# Dyslipidemia and Type-2 Diabetics

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Objective: To determine the frequency of type-2 diabetics who have target lipoprotein blood levels and to study these levels in patients with ischemic heart disease and cardiovascular disease risk factors.

Design: Retrospective analytical study of hyperlipidemic type-2 diabetics between January 2000 and January 2001 was undertaken.

Setting: King AbdulAziz University Hospital.

Method: We studied hyperlipdemic type-2 diabetics who were on regular follow up to the medical outpatient clinic of King Abdulaziz University Hospital from January 2000 to January 2001. The mean lipoprotein levels, duration of hyperlipidemia and it's treatment, duration of diabetes, it's treatment and degree of blood glucose control, presence of hypertension, ischemic heart disease, and smoking were recorded.

Results: A total of 202 patients were studied with mean age of 60 years and equal male to female ratio. The mean duration of diabetes was 10 years and it was 7 years for hyperlipidemia. The mean level of LDL was 3.15 mmol/1 and it was 1.0 mmol/1 and 2.47 mmol/1 for LDL and TG respectively. Only 31% of patients had LDL<2.6 mmol/1, 28% had HDL>1.1 mmol/1 and 37% had TG <1.7 mmol/1. No significant difference was found in the frequency of target level of LDL in patients with IHD and those without; 26% versus 34% (0.4). Similarly, no difference was found in those with hypertension, obesity, and patients with family history of IHD compared to those without these risk factors; 30%, 26%, 16% versus 34%, 36%, 33% (p 0.2,0.1,0.4 respectively). Males were found to have a higher frequency of target LDL level compared to females; 38% versus 25% (p 0.04).

Conclusions: A low frequency of type-2 diabetics has target levels of lipoproteins and this was also evident in diabetics with IHD and CVD risk factors.

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Diabetes mellitus (DM) is one of the most common endocrine disorders. It is closely associated with coronary heart disease (CHD)¹. Diabetes is associated with two to four fold excess risk of CHD². Eighty percent of patients with type 2 diabetes will die of cardiovascular diseases³. The most common pattern of dyslipidemia in diabetics is elevated triglyceride (TG), low high density lipoprotein cholesterol (HDL), and predominance of small dense particle low density lipoprotein (LDL). Baseline data from the UKPD (United Kingdom Prospective Diabetes) Study showed that both low HDL and elevated LDL predicted CHD⁴. According to the American Diabetes Association (ADA) recommendations⁵; treatment goal for lipoprotein therapy in diabetics is as follow: LDL <2.6 mmol/l, TG <1.7 mmol/l. The aim of this study is to determine the frequency of type 2 diabetics who have target lipoprotein blood levels and to study these levels in

patients with ischemic heart disease and cardiovascular disease risk factors.

# METHOD

The study was carried out at King Abdulaziz University Hospital (KAUH), a teaching hospital in the western province of Kingdom of Saudi Arabia. During a one year period—January 2000 through January 2001, type 2 diabetics who were on regular follow up to the medical outpatient clinic were studied. Patients with dyslipidemia (defined as patient is known hyperlipidemic or has LDL >2.6 mmol/l, HDL <1.1 mmol/l, TG >1.7 mmol/l) were included in the study. For those who were recently discovered, they were reassessed after 6 months period. The mean lipoprotein levels of the last two visits were calculated and the duration of hyperlipidemia and type of

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treatment (diet, statin, fibrates, or combined) were recorded. Data were collected from the duration of DM, type of treatment, degree of blood glucose control (well controlled defined as H1c <7%, fasting <7 mmol/l, post-prandial <9 mmol/l), presence of hypertension (patient is known or having blood pressure >140/90 mmHg), its duration, presence of ischemic heart disease (IHD) (assessed by patients history or changes on electrophysiological studies), smoking (whether active or passive) and mortality.

The frequency of target lipoprotein level were studied as well as its level in patients with (CHD) and (CHD) risk factors. Statistical analysis was performed using the SPSS software. Mean SD was determined for quantitative data, and frequency for categorical variables. Chi-square was used to analyze group difference for categorical variables. For continuous variables t-test were used if comparing two groups. P value <0.05 was considered significant.

