Lipid profile in Diabetes Mellitus

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ABSTRACT
Diabetes mellitus is the most common metabolic disorder affecting the people all over the world. Diabetes mellitus has been known to be associated with lipid disorders and cardiovascular complications. This study is planned to assess the lipaemic changes in diabetes mellitus patients attending the out-patient department of medicine department in HSK Hospital and Research Centre. Total Cholesterol (TC), Triglycerides (TG), LDL Cholesterol (LDL-C), HDL Cholesterol (HDL-C) levels were studied in serum of diabetes patients. This is a case control study which included 76 patients of diabetes as cases and 30 controls of the same age group and sex. All the samples were taken from subjects who fasted for at least 12 hours before the blood collection. The parameters were determined by manual method using spectrophotometer by liquid chemistry. The Triglycerides, Total cholesterol, LDL Cholesterol were higher in cases as compared to controls in both IDDM and NIDDM. The HDL-C was lower in NIDDM subjects who were on sulfonylurea or biguanides and was not significantly altered in IDDM patients on insulin. There was significant correlation between FBS and TC, TG, LDL-C and HDL-C in NIDDM patients. TC, TG, LDL-C showed significant correlation in IDDM subjects.

Key words: Diabetes mellitus, IDDM, NIDDM, lipid profile.

INTRODUCTION
Diabetes mellitus (DM) is a group of metabolic disease characterized by increase blood glucose level resulting from defects in insulin secretion, insulin action, or both1. The chronic hyperglycaemia of diabetes is associated with long-term damage, dysfunction and disturbance in failure of various organs, especially the eyes, kidneys, nerves, heart and blood vessels2. Patients with type-2 diabetes have increased risk of cardiovascular disease associated with atherogenic abnormalities and dyslipidaemia. Coronary artery disease, especially myocardial infarction is the leading cause of morbidity and mortality worldwide3. Hyperglycaemia and atherosclerosis are related in type-2 diabetes4.

Persistent hyperglycaemia causes glycosylation of all proteins, especially collagen cross linking and matrix proteins of arterial wall. This eventually causes endothelial cell dysfunction, contributing to atherosclerosis. The prevalence of dyslipidaemia in diabetes mellitus is 95%-5. Early detection and treatment of hyperlipidemia in diabetic patients reduces the risk for cardiovascular and cerebrovascular diseases. Lifestyle changes such as diet and exercise are very important in improving diabetic dyslipidemia, but often pharmacological therapy is needed6. Lipoprotein metabolism?

The rationale of the study was to detect the lipid abnormalities associated with chronic hyperglycaemia due to IDDM or NIDDM.

MATERIALS
This study was conducted in diabetic clinic, HSK Hospital and Research centre Bagalkot. A total number of 50 control who were healthy non smokers non alcoholics and at the time of study all of them were keeping good health and 76 diabetics who were on treatment were studied. The diabetics were on either sulfonylurea/biguanides or insulin treatment for type I or type II diabetes. In our study, we excluded diabetic subjects who were smokers, alcoholics and who were hypertensives, familial hyperlipidemia patients and patients with complications.

METHODS
1. Serum glucose estimation by Ortho-Toluidine method8.
5. Serum LDL-Cholesterol was calculated by Friedwald's Formula12.

These tests were done by manual methods using stat fax 3300 spectrophotometer by liquid chemistry.
Summary of metabolism of lipoproteins (Apoproteins-A, B, C, D, E; CII and E):
TG—Triglyceride; C—Cholesterol; P—Phospholipid; VLDL—Very low density lipoprotein;
IDL—Intermediate density lipoprotein; LDL—Low density lipoprotein; HDL—High density lipoprotein.

Metabolism of high density lipoproteins (P—Phospholipid; C—Cholesterol; CE—Cholesteryl ester; A, CII, E—Apoproteins; LCAT—Lechithin cholesterol acyltransferase).
RESULTS

Table 1: Showing Comparison of FBG, TC, TG, HDL-c, LDL-c Between Insulin Treated Diabetic subjects (IDDM and Control Subjects)

<table>
<thead>
<tr>
<th>Groups</th>
<th>FB Glucose Mg/dl</th>
<th>TC mg/dl</th>
<th>TG mg/dl</th>
<th>HDL-c Mg/dl</th>
<th>LDL-c Mg/dl</th>
</tr>
</thead>
<tbody>
<tr>
<td>Control</td>
<td>92.2±11.0</td>
<td>173.0±31.6</td>
<td>122.5±28.7</td>
<td>56.9±17.9</td>
<td>93.3±36.3</td>
</tr>
<tr>
<td>IDDM Subjects</td>
<td>229.8±46.2</td>
<td>222.2±23.3</td>
<td>197.9±49.7</td>
<td>58.4±14.6</td>
<td>124.2±25.0</td>
</tr>
<tr>
<td>p-value</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
<td>&gt;.05</td>
<td>&gt;.01</td>
</tr>
<tr>
<td>Statistical Significance</td>
<td>H. S</td>
<td>H. S</td>
<td>H. S</td>
<td>N. S</td>
<td>S. S</td>
</tr>
</tbody>
</table>

Table 2: Showing Comparison of FBG, TC, TG, HDL-c, LDL-c Between Diabetic (NIDDM) and Control Subjects.

