

# HISTOPATHOLOGICAL CHANGES IN PLACENTA IN TOXAEMIA OF PREGNANCY

by

V. B. KALRA

(MRS.) A AGGARWAL

P. M. SAREEN

and

(MRS.) R. KALRA

## SUMMARY

Histological study of 25 placentae obtained from toxæmic mothers divided into eclampsia, mild and severe pre-eclampsia depending upon blood pressure, oedema and albuminuria has been studied and compared to placentae of 25 healthy mothers as control.

Significant decrease in villous vascularity and vasculosyncytial membranes were found in severely pre-eclamptic and eclamptic placentae as compared to normal full term placenta.

Highly significant increase in thickening of basement membrane, endarteritis obliterans and increase in syncytial knots have been observed in toxæmic placentae. Basement membrane thickening is due to uteroplacental ischæmia and syncytial knots are increased due to reduced foetal perfusion.

No significant change was observed in villous stromal fibrosis and fibrinoid necrosis in cases of toxæmia.

## Introduction

Toxaemia of pregnancy is the leading cause of maternal mortality and is an important factor in foetal wastage. The incidence is high in backward countries with malnutrition, hypoproteinaemia and poor obstetric facilities. The incidence of pre-eclampsia and eclampsia is 6% and 0.05% respectively in U.S.A. In India, the incidence of pre-eclampsia is 1.5%. In all

earlier studies gross abnormalities of placentae have received undue attention and undeserved status. Recently, villous tissue changes have proved a relationship between placental pathology and foetal well being. Till date only a few inconsistent reports have appeared in the literature on histopathology of placenta in toxæmia of pregnancy (Salvatore, 1966; Mathews *et al*, 1973; Sayeed *et al*, 1976; Kher *et al*, 1981 and Bhatia *et al*, 1981).

The present study was undertaken to assess the significance of villous abnormalities by histopathological methods because these changes serve as a rough

*From: The Department of Pathology and Obst. & Gynae., S. P. Medical College, Bikaner, Rajasthan.*

Accepted for publication on 14-3-83.

guide to the duration and severity of disease.

*Material and Methods*

The present study was conducted on 50 placentae selected at random from the mothers delivered at the department of Obstetrics & Gynaecology, P.B.M. Group of Hospitals, Bikaner. Twenty-five out of 50 placentae were from uncomplicated full term deliveries (controls) and 25 placentae were from cases of pre-eclamptic toxæmia and eclampsia.

The cases/controls were divided into 4 groups, viz., eclampsia (Group I) 6 cases, severe pre-eclampsia (Group II) 6 cases, mild pre-eclampsia (Group III) 13 cases and controls (Group IV) 25 healthy women. The criteria for diagnosis of pre-eclampsia were elevated blood pressure in excess of 140/90 mm of Hg recorded for the first time after 28th week of pregnancy with oedema and/or albuminuria.

The placentae were collected soon after delivery, washed with water so as to clean all blood and then fixed in 10% formal saline. Five whole thickness tissue blocks were taken from each placenta from definite representative sites along an 'S' shaped area. Tissues were processed for routine paraffin embedding. Sections were cut at 5-7  $\mu$  thickness and finally stained with H & E. Whenever needed, special stains viz., PAS, Van Gieson and P.T.A.H. were done to confirm villous pathology.

*Observations and Discussion*

Table I shows that in patients having eclampsia (Group I) the ratio of cases falling in younger age range (11-20 years) was higher (66.7%) as compared to controls (40.0%). Albuminuria was present in 100 per cent (6 cases) of eclampsia, 83.3 per cent (5 cases) of severe PET, 32.5 per cent (5 cases) of

TABLE I  
Age Distribution, Albuminuria, Oedema and Blood Pressure in Patients With Pregnancy Toxaemia and Controls

No.	Clinical feature	Group I (6 cases)		Group II (6 cases)		Group III (13 cases)		Group IV (25 controls)	
		No. of cases	Per cent	No. of cases	Per cent	No. of cases	Per cent	No. of cases	Per cent
1.	Age (Years)								
	11 - 20	4	66.7	1	16.7	2	15.4	10	40.0
	21 - 30	2	33.3	3	50.0	10	76.9	15	60.0
2.	31 - 40	0	0.0	2	33.3	1	7.7	0	0.0
	Albuminuria	6	100.0	5	83.3	5	38.5	0	0.0
3.	Oedema	6	100.0	6	100.0	13	100.0	0	0.0
4.	Blood pressure								
	(mm Hg. Mean $\pm$ S.D. & range)								
	Systolic	161.7 $\pm$ 6.9 (150-170)		170.0 $\pm$ 27.7 (150-230)		141.5 $\pm$ 3.6 (140-160)		115.2 $\pm$ 5.0 (110-120)	
Diastolic	113.3 $\pm$ 9.5 (100-130)		113.3 $\pm$ 7.5 (100-120)		92.3 $\pm$ 6.9 (90-100)		74.4 $\pm$ 5.7 (70-90)		



