An Observational Study on Serum Ferritin Level in Dengue Fever and Its Correlation with Severity of Dengue Fever
Authors
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Abstract
Aims and Objectives: To study the serum ferritin level in dengue fever and its correlation with severity of dengue fever
Methods: The study was performed on patients admitted for dengue fever in Rajah Muthiah Medical College and Hospital under the Division of General Medicine from October 2017 to September 2019. After satisfying the inclusion and exclusion criteria 50 dengue IgM positive patients were included in this study. Their serum ferritin levels on the day of admission was measured. The relationship between serum ferritin levels and platelets and severity of dengue fever was observed.
Results: There was significant correlation between serum ferritin levels and thrombocytopenia (p=0.006). Patients with increased ferritin levels had significant decrease in platelet count. Bleeding tendencies were found to be higher in patients who had increased ferritin levels and thrombocytopenia.
Conclusion: Serum ferritin levels were increased in dengue fever. Patients with high ferritin levels were found to have increased incidence of thrombocytopenia. Hence high ferritin levels can be used as a indicator for assessing disease severity and prognosis of dengue fever patients. Those patients with high serum ferritin levels should be monitored carefully.

Introduction
Ferritin is a acute phase reactant produced by reticulo-endothelial cells in response to inflammation and infection. It is produced in significant amount by monocytes, macrophages and hepatic cells. It has been shown that synthesis of ferritin can be induced by cytokines.

The reticulo endothelial system plays a critical role in iron metabolism by processing hemoglobin from senescent red blood cells. Acute inflammation and infection induce the blockade of iron release resulting in a decreased in serum iron, a virulence factor for many microorganisms. Elevated levels of serum ferritin, an acute phase reactant reflect the clinical response to deprive micro organisms of serum iron.

Hyperferritinemia is associated with immune activation and coagulation disturbances. A raise in serum ferritin level has been observed to have linear correlation with severity of dengue fever

Aims and Objectives
- To measure the serum ferritin levels in dengue fever patients
- To determine the relationship between serum ferritin level and thrombocytopenia in patients with dengue fever

Inclusion Criteria
Confirmed cases of Dengue IgM positive, IgG positive and NS1 Ag positive (Age > 13 yrs)
Exclusion Criteria
- Pregnancy
- Other febrile illness
- Mixed infections
- Patients with known hematological disorders
- Patients on heparin
- Patients on chemotherapy and radiotherapy / immuno compromised patients.
- Known patients with qualitative and quantitative platelet disorders

Methods
The study was performed on patients admitted for dengue fever in Rajah Muthiah Medical College and hospital under the Division of General Medicine from October 2017 to September 2019. After satisfying the inclusion and exclusion criteria 50 dengue IgM positive patients were included in this study. Their serum ferritin levels on the day of admission was measured. The relationship between serum ferritin levels and platelets and severity of dengue fever was observed. Statistical analysis was done using the SPSS software.

Patient Characteristics
Table 1: Sex Distribution of Study Participants

<table>
<thead>
<tr>
<th>Sex</th>
<th>No. of patients</th>
</tr>
</thead>
<tbody>
<tr>
<td>Male</td>
<td>28</td>
</tr>
<tr>
<td>Female</td>
<td>22</td>
</tr>
<tr>
<td>Total</td>
<td>50</td>
</tr>
</tbody>
</table>

In a total number of 50 patients 56% (n=28) were males and 44% (n=22) were females.

Figure 1: Sex Distribution of Study Participants

Table 2: Age Distribution of Study Participants

<table>
<thead>
<tr>
<th>Age</th>
<th>No. of patients (n)</th>
<th>Percentage (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>11-25</td>
<td>19</td>
<td>38</td>
</tr>
<tr>
<td>26-40</td>
<td>12</td>
<td>24</td>
</tr>
<tr>
<td>41-55</td>
<td>13</td>
<td>26</td>
</tr>
<tr>
<td>56-70</td>
<td>2</td>
<td>4</td>
</tr>
<tr>
<td>71-85</td>
<td>4</td>
<td>8</td>
</tr>
<tr>
<td>Total</td>
<td>50</td>
<td>100</td>
</tr>
</tbody>
</table>

In a total number of 50 patients most of them were in the age group of 11-25 (n=19) and least number was in the age group 56-70 (n=2).
Table 3: Sex Distribution and Ferritin Co-Relation among Study Participants

<table>
<thead>
<tr>
<th>SEX</th>
<th>FERRITIN</th>
<th>TOTAL</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>HIGH</td>
<td>NORMAL</td>
</tr>
<tr>
<td>MALE</td>
<td>19</td>
<td>9</td>
</tr>
<tr>
<td>FEMALE</td>
<td>11</td>
<td>11</td>
</tr>
<tr>
<td>TOTAL</td>
<td>30</td>
<td>20</td>
</tr>
</tbody>
</table>

Out of 50 patients 28 were male and 22 were female. Chi square test was applied to test the significance between ferritin levels and sex distribution. There was no significant difference (p=.201) found between the variation in ferritin levels and sex distribution. This implies ferritin levels does not depend upon sex distribution.

Figure 3: Sex Distribution of Ferritin Co-Relation among Study Participants
Table 4: Significance between Ferritin Levels and Thrombocytopenia

<table>
<thead>
<tr>
<th>FERRITIN</th>
<th>THROMBOCYTOPENIA</th>
<th>TOTAL</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>PRESENT</td>
<td>ABSENT</td>
</tr>
<tr>
<td>HIGH</td>
<td>16</td>
<td>14</td>
</tr>
<tr>
<td>NORMAL</td>
<td>3</td>
<td>17</td>
</tr>
<tr>
<td>TOTAL</td>
<td>19</td>
<td>31</td>
</tr>
</tbody>
</table>

Chi square test was applied to test significance between serum ferritin levels and thrombocytopenia. There was significant correlation ($p=0.006$) between serum ferritin levels and thrombocytopenia. This implies that increase in ferritin levels is associated with significant decrease in platelet count.

Figure 4: Significance between Ferritin Levels and Thrombocytopenia

Results and Analysis

1) In this study 50 dengue IgM Elisa POSITIVE patients were studied.
2) Out of 50 patients, 28 were males and 22 were females.
3) Most of the patients were in the age group of 11-25 and least number of patients were in the age group of 56-70.
4) There was significant decrease in platelet count in patients with high serum ferritin levels.
Discussion
Currently dengue is causing major public health concern throughout the World particularly in South East Asian countries. Serum ferritin levels are elevated in patients with dengue fever and elevated serum ferritin levels are associated with increased incidence of thrombocytopenia.
In a study conducted by Petchaiappan V et al., serum ferritin levels negatively correlated with the platelet count ($r = 0.51$, $p < 0.001$).
Study conducted by Soundravally R et al. concluded that raised ferritin levels could predict the dengue severity with sensitivity of 76.9% and specificity of 83.3% on the day of admission.
Study conducted in Aruba and Brazil showed increased levels of serum ferritin were significantly associated with disease severity in dengue virus infection.
In a study conducted by Diwakar TN et al., concluded that significant association was found between raised serum ferritin, AST, ALT and severe Dengue ($p<0.001$)

Conclusion
Hyperferritinemia can be used as an indicator for assessing disease severity and prognosis of dengue fever patients. Those patients with high serum ferritin levels should be monitored carefully than patients with normal serum ferritin levels.

References