Introduction

The prevalence of gallstone disease is high in many European countries as well as in the USA (15). Serum lipids and lipoproteins were suggested to be related with the pathogenesis of gallstones (18).

Although there have been numerous studies evaluating the serum lipid status of gallstone patients, these studies yielded inconsistent results. For example, decreased (15) and increased (17) total cholesterol (TC) levels have been reported in gallstone patients. The highest gallstone formation has been reported to be observed at low HDL-C levels and high triglyceride levels (18). Conversely, an unchanged high density lipoprotein cholesterol (HDL-C) and triglyceride (TG), and reduced serum low density lipoprotein cholesterol (LDL-C) were reported in gallstone patients (17).

The aim of this study was to investigate the possible relationship between serum lipids, lipoproteins and gallstone disease, and compare it with previous studies.

Materials and Methods

In this study 32 hospitalised gallstone patients undergoing elective cholecystectomy operation (27 women, 5 men; 28–65 years; mean: 47.6 years) and 32 healthy donors (control group, 27 women and 5 men; 26–60 years; mean: 43.4 years) were allocated to the study. Those having diseases other than cholelithiasis were not included to the study in gallstone patients group. In gallstone group patients, there were no cholestasis, and the number of stones were two or more. There were no differences in demographic characteristics of patients between groups, including body mass index, and fasting blood sugar, as morbid obese patients, hypertensive patients and patients with fasting blood sugar of more than 120 g/dL were not included to the study.

Blood was obtained in the fasting state and serum samples were analysed for TC, HDL-C, TG, lipoprotein(a) (Lp(a)), apoprotein A (Apo-A), and apoprotein B (Apo-B) concentrations. The serum TC and TG concentrations were determined using an enzymatic method (UV test, Boehringer Mannheim) and by using auto-analyser (Hitachi 705). Lp(a), Apo-A, and Apo-B values were determined by nephelometry. LDL-C values were calculated by the Friedewald formula using TC, HDL-C and TG values obtained previously (4).

All the results are presented as mean value ± standard error (SE). The statistical analyses were performed using the analysis of variance (ANOVA).

Results

There was no difference in patient characteristics between groups. Mean (±SE) values for fasting serum lipid fractions (TC, HDL-C, LDL-C, TG) and the concentra-
malignant between groups, determinant factors of gallstone metabolism. A disturbance in the lipid and lipoprotein metabolism may increase the risk of developing cholelithiasis in certain subjects. It was reported that serum apolipoproteins might be more sensitive parameters than serum lipids in distinguishing patients with stones from those without stones (5, 19). Similar to these findings we have found increased concentrations of serum Apo-A, Apo-B, LDL-C, TC, and TG in gallstone patients. Stenos et al (6) reported that obese persons were at risk for cholesterol gallstones because their bile is saturated with calcium bilirubinate (12). This study confirms results of earlier studies of serum TG (7,15,18) and HDL-C (5,18) levels in gallstone patients.

Contradictory to some studies (17,15) and in accordance with others (5,7), we observed higher concentrations of TC and LDL-C in gallstone patients than did control subjects, although the difference for TC was not statistically significant.

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Data on the association between cholelithiasis and diabetes often are controversial and are mostly based on autopsies or on hospital series. However De Santis et al (2) reported that an altered glucose metabolism may increase the risk of developing cholelithiasis in certain subjects. Gebhard et al (6) reported that obese persons were at risk for cholesterol gallstones because their bile is saturated with cholesterol. Nevertheless, neither diabetic nor obese patients were included in our study to get the accurate results.

In conclusion, the results of this study may indicate that a disturbance in the lipid and lipoprotein metabolism may have a role in gallstone formation.

References


Discussion

It is a widely accepted idea that the gallstone formation is closely related to a disorder in the lipid and lipoprotein metabolism (1). It was reported in a study that the average levels of serum lipids in patients with gallstones were higher than subjects without gallstones (11). However many metabolic factors may affect the gallstone formation (13). In order to standardize the groups in this study hypertensive, atherosclerotic and morbid obese patients were not included to the study. Since other patient characteristics were similar between groups, determinant factors of gallstone formation in these patients seem to be related to the lipid metabolism.

Triglycerides are converted by fatty tissue lipase into free fatty acids and transported in the blood stream to the various organs where, in the presence of glucose, they are resynthesized to triglycerides. In the liver, a large part of the free fatty acids is utilized for the synthesis of the lipids, primarily phospholipids. When the supply of free fatty acids exhausts the metabolic capacity of the liver, as, for example, in fat-rich diets, triglycerides accumulate in the liver cells and may possibly be excreted in the bile. Free fatty acids also stimulate mucin hypersecretion in the gallbladder (12). This mucosal hypersecretion has been assigned a significant role in the formation of gallbladder stones (8). It is suggested that HDL-C might be a protective factor against gallstone formation in gallbladder (5). HDL-C has been reported to be negatively associated with gallbladder disease or the presence of gallstones (16). Also, Carel found the highest gallstone risk at low HDL-C levels and high TG levels (18). This study confirms results of earlier studies of serum TG (7,15,18) and HDL-C (5,18) levels in gallstone patients.

The values for LDL-C, TG, TC, Apo-A, and Apo-B in the gallstone patients were found to be higher than the values obtained in the control group, and these differences were statistically significant (p<0.05). On the other hand, HDL-C, Lp(a) levels were found to be lower in gallstone patients, but this difference was not statistically significant.

### Tab 1: Serum lipid concentrations in control and gallstone groups.

<table>
<thead>
<tr>
<th>Group</th>
<th>TC (mg/dL)</th>
<th>HDL-C (mg/dL)</th>
<th>LDL-C (mg/dL)</th>
<th>TG (mg/dL)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Control</td>
<td>185.60±19.3</td>
<td>48.40±7.8</td>
<td>95.35±13.7</td>
<td>136.40±23.2</td>
</tr>
<tr>
<td>Gallstone</td>
<td>209.23±11.6</td>
<td>43.35±3.6</td>
<td>132.70±8.0</td>
<td>186.91±9.4</td>
</tr>
</tbody>
</table>

TC indicates total cholesterol; LDL, low density lipoprotein; HDL, high density lipoprotein; TG, triglyceride. NS, Not significant.

### Tab 2: Serum lipoprotein concentrations in control and gallstone groups.

<table>
<thead>
<tr>
<th>Group</th>
<th>Lp(a) (mg/dL)</th>
<th>Apo-A (mg/dL)</th>
<th>Apo-B (mg/dL)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Control</td>
<td>46.72±13.6</td>
<td>151.46±8.3</td>
<td>91.12±8.3</td>
</tr>
<tr>
<td>Gallstone</td>
<td>42.63±7.2</td>
<td>158.94±9.4</td>
<td>98.60±6.2</td>
</tr>
</tbody>
</table>

Lp(a) indicates lipoprotein(a); Apo-A, apoprotein A; Apo-B, apoprotein B. NS, Not significant.


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