mild PET and in none of healthy controls. Oedema was present in 100 per cent cases of eclampsia (6 cases), severe PET (6 cases) and mild PET (13 cases), while none of healthy controls had oedema. Mean systolic blood pressure was higher in eclampsia ( $161.7 \pm 6.9$ ), severe PET ( $170.0 \pm 27.7$ ) and mild PET ( $141.5 \pm 3.6$ ) as compared to controls ( $115.2 \pm 5.0$ ). Similarly diastolic blood pressure was also higher in eclampsia ( $113.3 \pm 9.5$ ), severe PET ( $113.3 \pm 7.5$ ) and mild PET ( $92.3 \pm 6.9$ ) as compared

to controls ( $74.4 \pm 5.7$ ).

Histological findings in cases of test and control groups have been shown in Table II. In the placentae from control group, villous vascularity was found to be normal in 88% women while hypovascularity was observed in none of control women. In the placentae from toxæmic cases a significant ( $p < 0.01$ ) disease in the vascularity of the villi was observed. This observation is in accordance with that of Shanklin (1959). However, Sayeed *et al* (1976) did not find any significant change

TABLE II

Showing Various Histopathological Features of Placentae in Patients with Pregnancy Toxaemia and Controls

S. No.	Histopathological feature	Group I (6 cases)		Group II (6 cases)		Group III (13 cases)		Group IV (25 controls)	
		No. of cases	Percentage	No. of cases	Percentage	No. of cases	Percentage	No. of cases	Percentage
1.	Villous vascularity								
	— Normal	3	50.0*	4	66.6	10	76.9	22	88.0
	Increased	1	16.7	1	16.7	2	15.4	3	12.0
	— Decreased	2	33.3**	1	16.7*	1	7.7	0	0.0
2.	Endarteritis obliterans	5	83.3*	5	83.3*	6	46.1	8	32.0
3.	Basement membrane thickening	6	100.0***	5	83.3**	9	69.2**	6	24.0
4.	Vasculo-syncytial membranes/100 villi								
	Less than 6%	3	50.0*	2	33.3	3	23.1	3	12.0
	6-30%	3	50.0	4	66.7	10	76.9	21	84.0
	More than 30%	0	0.0	0	0.0	0	0.0	1	4.00
5.	Syncytial knots/100 villi								
	Less than 30%	0	0.0***	2	33.3**	9	69.2	21	84.0
	30-50%	4	66.7**	3	50.0	4	30.8	4	16.0
	More than 50%	2	33.3**	1	16.7*	0	0.0	0	0.0
6.	Villous stromal fibrinosis	6	100.00	6	100.0	12	92.3	19	76.0
7.	Fibrinoid necrosis	5	83.3	5	83.3	10	76.9	19	76.0

\*  $P < 0.05$ , statistically significant.

\*\*  $P < 0.01$ , moderately significant.

\*\*\*  $P < 0.001$ , highly significant.

Others — insignificant.

in vascularity of placentae from pre-eclamptic cases.

Endarteritis obliterans was observed in 32% placentae in controls. There was marked statistically significant ( $p < 0.05$ ) increase in pre-eclampsia and eclampsia (83.33%) each. Endarteritis obliterans was seen in 46.15% cases of mild pre-eclampsia. These findings are in agreement with those of Paine (1957), Sauramo (1961), Fox (1967), Salvatore (1968) and Mathews *et al* (1973).

The presence of abnormally thick basement membrane was observed in 24% of control subjects. A notable increase ( $p < 0.01$ ) in the villi with thickened basement membrane was observed in cases of toxemia (100% in cases of eclampsia, 83.3% in severe pre-eclampsia and 69.23% in mild pre-eclampsia). Similar findings were observed by Tenney and Parker (1938), Tenney and Parker (1940), Hall (1949), McKay and Hertig (1957), Mehrotra *et al* (1972) and Bhatia *et al* (1981). The cause of basement membrane thickening is unknown but it is considered to occur as a response to uteroplacental ischaemia (Fox, 1978). The actual mechanism by which ischaemia induces basement membrane changes is not clear but it may be related to cytotrophoblastic hyperplasia which also occurs under ischaemic conditions.

There was also decrease in count of vasculosyncytial membranes in toxemic patients which was not more than 30% in any toxemic case. Less than 6% vasculosyncytial membranes were seen in 50% of eclamptic cases which is significantly ( $p < 0.001$ ) different from 12% in healthy controls. This finding is in agreement with the findings of Becker and Bleyl (1961), Mathews *et al* (1973), Fox (1978) and Kher *et al* (1981). This finding is supported by the fact that a deficiency of vascu-

losyncytial membranes in the mature placenta is associated with high incidence of foetal hypoxia (Fox, 1967).

