, and intrauterine growth limitation. Pre-eclampsia is diagnosed with a systolic or diastolic BP of 140 or 90 mmHg and proteinuria of 0.3 g or greater in a 24-hour urine specimen. This begins after 20 weeks and can lead to premature delivery and increase mother and baby risk<sup>27</sup>.

## **CONCLUSION**

The most prevalent hematological profile was thrombocytopenia, followed by neutropenia and polycythemia. The sample population represents one institutional experience, but it covers a wide range of Pakistani regions.

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