

## Prevalence of Retinopathy and Its Associated Factors in Type-2 Diabetes Mellitus Patients Visiting Hospitals and Diabetic Clinics in Faisalabad, Pakistan

Muhammad Hassan, Mobeen Akhtar\* and Naeem Akhtar

Department of Zoology, G.C. University, Faisalabad (MH, MA) and Allied Hospital, Punjab Medical College Faisalabad (NA)

**Abstract.-** Five hundred patients, visiting the government hospitals and diabetic clinics at Faisalabad were examined for retinopathy and their blood samples were analyzed for HbA<sub>1c</sub> and lipid profile. Retinopathy was found 41.4% and its prevalence was found higher in patients with HbA<sub>1c</sub> >8%. Besides that the age of patients, duration of diabetes, history of hypertension and smoking status were found to be significantly (P<0.05) associated with the prevalence of retinopathy in the diabetic patients. It was concluded that the prevalence of diabetic retinopathy was higher in patients with poor glycaemic control, duration of diabetes and history of hypertension.

**Key words:** Retinopathy, glycosylated haemoglobin, HbA<sub>1c</sub>, diabetes mellitus type-2.

### INTRODUCTION

**D**iabetes mellitus (DM) is a metabolic syndrome characterized by hyperglycaemia due to absolute or relative deficiency of insulin. Lack of insulin affects the metabolism of carbohydrate, protein and fat, and causes a significant disturbance of water and electrolyte homeostasis. Death may result from acute metabolic decompensation, while long-standing metabolic derangement is frequently associated with permanent and irreversible functional and structural changes in the cell of the body, with those of the vascular system being particularly susceptible. It damages the vessels and the basement membranes, causing impaired delivery of nutrients and hormones to the tissues, resulting in tissues damage. The most affected sites are the retina, renal glomerulus and the nerve sheath (Edwards *et al.*, 1991).

Diabetic retinopathy is the most common cause of legal blindness in adults under the age of 65 in most countries (NIH, 1985). Insulin deficiency results in the elevated blood glucose level that is accompanied by and causes structural, physiologic and hormonal changes that affect retinal capillaries, causing the capillaries to become functionally less competent (Shimizu *et al.*, 1981; Niki *et al.*, 1984;

Bresnick, 1989).

The incidence of retinopathy increases alongwith the duration of diabetes and 60 percent of patients with type-2 diabetes have retinal changes after 20 years of the disease. The first signs of retinopathy are microaneurysms (minimal retinopathy), which are followed by haemorrhages and lipid exudates (background retinopathy), cotton-wool spots and venous reduplication (proliferative retinopathy). Non-enzymatic glycation and changes in growth factor profiles cause deposition of further extracellular matrix material along the basement membranes of small vessels. Also, an excess of glucose activates the polyol pathway which causes accumulation of sorbitol in the lens and is accompanied by cataracts (Esperance *et al.*, 1990; Kinoshita *et al.*, 1990). The present paper describes the prevalence of retinopathy and its associated factors in type-2 diabetes mellitus as determined in 500 patients visiting Government hospitals and private clinics at Faisalabad, Pakistan.

### MATERIALS AND METHODS

A total of 500 patients with type-2 diabetes belonging to different socio-economic groups and attending the Government Hospitals (District Headquarter Hospital and Allied Hospital) and private clinics in Peoples Colony and Hajwary Town in Faisalabad for a follow-up visit during the study period from November, 2004 to December, 2005 were

\* Corresponding author: [mianmobeenakhtar@gmail.com](mailto:mianmobeenakhtar@gmail.com)  
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included in this study.

Blood samples from all the diabetic patients with type-2 were analysed for glycosylated Haemoglobin (HbA<sub>1c</sub>) according to Bisse and Abraham (1985), cholesterol, according to Artiss and Zak (1997), triglycerides according to Cole *et al.* (1997), high-density lipoprotein cholesterol (HDL-C) according to Burstein *et al.* (1970) and low density lipoprotein cholesterol (LDL-C) according to Friedewald *et al.* (1972).

#### Patient's medical examination

All the diabetic patients included in this study were examined by registered practitioners for retinopathy. Other details such as gender, age, education, and socio-economic status, family history of diabetes, duration of diabetes, smoking status, blood pressure, weight, height, exercise status, waist girth and hip girth of all diabetic patients were recorded on a structural proforma. The data was subjected to statistical analysis (chi-square test) in order to determine the relationship among the various risk factors and complications of diabetes.

## RESULTS AND DISCUSSION

Out of the 500 diabetic patients, 207 (41.4%) were observed with complication of retinopathy, while 293 (58.6%) showed no indication of retinopathy. Table I shows characteristics of all 500 persons suffering from diabetes mellitus type-2 included in this study. Table II shows results of biochemical tests of all diabetic patients included in this study. Prevalence of retinopathy in all diabetic patients included in this study with reference to different associated factors is shown in Table III. Results of chi-square test showed that age of patient, duration of diabetes, history of hypertension and smoking status are significantly ( $P < 0.05$ ) associated with the prevalence of retinopathy in diabetic patients.