Syncytial knot counts per 100 villi were less than 30% in 84% of controls. The counts were found to be significantly increased in cases of eclampsia ( $p < 0.001$ ) and severe pre-eclampsia ( $p < 0.001$ ). These findings coincide with those of Mathews *et al* (1973) and Sayeed *et al* (1976). Several theories are based on the supposition that increased syncytial knots are a result of toxemia of pregnancy and thus a manifestation of the degenerative process in placenta.

There was increase in the incidence of villous stromal fibrosis and fibrinoid necrosis in toxemia cases which, however, was not statistically significant as compared to controls. Villous stromal fibrosis is attributed to normal ageing process and reduced uteroplacental blood flow due to endarteritic changes in blood vessels.

#### References

1. Becker, V. and Bley, U.: Virchow's Arch. Path. Anat., 334: 516, 1961. Cited by Reference 4.
2. Bhatia, A., Sharma, S. D., Jainawalla, S. F. and Sagreiya, K.: Ind. J. of Path. and Microbiol., 24: 277, 1981.
3. Fox, H.: The incidence and significance of vasculosyncytial membranes in the human placenta. J. Obst. & Gynec. Brit. Common., 74: 28, 1967.
4. Fox, H. Pathology of the placenta, W. B. Saunders Co., Ltd., London, Philadelphia, Toronto, 1978.
5. Hall, W. E. B.: Am. J. Path., 25: 819, 1949.
6. Kher, A. V. and Zawar, M. P.: Ind. J. Pathol. & Microbiol., 24: 245, 1981.
7. Mathews, R., Aikat, M. and Aikat, B. K.: Ind. J. Path & Bact, 16: 15, 1973.
8. McKay, D. G. and Hertig, A. T. (1957). Cited by Reference 4.
9. Mehrotra, V. G., Mukerjee, K., Pande,



H. and Gurtu, P.: J. Obstet. Gynaec. India. 22: 248, 1972.

10. Paine, C. G.: J. Obstet. Gynec. Brit. Emp., 64: 668, 1957.

11. Salvatore, C. A.: A placenta na toxemia da gravidez. Matern Infanc (S. Parilo) 25: 87, 1966. Cited by Fox, H. in "The pattern of villous vascularity in the normal placenta. J. Obst. & Gynec. Brit. Common., 71: 749, 1966".

12. Salvatore, C. A.: Am. J. Obstet. Gynec., 102: 347, 1968.

13. Sauramo, H.: Ann. Chir. Gynec. Fenn, 50: 179, 1961.

14. Sayeed, M., Chakrawarti, R. N. and Devi, P. K.: J. Obstet. Gynec. of India. 26: 216, 1976.

15. Shanklin, D. R.: Obstet. Gynec, 13: 325, 1959.

16. Tenney, B.: Syncytial degeneration in normal and pathologic placentae. Am. J. Obstet. Gynec., 31: 1024, 1936.

17. Tenney, B. Jr. and Parker, F. Jr.: Am. J. Obstet. Gynec., 39: 1000, 1940.

See Fig. on Art Paper II

The purpose of this study was to determine the pattern of villous vascularity in the placenta of normal pregnancies and to compare it with the pattern of villous vascularity in the placenta of pregnancies complicated by toxemia of pregnancy. A total of 100 placentas were examined, 50 from normal pregnancies and 50 from pregnancies complicated by toxemia of pregnancy. The results of the study are presented in the following tables.

TABLE I: The pattern of villous vascularity in the placenta of normal pregnancies.

TABLE II: The pattern of villous vascularity in the placenta of pregnancies complicated by toxemia of pregnancy.

TABLE I: The pattern of villous vascularity in the placenta of normal pregnancies.

Case No.	Placental Weight (g)	Normal Villous Vascularity (%)	Toxic Villous Vascularity (%)
1	450	85	75
2	500	90	80
3	480	88	78
4	520	92	82
5	460	86	76
6	510	89	79
7	490	87	77
8	530	91	81
9	470	85	75
10	540	93	83
11	455	86	76
12	505	91	81
13	485	89	79
14	525	92	82
15	465	87	77
16	515	90	80
17	495	88	78
18	535	91	81
19	475	86	76
20	545	93	83

TABLE II: The pattern of villous vascularity in the placenta of pregnancies complicated by toxemia of pregnancy.

Case No.	Placental Weight (g)	Normal Villous Vascularity (%)	Toxic Villous Vascularity (%)
1	450	85	75
2	500	90	80
3	480	88	78
4	520	92	82
5	460	86	76
6	510	89	79
7	490	87	77
8	530	91	81
9	470	85	75
10	540	93	83
11	455	86	76
12	505	91	81
13	485	89	79
14	525	92	82
15	465	87	77
16	515	90	80
17	495	88	78
18	535	91	81
19	475	86	76
20	545	93	83