Original Article

Histopathological study of placentae in intrauterine growth retardation pregnancies in a tertiary care hospital and correlation with fetal birth weight

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Keywords:
Birth weight;
Chorionic villi;
Cytotrophoblasts;
Fetus;
IUGR;
Placenta;
Syncytial knotting

ABSTRACT

Background: Intra uterine Growth Retardation is the most significant factor of perinatal mortality. The aim of this study was to assess the histopathological changes in the placenta in association with IUGR and correlation with fetal birth weight.

Materials and methods: A total of 100 placentae were included. Twenty five normal placentae and 75 placentae were from IUGR pregnancies were included.

Results: Intervillous fibrin deposition (64%), increased syncytial knotting (64%), stromal fibrosis (65%), cytotrophoblastic hyperplasia (44%) and basement membrane thickening (40%) were seen along with hypovascular villi and infraction were present in 32% and 28% respectively. These changes were seen less in the control group (p<0.001). Statistically significant association between the birth weight and microscopic changes (chi square=19.543, degree of freedom=4, p<0.005) was observed.

Conclusion: Severity of intrauterine growth retardation is related to the microscopic change in the placenta. The number and severity of microscopic changes in IUGR placentas increased with decreasing fetal birth weight.

INTRODUCTION

The antenatal health care given to pregnant women has great influence on the rates of perinatal death and morbidity. Amongst the different causes of perinatal mortality, Intra Uterine Growth Retardation (IUGR) is the single most significant factor and about 2/3 deaths among infants with less than 2500 g birth weight.¹ Causes of IUGR can be divided into maternal, fetal, placental and unknown causes, but the basic pathophysiology is due to reduced availability of nutrients in mother or its reduced transfer by the placenta to the fetus. It may also be due to reduced utilization by the fetus.²
Placenta is the life line of the fetus and is a versatile organ with multiple functions, which are vital to the normal development of the fetus. Any disease adversely affecting the mother or the fetus brings about morphological changes in the placenta. These morphological changes can be studied by gross and microscopic examination of the placentas. The information obtained will be of great value in understanding the causes of adverse maternal, fetal and neonatal outcomes.

The present study was undertaken to assess the significance of placental changes in causing the reduction in birth weight of the babies in IUGR pregnancies.

MATERIALS AND METHODS

The present study was a retrospective and was conducted on 100 placentas collected in 2001 to 2002. Permission from the institutional review committee was obtained. Among these 75 placentas were from clinically diagnosed IUGR pregnancies and 25 were from normal pregnancies. Fetal birth weight was noted in all cases. Placentas were collected soon after delivery, washed with water so as to clean all blood. For fixation, the placenta was cut into the multiple longitudinal slices of 2 cm and kept in 10% formalin for 2-3 days. After fixation, In addition to the sections from membranes, four whole thickness sections were taken from placenta. Additional sections were taken whenever gross lesions were detected along with the transverse section of the cord near to its attachment over the placental surface. Paraffin embedded blocks were made after processing and sections were cut approximately 2-3 microns thick and stained routinely by hematoxylin and eosin. Whenever needed, special stain viz. PAS, Masson’s trichrome, Von Gieson were done to confirm villous pathology. All the data were collected from the medical reports.

Table 1: Distribution of various microscopic lesions in placenta

<table>
<thead>
<tr>
<th>Microscopic lesion</th>
<th>Study (n=75)</th>
<th>Group %</th>
<th>Control (n=25)</th>
<th>Group %</th>
</tr>
</thead>
<tbody>
<tr>
<td>Inter villous fibrin deposition</td>
<td>48</td>
<td>64***</td>
<td>3</td>
<td>12</td>
</tr>
<tr>
<td>Increased syncytial knotting</td>
<td>48</td>
<td>64***</td>
<td>2</td>
<td>8</td>
</tr>
<tr>
<td>Stromal fibrosis</td>
<td>42</td>
<td>56***</td>
<td>3</td>
<td>12</td>
</tr>
<tr>
<td>Cytotrophoblastic hyperplasia</td>
<td>33</td>
<td>44***</td>
<td>0</td>
<td>0</td>
</tr>
<tr>
<td>Basement membrane thickening</td>
<td>30</td>
<td>40***</td>
<td>0</td>
<td>0</td>
</tr>
<tr>
<td>Infraction (fig.3)</td>
<td>21</td>
<td>28***</td>
<td>0</td>
<td>0</td>
</tr>
<tr>
<td>Fibromuscular sclerosis of fetal stem arteries</td>
<td>21</td>
<td>28**</td>
<td>1</td>
<td>4</td>
</tr>
<tr>
<td>Hypovascular villi</td>
<td>24</td>
<td>32***</td>
<td>0</td>
<td>0</td>
</tr>
<tr>
<td>Fibrinoid necrosis of villi</td>
<td>15</td>
<td>20*</td>
<td>1</td>
<td>4</td>
</tr>
<tr>
<td>Calcification</td>
<td>27</td>
<td>36</td>
<td>8</td>
<td>32</td>
</tr>
<tr>
<td>Chronic villitis</td>
<td>9</td>
<td>12*</td>
<td>2</td>
<td>8</td>
</tr>
</tbody>
</table>

P < 0.05 - Satisfactory significant
P < 0.01 - Moderately significant
P < 0.001 - Highly significant
Others - Insignificant

Statistics

Statistical correlation of was carried out between them by using SPSS 18 version. Chi-square test with or without yate’s correction was used as and when required. P < 0.05 was taken as critical level of significance.