#### Sex wise prevalence

In the present study, a high sex specific prevalence was reported in females in type-2 diabetes mellitus. Within type-2 diabetes mellitus 43.2% were males and 56.8% were females. In

**Table I.- Characteristics of 500 persons with type-2 diabetes mellitus.**

| Variable                          | Total (n=500) |      |
|-----------------------------------|---------------|------|
|                                   | No.           | %    |
| Sex                               |               |      |
| Female                            | 284           | 56.8 |
| Male                              |               |      |
| Age (Years)                       |               |      |
| ≤49                               | 145           | 29.0 |
| 50-59                             | 187           | 37.4 |
| 60-69                             | 137           | 27.4 |
| ≥70                               | 31            | 6.2  |
| Education                         |               |      |
| Gragaute/Postgraduate             | 88            | 17.6 |
| Higher Secondary                  | 84            | 16.8 |
| Secondary                         | 91            | 18.2 |
| Primary                           | 59            | 11.8 |
| Illiterate                        | 178           | 35.6 |
| Monthly income (Pakistani Rupees) |               |      |
| >10,000                           | 90            | 18.0 |
| 5,000-10,000                      | 115           | 23   |
| <5,000                            | 181           | 36.2 |
| Not mentioned                     | 114           | 22.8 |
| Duration of diabetes              |               |      |
| 1-5 years                         | 206           | 41.2 |
| >5 years                          | 294           | 58.8 |
| History of smoking                |               |      |
| Never smoked                      | 418           | 83.6 |
| Past smoker                       | 38            | 7.6  |
| Current smoker                    | 44            | 8.8  |
| Exercise status                   |               |      |
| Regular                           | 173           | 34.6 |
| Irregular                         | 327           | 65.4 |
| History of hypertension           |               |      |
| Yes                               | 196           | 39.5 |
| No                                | 304           | 60.8 |
| Body mass index (BMI)             |               |      |
| ≥25                               | 331           | 66.2 |
| <25                               | 169           | 33.8 |
| Waist to hip ratio (WHR)          |               |      |
| More                              | 417           | 83.4 |
| Less                              | 83            | 16.0 |

agreement with the present study, previously Shera *et al.* (1999) have estimated that the type-2 diabetes was more in females (11.1%) than in males (9.4%) in

**Table II.- Biochemical tests of 500 diabetic patients.**

|                              | No. of patients | Percent |
|------------------------------|-----------------|---------|
| HbA1c                        |                 |         |
| Good glycemic control <7%    | 248             | 49.6    |
| Acceptable 7-8%              | 93              | 18.6    |
| Poor glycemic control >8%    | 159             | 31.8    |
| Cholesterol                  |                 |         |
| Desireable ≤200 mg/dl        | 119             | 23.8    |
| Borderline 200-240mg/dl      | 136             | 27.2    |
| High risk >240 mg/dl         | 245             | 49.0    |
| Total triglycerides          |                 |         |
| Desireable ≤200 mg/dl        | 121             | 24.2    |
| Borderline 200-240mg/dl      | 114             | 22.8    |
| Elevated/high risk>240 mg/dl | 265             | 53.0    |
| HDL-cholesterol              |                 |         |
| Desireable/normal >40 mg/dl  | 61              | 12.2    |
| Borderline 30-40mg/dl        | 54              | 10.8    |
| High risk >30 mg/dl          | 385             | 77.0    |
| LDL-cholesterol              |                 |         |
| Desireable ≤130 mg/dl        | 128             | 25.6    |
| Borderline 130-160mg/dl      | 79              | 15.8    |
| High risk <240 mg/dl         | 293             | 58.6    |

the rural areas of NWFP. Rosenbloom *et al.* (1998) also reported that females were more frequently affected than the males by non insulin dependent diabetes mellitus (NIDDM) in Native American and African-American youth. Female preponderance in prevalence of diabetes mellitus in Pakistani patients has also been reported previously (Shera *et al.*, 1999b). In a survey of consecutive households in a relatively prosperous and a poor area in Karachi, it was documented that diabetes was more common in females in both populations (Hameed *et al.*, 1995).

