Serum Lipids and Diabetic Retinopathy in Newly Diagnosed Type 2 Diabetic Subjects

Md. Rezwanur Rahman¹, M Iqbal Arslan², Md. Mozammel Hoque³, Forhadul Hoque Mollah⁴, Shahana Shermin⁵

Abstract

Background: Diabetic retinopathy is the commonest and usually the first observable vascular complication of diabetes mellitus. Along with hyperglycaemia, dyslipidaemia is a contributing factor for the occurrence of diabetic retinopathy. It is postulated that dyslipidaemia results in formation of hard exudate by increasing blood viscosity and altering the fibrinolytic system. A case control study was carried out in the department of Biochemistry, Bangabandhu Sheikh Mujib Medical University, Dhaka during the period of January 2006 to December 2007 to evaluate the serum lipid profile in newly diagnosed type 2 diabetic subjects with diabetic retinopathy. Materials and Methods: Total 85 newly diagnosed type 2 diabetic subjects were included in this study, 40 were cases having retinopathy and 45 were age and sex matched controls without retinopathy. Serum triglyceride (TG), total cholesterol (TC), low density lipoprotein cholesterol (LDL-C) and high density lipoprotein cholesterol (HDL-C) were compared between cases and controls. Unpaired t-test and chi-square test were done between groups as tests of significance. **Results:** All the parameters of lipid profile showed dyslipidaemic trend both in cases and controls. In the cases TG was significantly higher and HDL-C was significantly lower than that of controls (p < 0.05) whereas no significant difference was found between cases and controls with respect to serum TC and LDL-C. **Conclusion:** It can be concluded that high TG and low HDL-C are associated with diabetic retinopathy in newly diagnosed type 2 diabetes.

Keywords: Lipid profile, Dyslipidaemia, Diabetic retinopathy

J Enam Med Col 2011; 1(2): 63-66

Introduction

Diabetes mellitus (DM) is one of the common chronic disorders worldwide with a number of irreversible complications and diabetic retinopathy is usually the first observable vascular complication in DM.^{1,2} Diabetic retinopathy is a well characterized, sight threatening, chronic, ocular disorder that eventually develops to some degree in nearly all patients with diabetes mellitus.³ At the time of diagnosis of type 2 diabetes, about 21% patients have established retinopathy. Almost all patients with type 1 diabetes and >60% of patients with type 2 diabetes develop retinopathy within 20 years of diagnosis.⁴

Several decades of basic and clinical research have established that though hyperglycaemia is a major

Correspondence Md. Rezwanur Rahman, Email: rezwanrahman5@yahoo.com

^{1.} Associate Professor, Department of Biochemistry, Delta Medical College, Dhaka

^{2.} Professor, Department of Biochemistry, BSMMU, Dhaka

^{3.} Professor, Department of Biochemistry, BSMMU, Dhaka

^{4.} Associate Professor, Department of Biochemistry, BSMMU, Dhaka

^{5.} Senior Medical Officer, Department of Gynaecology & Obstetrics, BIRDEM Hospital, Dhaka

causative factor in the development of diabetic retinopathy, there are also some other factors that accelerate its development and progression. Among the contributing factors, diabetic dyslipidaemia characterized by elevated triglycerides (TG), low levels of HDL cholesterol (HDL-C), and increased presence of small, dense LDL particles, has been proposed as a probable risk factor of diabetic retinopathy.⁴⁻⁶

The mechanism by which serum lipids may cause the progression of diabetic retinopathy is not clearly understood. It has been postulated that elevation of blood viscosity and alterations in the fibrinolytic system occur in dyslipidaemia causing hard exudate formation. There may also be incorporation of triglycerides into the cell membrane leading to changes in membrane fluidity and leakage of plasma constituents into the retina. This results in haemorrhage and oedema in the retina.⁷ Epidemiological studies reveal a strong inverse relation between HDL-C levels and atherogenicity.⁸

We conducted this study in our population with the aim to compare total cholesterol (TC), triglycerides (TG), HDL cholesterol (HDL-C) and LDL cholesterol (LDL-C) concentrations between newly diagnosed type 2 DM patients with retinopathy and newly diagnosed type 2 DM patients without retinopathy to determine whether dyslipidaemia has any role in the occurrence of this sight threatening disorder of diabetes mellitus.

Materials and Methods

This case control study was carried out in the department of Biochemistry, Bangabandhu Sheikh Muiib Medical University (BSMMU), Dhaka during January 2006 to December 2007. A total of 85 newly diagnosed type 2 diabetic patients were included purposively on the basis of predefined inclusion and exclusion criteria with special attention to having no history of taking lipid lowering drugs. Study subjects were categorized into cases (40 in number) having retinopathy and age and sex matched controls without retinopathy (45 in number). With ethical considerations and aseptic precaution, 5 mL fasting venous blood was collected from the study subjects. Serum TC, TG and HDL-C were measured by CHOD-PAP⁹, GPO¹⁰ and precipitant¹¹ methods respectively. Serum LDL-C was calculated by Friedewald's formula¹² and subjects having TG > 400 mg/dL were excluded from the study. Diagnosis of diabetic retinopathy was done by ophthalmoscopy. All the data were recorded in a preformed data collection sheet and were analyzed by using SPSS 12.0 for Windows. Unpaired t-test and chi-square test were done between groups as tests of significance.

