ASSOCIATION OF LOW VITAMIN D WITH PRE ECLAMPSIA.

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Abstract:
Objective: To determine the association of preeclampsia and vitamin D deficiency. Study design: Cohort study. Place and duration of study: Gynecology Department, Khwaja Muhammad Safdar Medical College, during 01-07-2018 to 31-12-2018. Material and Methods: In this study the pregnant females irrespective of their gravida and parity status and having gestational age more than 20 weeks were included. The cases that had vitamin D level lower were labelled as cases and those with normal levels as controls. They were followed monthly until delivery and during this period they were followed for protein urea and BP to label pre eclampsia. Results: In this study 100 cases were selected. 50 in each group. Mean age in case and control group was 29.57±4.57 vs 28.43±3.79 years and mean duration of gestation at presentation was 26.43±5.11 vs 27.42±5.23 weeks. There were 39 vs 41 multigravida and 37 vs 38 uneducated females in cases and control group respectively. Preeclampsia was seen in 4 (08%) of control and 14 (28%) cases with odd ration of 4.47 (1.35 to 14.75) with p value 0.01. Conclusion: There is significant association of low vitamin D level and risk of pre eclampsia. Key words: pre eclampsia, vitamin D, Hypertension.

Introduction

Preeclampsia (PE) is one of the highly morbnd and can be fatal entity if converted into eclampsia. Its one of the hypertensive disorders of the pregnancy and seen in the later stages of pregnancy. Its incidence is relatively low; though not that uncommon and is seen in less than 10% of the cases.1,2

There are number of factors that can lead to its development and can be both maternal, fetal or mixed. Furthermore, they can also be attributed on the basis of underlying pathophysiology comprising angio-genetic, endothelial dysfunction, syncyto-tropho blastic microparticles (STMP), and inflammatory cells activation that results in wide cascade of interactions and lead to pre eclampsia. Still there are around hundred factors that can predispose to this entity.3,4

Dietary factors are one important aspect of the offending factors that can result in hypertensive disorders like pre eclampsia and among them role of vitamin A, C and D have been seen in the past and great emphasis is being given to the latter one. It has shown to impact the angiogenesis and anti inflammatory properties and also inhibition of various factors that can result in better control of the disease, however the mechanism is complex.5,6

There is further data regarding the co administration of calcium along with vitamin D and that is even been recommended by World Health Organization (WHO) in the prevention of pre eclampsia.7,8

Objective:

To determine the association of preeclampsia and vitamin D deficiency.

Patients and methods;

This cohort study was carried out at Department of Gynecology Department, Khwaja Muhammad Safdar Medical College during 01-07-2018 to 31-12-2018. In this study the pregnant females irrespective of their gravida and parity status and having gestational age more than 20 weeks were included. The cases with prior history of pre eclampsia or eclampsia, the known cases of hypertension and those with end organ liver, renal or hepatic failure were excluded from this study. Then two groups were made. Group 1 as cases and group 2 as healthy controls. The vitamin D levels of all the subjects were assessed at 1st visit and then these cases that had vitamin D level lower were labelled as cases and those with normal levels as controls. They these cases were followed monthly until delivery and during this period they were followed for protein urea and BP to label pre eclampsia.
Statistical analysis;

The data was entered and analysed by SPSS-version 23.0. Qualitative variables were presented as frequencies and quantitative as mean and standard deviation. Odd ratios were calculated to look for association.

Results;

In this study 100 cases were selected. 50 in each group. Mean age in case and control group was 29.57±4.57 vs 28.43±3.79 years and mean duration of gestation at presentation was 26.43±5.11 vs 27.42±5.23 weeks. There were 39 vs 41 multigravida and 37 vs 38 uneducated females in cases and control group respectively as in table I. Preeclampsia was seen in 4 (08%) of control and 14 (28%) cases with odd ration of 4.47 (1.35 to 14.75) with p value 0.01 as in table II.