### RESULTS

A total of 202 patients were included in the study. The mean age was  $59.9\pm12.9$  years with equal male to female ratio and mean duration of diabetes was  $10.3\pm7$  years. Most of the patients were using oral hypoglycemic agents for blood glucose control followed by insulin and diet; 121/202 (60%), 59/202 (29%), 22/202 (11%) respectively. The majority of patients, 190/202 (94%), had poor blood glucose control. The mean duration of hyperlipidemia was  $6.6\pm1.7$  years. Hypertension, obesity and IHD were found with high frequency in the study group (Table 1). A low frequency of diabetics has goal of lipoprotein levels ie, only 31% had goal LDL, 28% had goal HDL and 37% had goal triglyceride level(Table 2). As shown in Table 3; both patients with CHD, CHD risk factors and those without have poor target lipoprotein levels. Males are more likely to have target LDL compared to females. A significant relation was found between poor lipid control and long duration of hyperlipidemia; mean duration of hyperlipidemia in patients with poor control was  $6.9\pm1.8$  years versus  $5.2\pm1.4$  years in those with good control (p=0.004). Patients with poor glycemic control has poor Tigo (37%) goal HDL versus 130/190 (68%), Patients with poor blood glucose control has goal level of LDL, 70/190 (37%) goal TG, 52/190 (27%) goal HDL versus 130/190 (68%) has low HDL levels respectively (p <0.001, 0.02, 0.001 respectively) Most of patients, 112/202 (55.4%), were not taking pharmacological treatment for lipid control while those on statin were 66/202(32.7%), on fibrates 18/202(8.9%) and on combined statin and fibrates 6/202(3.9%).

Table 1. Some characteristics of the study group

| Variable   | Total | Number=202<br>N.(%) |
|--|-------|---------------------|
| Ischemic heart disease   |       | 81(40)              |
| Hypertension   |       | 140(69)             |
| Duration of hypertension in<br>years (mean±SD)<br>Body mass index in |       | 8.25±6.8            |
| Kg/m² (mean± SD)   |       | $29.3 \pm 6.3$      |
| <30 Kg/m²  |       | 121 (60)            |
| >30 Kg/m <sup>2</sup>  |       | 81(40)              |
| Smoking  |       | 37(18)              |
| Family history of ischemic   |       |                     |
| heart disease  |       | 38(19)              |
| Mortality  |       | 17(8)               |

Table 2. Lipoproteins blood levels

| Lipoprotein type   | Lipoprotein level<br>N(%)   |  |  |
|--|---|--|--|
| LDL in mmol/l (mean±SD)  <2.6 mmol/l  2.6 -3.4 mmol/l  3.4 mmol/l  HDL in mmol/l (mean ±SD)  >1.1 mmol/l  Triglyceride in mmol/l (mean±SD)  <1.7 mmol/l  >1.7 mmol/l | 3.15 ±1.9<br>62(3)<br>60(30)<br>80(39)<br>±0.29<br>57(28)<br>145(72)<br>2.47±1.9<br>75(37)<br>127(63) |  |  |

Table 3. Relation of target lipoprotein level to ischemic heart disease and cardiovascular disease risk factors

| Variables                        | LDL<br>N (%)     | P    | HDL<br>N(%)      | P    | TG<br>N(%)       | P             |  |
|----------------------------------|------------------|------|------------------|------|------------------|---------------|--|
| IHD (N=81)<br>No IHD (N=121)     | 23(28)<br>41(34) | 0.4  | 16(20)<br>41(34) | 0.02 | 27(33)<br>48(40) | 0.3           |  |
| Hypertension<br>(N=140)          | 42(30)           | 0.2  | 36(26)           | 0.4  | 49(34)           | 0.2           |  |
| No hypertension                  | 21(34)           |      | 20(33)           |      | 25(42)           |               |  |
| (N=62)<br>Obese (N=81)           | 21(26)           | 0.1  | 24(30)           | 0.7  | 23(28)           | 0.03          |  |
| Non-obese<br>(N=121)             | 43(36)           |      | 33(27)           |      | 52(43)           |               |  |
| Smokers (N=37)                   | 6(16)            |      | 9(24)            | 0.5  |                  | 6(16) < 0.001 |  |
| Non-smokers<br>(N=165)           | 58(35)           |      | 48(29)           |      | 69(42)           |               |  |
| Family history<br>of IHD (N=38)  | 10(26)           | 0.4  | 10(26)           | 0.7  | 10(26)           | 0.1           |  |
| No family history<br>(N=164)     | 54(33)           |      | 47(29)           |      | 65(40)           |               |  |
| Males (N=102)<br>Females (N=100) | 39(38)           | 0.04 | 25(25)<br>32(32) | 0.2  | 38(37)<br>37(37) | 0.9           |  |