Results

Among the 85 newly diagnosed type 2 diabetes mellitus patients 40 had nonproliferative retinopathy designated as cases with mean age of 44.65 ± 8.80 years (29 to 65 years). The remaining 45 were without retinopathy regarded as controls with mean age of 44.04 ± 9.09 years (30 to 65 years). Among the cases 17 were male and 23 were female whereas 20 were male and 25 were female among the controls. No significant difference (p > 0.05) was observed regarding age and sex distribution between cases and controls (Table I).

Table I: Distribution and comparison of age and sex between cases and controls

| Study | Mean age | t/n | Sex | | |
|--------------------|---|------------------|------|--------|------------------|
| subjects | (yrs) | νp | Male | Female | χ-/p |
| Case $(n = 40)$ | $\begin{array}{c} 44.65 \pm 8.80 \\ (2965) \end{array}$ | 0.756/ > 0.05 | 17 | 23 | 0.033/ > 0.05 |
| Control $(n = 45)$ | 44.04 ± 9.09 (30-65) | | 20 | 25 | |

Table II shows the comparison of lipid profile parameters between cases and controls. In the cases TG was significantly higher and HDL-C was significantly lower than those of controls (p < 0.05) whereas no significant difference was found between cases and controls with respect to TC and LDL-C.

Table II: Comparison of lipid profile between cases and controls

| Parameters (in mg/dL) | $Cases Mean \pm SD (n = 40)$ | Controls Mean \pm SD (n = 45) | t value | p value |
|--------------------------|------------------------------|---------------------------------------|---------|---------|
| TC | 210.48 ± 31.91 | 206.60 ± 33.56 | 0.544 | > 0.05 |
| | (155–273) | (150 - 268) | | |
| TG | 205.30 ± 50.64 | 178.96 ± 41.55 | 2.633 | < 0.05 |
| | (134–360) | (110 - 310) | | |
| HDL -C | 36.42 ± 7.72 | 40.82 ± 7.57 | -02.649 | < 0.05 |
| | (22–53) | (24-60) | | |
| LDL -C | 133.18 ± 28.71 | 130.16 ± 30.55 | 0.468 | > 0.05 |
| | (84-197) | (74 - 187) | | |

Discussion

This case control study was designed to evaluate the components of lipid profile in diabetic retinopathy in newly diagnosed type 2 diabetes mellitus subjects. We have compared serum TC, TG, HDL-C and LDL-C levels between 40 newly diagnosed type 2 diabetes mellitus cases with nonproliferative retinopathy and age and sex matched 45 controls of newly diagnosed type 2 diabetes mellitus patients without retinopathy.

We have found raised TC, TG and LDL-C and lower HDL-C concentrations in both the study groups with notable dyslipidaemic pattern in cases. Several discrepant findings were observed in different studies. In our study the concentration of TG was significantly higher in cases compared to controls and no significant difference in TC and LDL-C concentrations was observed between the groups, which is partly supported by the studies done by Kareem et al¹³ and Rema et al¹⁴ where the former group found higher TC and TG concentrations and the latter found significantly higher TC, TG and LDL-C concentrations. Our finding of no significant difference in TC and LDL-C concentrations between the groups is in accordance with that of several other studies.¹⁵⁻¹⁹ This study observed lower HDL-C concentration in both the study groups compared to reference value and comparison revealed statistically significant lower level in cases. Similar finding was observed by Kordonouri et al²⁰ and Sinav et al²¹ but the studies done by van Leiden HA et al²² and Wong et al²³ differ from our findings. These conflicting findings regarding different components of lipid profile may be due to different dietary habit, life style and ethnicity of our study subjects than that of the studies done abroad. Sample size might also play a role.

It may be concluded that hypertriglyceridaemia and low HDL-C are associated with diabetic retinopathy in newly diagnosed type 2 diabetes mellitus. Subjects with diabetes and altered lipid levels should undergo more frequent ophthalmologic examinations and appropriate lipid lowering therapy in an early phase to prevent other atherosclerosis related morbidity and mortality, and also to contribute to the prevention of retinopathy.