RESULTS

Several major histologic changes occur in IUGR placentas compared to the normal placentas. These changes include intervillous fibrin deposition (64% cases, fig. 1), increased syncyial knot-ting (64% cases, fig. 2), stromal fibrosis (56% cases), cytotrophoblastic hyperplasia (44% cases) and basement membrane thickening (40% cases).

In addition, hypovascular villi and infraction were present in 32% and 28% of placentas of IUGR pregnancies. All these changes were present in a lower number of cases of the control group (p<0.001). Fibromuscular sclerosis of fetal stem arteries was present in 28 percent of IUGR placentas (P<0.01). (Table 1)

Table 2 shows the association of fetal birth weight with the microscopic changes in placenta. For this purpose IUGR placentas are divided into four groups depending upon the number of microscopic findings. Group I placentas showed one significant microscopic change, and similarly

Group II to IV showed two, three and four or more microscopic features. The table shows that the number of microscopic changes increases with the decrease in birth weight, as 13 cases (61.9%) out of 21 cases having birth weight less than 2 kg showed grade IV microscopic change. However when the birth weight is above 2 kg only 14.8 % cases showed grade IV change. Since group I is having only one case, association was studied with group II and above. Thus there is statistically significant association between the birth weight and microscopic changes (chi square value...
DISCUSSION

IUGR is associated with medical conditions that interfere with the circulation and efficiency of the placenta, with the development or growth of the fetus or with the general health and nutrition of the mother. IUGR may be a normal fetal response to nutritional or oxygen deprivation. Therefore, the issue is not the IUGR, but rather the ongoing risk of malnutrition or hypoxia. IUGR infants more frequently show multiple types of lesions in their placentas. Relationship of all placental lesions to IUGR was independent of each other. It is evident that there is not a single utero placental or villous lesion that result in fetal growth restriction. It is more likely that it is the accumulation (or total burden) of placental injury that when present for a sufficient time interval leads to IUGR. So, the patterns of lesions are more important in elucidating the casual pathways by which placental histopathology is translated into IUGR and histology examination of placenta may clarify a cause for delayed fetal growth. The severity of placental abnormalities expressed as cumulative number of placental lesions is a significant risk factor for IUGR. Redline et al demonstrated that five chronic patterns of placental injury are significantly increased in placentas from pregnancies complicated by IUGR. These are maternal and fetal vascular obstructive lesions, high grade villitis of unknown etiology (VUE), perivillous fibrinoid deposition and chronic abruption. In our study, several, major histopathological changes were observed in IUGR placentas. They were intervillous fibrin deposition, increased syncytial knotting, stromal fibrosis of villi, cytotrophoblastic hyperplasia and thickening of basement membrane. In addition, infraction and hypovascular villi were present in a significant number of IUGR placentae. All these findings indicate disturbance in placental maturation and morphological changes due to reduced utero placental blood flow. End result of all these placental changes is a growth retarded fetus. Similar observations were made in studies on IUGR placental by Dawson L et al. Redline et al also observed increased intervillous fibrin deposition and extent of increased intervillous fibrin showed the strongest correlation with both placental weight and fetal weight. After comparing the microscopic change

Table 2: Showing association of fetal birth weight with the number of microscopic findings

<table>
<thead>
<tr>
<th>S.No</th>
<th>Fetal weight (Kgs)</th>
<th>Group I</th>
<th>Group II</th>
<th>Group III</th>
<th>Group IV</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>1.</td>
<td>2-2.4</td>
<td>1</td>
<td>17</td>
<td>28</td>
<td>8</td>
<td>54</td>
</tr>
<tr>
<td>2.</td>
<td>1.5-2</td>
<td>0</td>
<td>1</td>
<td>5</td>
<td>6</td>
<td>12</td>
</tr>
<tr>
<td>3.</td>
<td>&lt;1.5</td>
<td>0</td>
<td>0</td>
<td>2</td>
<td>7</td>
<td>9</td>
</tr>
<tr>
<td>Total</td>
<td></td>
<td>1</td>
<td>18</td>
<td>35</td>
<td>21</td>
<td>75</td>
</tr>
</tbody>
</table>

Chi Square Value = 19.543
Degree of freedom = 4
P < 0.005

Figure 1: Photomicrograph showing placenta with massive intervillous fibrin deposition (HE stain, X40).

Figure 2: Photomicrograph showing placenta with increased syncytial knotting (HE stain, X40).

Figure 3: Photomicrograph showing placental infarct with ghost like villi (H&E, 40x)
with the fetal birth weights, it is found that severity of IUGR is related to the severity of microscopic change in the placenta. Both number and severity of microscopic changes in placenta increased with decreasing fetal birth weight. The increasing number and severity of microscopic changes indicate the increasing burden on the placenta resulting in chronic placental insufficiency and the end result of which is a growth retarded fetus. Masodkar et al\textsuperscript{10} and Moriohio Iwata et al\textsuperscript{11} have found similar association between placental changes and birth weight. Thus placental histological examination may clarify the cause for delayed fetal growth and contribute to the better understanding of the genesis of intrauterine growth retardation.\textsuperscript{5}

CONCLUSION

Pathological changes in placenta were higher amongst IUGR infants as compared to control group. Placental pathology examination gives more accurately the cause of IUGR infants. The findings suggest that chronic ischemia and associated secondary changes probably lead to improper perfusion and IUGR. Comparison of the microscopic changes with the fetal birth weight, revealed that severity of IUGR is related to the severity of microscopic change in the placenta. Therefore, to know the potential pathologies in the placenta, all placentas of IUGR infants should be examined.

Conflict of interest: None

REFERENCES