Table I: Study variables (n= 50 each)

<table>
<thead>
<tr>
<th>Variables</th>
<th>Case</th>
<th>Control</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age</td>
<td>Mean ± SD</td>
<td>Range</td>
</tr>
<tr>
<td></td>
<td>29.57±4.57</td>
<td>28.43±3.79</td>
</tr>
<tr>
<td>Age of gestation</td>
<td>26.43±5.11</td>
<td>27.42±5.23</td>
</tr>
<tr>
<td>Variables</td>
<td>Frequency</td>
<td>Frequency</td>
</tr>
<tr>
<td>Primigravida</td>
<td>11</td>
<td>9</td>
</tr>
<tr>
<td>Multigravida</td>
<td>39</td>
<td>41</td>
</tr>
<tr>
<td>Educated</td>
<td>13</td>
<td>12</td>
</tr>
<tr>
<td>Uneducated</td>
<td>37</td>
<td>38</td>
</tr>
<tr>
<td>Rural</td>
<td>40</td>
<td>37</td>
</tr>
<tr>
<td>Urban</td>
<td>10</td>
<td>13</td>
</tr>
<tr>
<td>Smoker</td>
<td>4</td>
<td>3</td>
</tr>
<tr>
<td>Non smoker</td>
<td>46</td>
<td>47</td>
</tr>
</tbody>
</table>

Table II: Association of vitamin D deficiency and pre eclampsia

<table>
<thead>
<tr>
<th>Pre eclampsia</th>
<th>Odds ratio (95% CI)</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Yes</td>
<td>No</td>
<td></td>
</tr>
<tr>
<td>Control</td>
<td>4 (8%)</td>
<td>46 (92%)</td>
</tr>
<tr>
<td>Case</td>
<td>14 (28%)</td>
<td>36 (72%)</td>
</tr>
</tbody>
</table>

Discussion;

Pre eclampsia is not uncommon and can be a precursor of a highly morbid and even fatal entity as eclampsia. There are number of maternal and fetal related factors that are addressed and being managed to decrease the risk and still there are many un noticed and work is being done to explore. Dietary deficiencies especially with lower levels of vitamin D is being highly considered for this and for which this study was planned to look its degree of association.8-9

Preeclampsia was seen in 4 (08%) of control and 14 (28%) cases with odd ration of 4.47 (1.35 to 14.75) with p value 0.01. These results were comparable to the findings of the previous studies. According to a study done by Bondar LM et al they also found that there was significant association of the risk of pre eclampsia and low vitamin D and they further described that the risk is even doubles if there is a 50 nmol/l decrease in its level with odd ration of 2.5 95% CI 1.1-54.10

The data from 2 other studies done by Crikshank DP and Seely EW et al also revealed increased risk of pre eclampsia in cases with low vitamin D with p values < 0.05.11-12 In another recent study from Iceland by Olafsdottir AS revealed a U shaped curve to look for risk of pre eclampsia and vitamin D levels.13

This was also supported by the data of the two randomized trials where they supported the role of vitamin D in the prevention of preeclampsia. In an uncontrolled trial, supplementation with a multivitamin/mineral supplement and halibut liver oil (containing 900 IU/d vitamin D) provided at 20 wk gestation reduced the odds of preeclampsia by 32% (95% CI, 11–47%).14-15 Marya et al randomized 400 women at 20–24 wk gestation to vitamin D (1200 IU/d) and calcium (375 mg/d) supplements or no treatment and found a significant reduction in blood pressure (P < 0.001) and a non significant reduction in the incidence of preeclampsia in the treated group compared with the untreated (6 vs. 9%).16 In a cohort study, investigators found that regular supplementation with vitamin D in the first year of life halved the risk of preeclampsia in the female offspring’s first pregnancy (28).

Studies of seasonal patterns in preeclampsia show the lowest incidence in summer, when sunlight is plentiful and serum 25(OH)D concentrations are at their peak, and the highest incidence in winter, when synthesis of vitamin D3 is limited in temperate zones and serum 25(OH)D levels are at their nadir.17,18

Conclusion;

There is significant association of low vitamin D level and risk of pre eclampsia.

References;

4. Gidlof S, Silva AT, Gustafsson S, Lindqvist PG. Vitamin D and the risk of preeclampsia--a nested case-