References

- Donnelly R, Emslie-Smith AM, Gardner ID, Morris AD. ABC of arterial and venous disease. Vascular complications of diabetes. BMJ 2000; 320: 1062-1066.
- Harris MI, Klein R, Welborn TA, Knuiman MW. Onset of NIDDM occurs at least 4–7 yrs before clinical diagnosis. Diabetes Care 1992; 15: 815-819.
- 3. Fong DS, Aiello LP, Ferris FL, Klein R. Diabetic Retinopathy. Diabetes Care 2004; 27(10): 2540-2553.
- Lydic TA, Reid GE, Esselman WJ, Busik JV. Dyslipidemia and diabetic retinopathy. mass spectrometry-based lipidomic analysis of rat and human retina. ARVO annual meeting abstract 2007.
- Miccoli R, Odello G, Giampietro O, Marchetti P, Cristofani R, Penno G, Meucci G, Navalesi R. Circulating lipid levels and severity of diabetic retinopathy in type I diabetes mellitus. Ophthalmic Research 1987; 19(1): 52-56.
- Leung H, Wang JJ, Rochtchina E, Wong TY, Klein R, Mitchell P. Dyslipidaemia and microvascular disease in the retina. Eye 2005; 19: 861-868.
- Barham D, Trinder P. An improved colour reagent for the determination of blood glucose by oxidase system. Analyst 1972; 97: 142-145.
- Fernandez ML, Jones JJ, Ackerman D, Barona J, Calle M, Comperatore MV, Kim JE, Anderson C, Leite JO, Volek JS, McIntosh M, Kalynych C, Najm W, Lerman RH. Low HDL cholesterol is associated with increased atherogenic lipoproteins and insulin resistance in women classified with metabolic syndrome. Nutr Res Pract 2010; 4(6): 492-498.
- Allain CC, Poon LS, Chan CSG, Richmond W, Fu PC. Enzymatic determination of total serum cholesterol. Clin Chem 1974; 20(4): 470-475.
- Bucolo G, David H. Quantitative determination of serum triglycerides by the use of enzymes. Clin Chem 1973; 19(5): 476-482.
- Lopes-Virella MF, Stone P, Ellis S, Colwell JA. Cholesterol determination in high-density lipoproteins separated by three different methods. Clin Chem 1977; 23(5): 882-884.
- Friedewald WT, Levy RI, Fredrickson DS. Estimation of the concentration of low density lipoprotein cholesterol in plasma, without the use of the preparative ultracentrifuge. Clin Chem 1972; 18(6): 499-502.
- Kareem I, Jaweed SA, Bardapurkar JS, Patil VP. Study of magnesium, glycosylated hemoglobin and lipid profile in diabetic retinopathy. Indian Journal of Clinical Biochemistry 2004; 19(2): 124-127.

- Rema M, Srivastava BK, Anitha B, Deepa R, Mohan V. Association of serum lipids with diabetic retinopathy in urban South Indians—the Chennai Urban Rural Epidemiology Study (CURES) Eye Study-2. Diabetic Medicine 2006; 23(9): 1029-1036.
- Dhir SP, Rajvir D, Jagat R, Dash RJ, Chakravarti RN. Serum lipoprotein cholesterol profile in diabetic retinopathy. Indian J Ophthalmol 1984; 32(2): 89-91.
- Rema M, Deepa R, Mohan V. Prevalence of retinopathy at diagnosis among type 2 diabetic patients attending a diabetic centre in South India. Br J Ophthalmol 2000; 84: 1058-1060.
- 17. Tarkun I, Arslan BC, Cantürk Z, Tarkun P, Kozdag G, Topsever P. Homocysteine concentrations in type 2 diabetes mellitus patients without cardiovascular disease: relationship to metabolic parameters and diabetic complications. Turkish Journal of Endocrinology and Metabolism 2003; 7(1): 11-17.
- Parvanova A, Iliev I, Filipponi M, Dimitrov BD, Vedovato M, Tiengo A, Trevisan R, Remuzzi G, Ruggenenti P. Insulin resistance and proliferative retinopathy: a cross-sectional, case-control study in 115 patients with type 2 diabetes. J Clin Endocrinol Metab 2004; 89(9): 4371-4376.

- 19. Nayak BS, Roberts L. Relationship between inflammatory markers, metabolic and anthropometric variables in the Caribbean type 2 diabetic patients with and without microvascular complications. J Inflamm (Lond) 2006; 3: 17.
- 20. Kordonouri O, Danne T, Hopfenmüller W, Enders I, Hövener G, Weber B. Lipid profiles and blood pressure: are they risk factors for the development of early background retinopathy and incipient nephropathy in children with insulin-dependent diabetes mellitus? Acta Paediatrica 1996; 85(1): 43-48.
- Sinav S, Onelge MA, Onelge S, Sinav B. Plasma lipids and lipoproteins in retinopathy of type I (insulindependent) diabetic patients. Ann Ophthalmology 1993; 25(2): 64-66.
- 22. van Leiden HA, Dekker JM, Moll AC, Nijpels G, Heine RJ, Bouter LM. Blood pressure, lipids, and obesity are associated with retinopathy: the Hoorn study. Diabetes Care 2002; 25: 1320-1325.
- Wong TY, Klein R, Sharrett AR, Manolio TA, Hubbard LD, Marino EK. The prevalence and risk factors of retinal microvascular abnormalities in older persons: The cardiovascular health study. Ophthalmology 2003; 110: 658-666.