Target LDL <2.6 mmol/l, target HDL >1.1 mmol/l, target TG <1.7 IHD = ischemic heart disease

### DISCUSSION

Diabetes is rapidly becoming a major public health problem worldwide<sup>2</sup>. A study conducted by Mokdad et al<sup>6</sup> detected 33% increase in the prevalence of diabetes in adults across all age-groups, races, education level, weight level, and level of smoking over 8 year period (1990-1998). Type 2 diabetes is a progressive disease and it is an independent risk factor for CHD. Patients with diabetes and no previous history of IHD have the same risk for cardiac events as patients with previous myocardial infarction<sup>1</sup>. It is also associated with a combination of CVD risk factors including; hypertension, high LDL, low HDL, high TG and abdominal obesity, which are primarily attributed to insulin resistance<sup>7,9</sup>. Patients with type 2 diabetes have an increased prevalence of lipid abnormalities that contribute to high rate of CVD. The Framingham offspring study had shown that there is twice the prevalence of low HDL, high TG, high LDL in diabetics compared to non-diabetics <sup>11,2</sup>. Clinical research has found that these lipid abnormalities to be an independent risk factor for CHD in diabetics <sup>11,2</sup>. It is clear in our study that minority of diabetics had goal lipoprotein level, where only 31% had goal LDL level. Similar result had been reported by Saaddine et al<sup>13</sup> and others<sup>14,6</sup>. High LDL level was evident in diabetics with CHD risk factors where they are supposed to have better care because of their high risk. Poor lipid control was associated with long

duration of hyperlipidemia and poor glucose control. This could be due to the patients being fed up from chronic use of multiple medications. Diabetes itself often requires 2 or 3 separate medications each day, without the addition of additional disease treatments. It is easy for us as physicians to tell a patient to take 5, 6, or 7 different pills us to 3 threat times and day, but for the actions it will be up to 3 three times per day, but for the patient it will be boring. Another reason could be financial; patient can't afford buying multiple medications. An interesting observation is better lipid control in males compared to females, which is also observed in other studies<sup>13,14</sup>, whether it is related to poor quality of care in diabetic females or other reasons, this needs to be studied. Studies had also other reasons, this needs to be studied. Studies had also showed a lower risk of death attributed to IHD in diabetic males compared to females! Lipid management had been shown to reduce the risk of CHD by 25% to 55% and the risk of death by 43% [8,19]. Almost half of the patients were not on any pharmacological theory and the risk of the patients were risk of death by 43%. Almost hair of the patients were not on any pharmacological therapy and statins were the most frequently used medication. In three secondary prevention studies using statins, diabetics achieved significant reduction in coronary events<sup>19-21</sup>. A primary prevention study also using statins showed similar trend of reduced events. The Helsinki Heart Study, the Veterans Affairs High-Density Lipoprotein Cholesterol Intervention Trial (VA-HIT) also showed reduction in CHD events using

#### CONCLUSION

It is clear from this study and other studies that lipid control is sub-optimal in diabetics. There is a gap between ADA recommendations and their clinical application. Patients with diabetes often lack sufficient knowledge about their disease and its complications. VHA (Veterans Health Administration) for example recommend reassessing patient's knowledge about diabetes at least three month after educational intervention25. Programs to improve diabetic's knowledge about the importance of controlling hyperlipidemia will allow them for better contribution to their care and the benefit of these programs should be reassessed. Also reinforcement and encouragement of physicians for better follow up and more aggressive management of hyperlipidemia is warranted.

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