<table>
<thead>
<tr>
<th>Groups</th>
<th>FB Glucose Mg/dl</th>
<th>TC mg/dl</th>
<th>TG mg/dl</th>
<th>HDL-c Mg/dl</th>
<th>LDL-c Mg/dl</th>
</tr>
</thead>
<tbody>
<tr>
<td>Control</td>
<td>92.2±11.0</td>
<td>173.0±31.6</td>
<td>122.5±28.7</td>
<td>56.9±17.9</td>
<td>93.3±36.3</td>
</tr>
<tr>
<td>IDDM Subjects</td>
<td>206.2±37.9</td>
<td>229.9±30.5</td>
<td>226.4±59.7</td>
<td>39.5±15.4</td>
<td>145.2±32.5</td>
</tr>
<tr>
<td>p-value</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Statistical Significance</td>
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<td>H. S</td>
<td>H. S</td>
<td>N. S</td>
<td>S. S</td>
</tr>
</tbody>
</table>

Table 3: Showing Comparison of FBG, TC, TG, HDL-c, LDL-c between control and Sulfonylurea Treated Diabetic subjects and also with Biguanide Treated Diabetics

<table>
<thead>
<tr>
<th>Groups</th>
<th>FB Glucose Mg/dl</th>
<th>TC mg/dl</th>
<th>TG mg/dl</th>
<th>HDL-c Mg/dl</th>
<th>LDL-c Mg/dl</th>
</tr>
</thead>
<tbody>
<tr>
<td>Control</td>
<td>92.2±11.0</td>
<td>173.0±31.6</td>
<td>122.5±28.7</td>
<td>56.9±17.9</td>
<td>93.3±36.3</td>
</tr>
<tr>
<td>IDDM Subjects</td>
<td>211.7±41.7</td>
<td>226.0±30.3</td>
<td>220.2±64.4</td>
<td>37.9±17.2</td>
<td>146.0±28.2</td>
</tr>
<tr>
<td>p-value</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Statistical Significance</td>
<td>H. S</td>
<td>H. S</td>
<td>H. S</td>
<td>H. S</td>
<td>H. S</td>
</tr>
</tbody>
</table>

1. The values are expressed as their mean ± S.D
2. H.S — highly significant, S. S-Statistically significant, N. S- not significant.

DISCUSSION

FASTING BLOOD GLUCOSE

The FBG levels in all the diabetics were highly significant (p<0.001) as compared to their respective controls\(^{13,14}\).

TOTAL CHOLESTEROL

Our study in IDDM and NIDDM are in accordance with earlier studies of John D Bagdade\(^ {15}\) and James M Falko\(^ {13}\). Diabetic state appears to be...
associated with increased synthesis of cholesterol. It has been hypothesized that hyperphagia of diabetes induces increased activity of HMG-CoA reductase of the intestine resulting in increased synthesis of cholesterol leading to raised levels in plasma. Dietary cholesterol also adds up to total cholesterol by increased absorption.

TRIGLYCERIDES

In our study the TG levels in IDDM as well as NIDDM on insulin as well as sulfonylurea are biguanide treated diabetics are raised and statistically highly significant. The hypertriglyceridemia may be due to higher rates of production of triglyceride rich VLDL by the liver and to decreased removal of TG by peripheral tissues-primarily adipose tissue and muscle. Insulin deficiency leads to high TG production and subsequent high packaging in VLDL. Several studies using radioactive substrates to trace the metabolism of plasma VLDL are consistent with their simultaneous overproduction and reduced clearance as the common etiologic mechanism for hypertriglyceridemia in poorly controlled IDDM. Furthermore, the structural composition of the VLDL itself may change with increase in protein components such as apolipoprotein C-III that inhibits the lipase enzyme and the uptake of VLDL remnants by the liver.

In NIDDM when TG are elevated above 200 mg/dl higher production rates of triglyceride and VLDL particles have been the most commonly identified metabolic abnormalities. Many hypertriglyceridemic NIDDM patients also appear to have defect in the clearance of triglyceride with lipoproteins.

HIGH DENSITY LIPOPROTEIN

The mean levels of HDL-C in our study in IDDM on insulin therapy are not statistically significant as compared to match controls which is in accordance the study of Kennedy and associates but not in variance with that reported by Nikkila et al. Celvert et al have reported low HDL-C in patients on sulfonylurea our study shows low HDL-C in both sulfonylurea and biguanide treated diabetics. Lower HDL-C in diabetes may be due to reduced Lipoprotein Lipase activity. The activity of cholesterol ester transfer protein is increased in IDDM. Hepatic TG lipase (HTGL) which lines the sinusoids of the liver which break TG added HDL is increased in the diabetic as an inversely correlated with HDL.

LOW DENSITY LIPOPROTEINS

Our study confirmed the stud. of Sosenko et al who have reported increase levels of LDL-C in IDDM. The mean LDL-c levels in total NIDDM subjects on sulfonylurea and patients on biguanides are increased (p<0.001) as compared to the matched controls. LDL production rates are reported to be elevated in IDDM but return to normal after insulin infusion. It may be due to increased synthesis of VLDL or impaired removal of VLDL remnant. Impaired receptor mediated clearance of LDL has also been postulated. In NIDDM there is alteration in the LDL lipid composition, and the LDL is enriched with triglycerides. LDL in patients with hypertriglyceridemia show decreased receptor binding to cultured skin fibroblasts, which may be the mechanism of increase in LDL-C in NIDDM.

CONCLUSION

This showed that every patient had at least one type of dyslipidemia. Overall diabetes mellitus is closely associated with dyslipidemia in both IDDM and NIDDM. It is mandatory to treat this dyslipidemia to prevent adverse lipemic status and long term complications to ensure a healthy and happy life inspite of diabetic dyslipidemia.

REFERENCES: