HEPATOCELLULAR DAMAGE DUE TO HALOTHANE ANAESTHESIA

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Abstract
To determine the hepatotoxic effect of halothane anaesthesia, liver function tests were done a day prior to surgery in 41 cases undergoing various elective operations. The tests were repeated on 7th post operative day to see the derangement. Statistically significant rise in serum bilirubin, ALT and alkaline phosphatase was found post-operatively. As certain groups of patients are more prone to develop halothane induced hepatotoxicity, damage in these cases could be prevented by careful pre-operative assessment (JPMA 38: 262, 1988).

INTRODUCTION
Halothane (Fluothane) a non-explosive halogenated hydrocarbon, which is structurally similar to chloroform is one of the most widely used general anaesthetic agents. It is also known to cause hepatic injury and post operative jaundice.1-7 Most studies suggest that a particular group of patients like obese elderly females, patients with unexplained spiking pre-operative fever, those exposed to halothane in the preceding 6 weeks, and those known to develop jaundice and liver damage after exposure to halothane are prone to develop such a damage8-11.

The present study was done to see the extent of liver damage following long halothane anaesthesia during a major surgery. There was no added risk to the patient because only liver function tests were added to the list of pre and post operative tests.

PATIENTS AND METHODS
Fortyone patients undergoing various elective surgical procedures under halothane anaesthesia were selected for the study. None of the patients were jaundiced or had history of jaundice in the past. Pre-operative liver function tests (serum bilirubin, ALT and alkaline phosphatase) a day prior to surgery were within normal limits in all the cases. Liver function tests were repeated on the 7th post operative day to see the changes indicating hepatocellular damage.

Statistical analysis was done using critical freedom ratio (F-ratio) or 99% probability level, and students’ T test.12

RESULTS
Of 41 cases, there were 21 males and 20 females. Except for 3 children under 12 years of age, all were adults. Significantly higher post operative values were obtained for serum bilirubin, ALT and alkaline phosphatase, suggesting hepatic damage by halothane (Table I).
<table>
<thead>
<tr>
<th></th>
<th>Mean</th>
<th>SD</th>
<th>F. Ratio</th>
<th>99% Probability Level or Critical ‘F’ Ratio</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mg%</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Serum</td>
<td>Pre-op</td>
<td>0.56</td>
<td>0.13</td>
<td>29.0</td>
</tr>
<tr>
<td>Bilirubin</td>
<td>Post-op</td>
<td>0.91</td>
<td>0.54</td>
<td></td>
</tr>
<tr>
<td>SGPT U/L</td>
<td>Pre-op</td>
<td>21.93</td>
<td>6.97</td>
<td>2.64</td>
</tr>
<tr>
<td></td>
<td>Post-op</td>
<td>37.27</td>
<td>11.37</td>
<td></td>
</tr>
<tr>
<td>Serum Alk. Phosphate U/L</td>
<td>Pre-op</td>
<td>28.22</td>
<td>12.44</td>
<td>6.85</td>
</tr>
<tr>
<td></td>
<td>post-op</td>
<td>54.34</td>
<td>32.57</td>
<td></td>
</tr>
</tbody>
</table>

SD = Standard Deviation
‘F’ Ratio = Freedom Ratio
Table II shows the percentage frequency of abnormal liver function tests seen post operatively. Of 123 liver function tests done post operatively in 41 cases, 29 (23.58%) tests were found to be abnormal in 22 patients.

**DISCUSSION**

The role of halothane in the production of post-operative hepatic damage and jaundice is controversial because hepatic damage following general anaesthesia was seen even before halothane was used as an anaesthetic\(^8\). Later it was thought that hypoxia caused by excessive use of nitrous oxide along with halothane is the cause of hepatic damage, rather than halothane alone\(^2\). Hepatotoxicity following the use of halothane usually mild to moderate with slight derangement has been reported in one or more components of liver function tests in about 20% cases\(^9,10\). Similar findings were seen in the present study (23.58%). Fulminating hepatotoxicity, though rare, is mostly seen in adults\(^11\). Children are often less vulnerable to hepatic damage than adults.\(^5,11\) Severe hepatic necrosis occurs in

<table>
<thead>
<tr>
<th>Parameters of LFTs and their normal values</th>
<th>Number of tests done</th>
<th>Number of tests in which post-op. values were found abnormal</th>
</tr>
</thead>
<tbody>
<tr>
<td>Bilirubin (0-1.2) mg%</td>
<td>41</td>
<td>7</td>
</tr>
<tr>
<td>SGPT (5-40) units/lit.</td>
<td>41</td>
<td>14</td>
</tr>
<tr>
<td>Alk. Phosphatase (15-69) units/lit.</td>
<td>41</td>
<td>8</td>
</tr>
<tr>
<td><strong>Total</strong></td>
<td><strong>123</strong></td>
<td><strong>29</strong></td>
</tr>
<tr>
<td><strong>%</strong></td>
<td><strong>23.58</strong></td>
<td></td>
</tr>
</tbody>
</table>

<https://example.com/table-II>
individuals who have previously been exposed to halothane. However, failure to produce similar
damage in animals, the rarity of hepatic damage in humans and the delayed appearance of hepatic
damage, suggests that halothane is not a direct hepatotoxin but is probably a sensitizing agent.\textsuperscript{12} It also
seems likely that the reduction of halothane through a minor metabolic pathway results in a toxic
metabolite that destroys the hepatocytes.\textsuperscript{12}
To reduce the frequency and severity of hepatotoxicity following the use of halothane, a minimum gap
of 3 months has been suggested between 2 halothane anaesthesias.

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