#### Age wise prevalence

Diabetes is known to be increasing in prevalence and incidence among the elderly (Walker *et al.*, 1997, Black *et al.*, 1999, Allessandro *et al.*, 1999). In the present study, it was found that diabetes mellitus type-2 was more prevalent in age group 50-59 years followed by patients of age group ≤49. Least prevalence was found in patients with age 70 years or more. It is due to the fact that diabetic patients rarely reach upto te age of 70 years or more. Due to

**Table III.- Prevalence of retinopathy in 500 diabetic patients and associated risk factors.**

| Vaiables                 | Complication present (n=207) |      | Complication not present (n=293) |      |
|--------------------------|------------------------------|------|----------------------------------|------|
|                          | No.                          | %    | No.                              | %    |
| Age in years             |                              |      |                                  |      |
| ≤49                      | 39                           | 18.8 | 106                              | 36.2 |
| 50-59                    | 77                           | 37.2 | 110                              | 37.5 |
| 60-69                    | 65                           | 31.4 | 72                               | 24.6 |
| ≥70                      | 26                           | 12.6 | 5                                | 1.7  |
| Duration of diabetes     |                              |      |                                  |      |
| 1-5                      | 70                           | 33.8 | 136                              | 46.4 |
| >5 years                 | 137                          | 66.2 | 157                              | 53.6 |
| Exercise status          |                              |      |                                  |      |
| Regular                  | 76                           | 36.7 | 97                               | 33.1 |
| Irregular                | 131                          | 63.3 | 196                              | 66.9 |
| Hypertension             |                              |      |                                  |      |
| Yes                      | 131                          | 63.3 | 65                               | 22.2 |
| No                       | 76                           | 36.7 | 228                              | 77.8 |
| Body mass index BMI      |                              |      |                                  |      |
| ≥25                      | 134                          | 64.7 | 197                              | 67.2 |
| < 25                     | 73                           | 35.3 | 96                               | 32.8 |
| Waist to hip ratio (WHR) |                              |      |                                  |      |
| More                     | 172                          | 83.1 | 245                              | 83.6 |
| Less                     | 35                           | 16.9 | 48                               | 16.4 |
| History of smoking       |                              |      |                                  |      |
| Never smoked             | 141                          | 68.1 | 277                              | 94.5 |
| Past smoker              | 27                           | 13.1 | 11                               | 3.8  |
| Current smoker           | 39                           | 18.8 | 5                                | 1.7  |

“Regular exercise” means if a person was doing at least 20 minutes exercise or at least 30 minutes brisk walk four or more times per week (Khuwaja *et al.*, 2004).

Hypertension was defined as the presence of systolic blood pressure ≥140 mmHg and diastolic blood pressure of ≥90 mmHg or a known hypertensive on treatment (Shera *et al.*, 2004). BMI = Weight in kilograms/(Height in meters)<sup>2</sup>. Those who had BMI ≥25 were labeled “high BMI” (Khuwaja *et al.*, 2004) WHR was calculated by = Waist girth in c.m/Hip girth in c.m. The normal figure for males was >0.95 and for females <0.85 (Alberti and Zimmet, 1998). Responders who were currently smoking and smoked at least 100 cigarettes in their life time were defined as current smokers (Kujwaja *et al.*, 2004).

the same reason only 31 out of 500 diabetic patients were of 70 years or more. Similarly, in a survey of relatively prosperous and a poor area in Karachi, Pakistan a maximal prevalence of 25% was seen in the affluent community aged 55-64 years (Hameed *et*

*al.*, 1995). In another study conducted in rural town of Shikarpur in Sindh Province, Pakistan, it was reported that prevalence of diabetes mellitus rose with age (Shera *et al.*, 1995).

#### *Literacy rate related incidence*

In this study, it was found that about one third (35.6%) of diabetic patients with complications were illiterate. This finding is supported by a multi-center cross-sectional study conducted in Karachi (Khuwaja *et al.*, 2004) that had found 31% prevalence of diabetic complications in illiterate patients. One possible reason for this overall high prevalence in illiterate diabetic patients is that they did not know the importance of proper management of diabetes mellitus, optimal glycemic control and imbalance nutrition.

#### *Socioeconomic status*

The role of socio-economic status in promoting diabetes management and improved glycemic control is a little-explored area. Lower income is one of the problems for proper glycemic control in black children with diabetes (Delamater *et al.*, 1999). Present study also supports the above findings because lower income was one of the problems for maintaining good glycemic control in most of the patients with poor glycemic control.

#### *Duration of disease*

The duration of diabetes is considered an important factor in the development of complications (Knuiman *et al.*, 1986). In our study, 58.8% patients had diabetes and associated complications for more than five years. The findings of present study are supported by a population based cross-sectional study in Pakistan, in which highest prevalence of complications was found in persons having diabetes for more than 13 years (Hashim *et al.*, 2004). Present study indicates that 18.8% of individuals who are current smokers have retinal complications as compared to 13.1% of individuals who were past smokers.

#### *Hypertension*

Hypertension is a well recognized factor associated with the development of complications especially cardiovascular and cerebrovascular

diseases, and this risk increases further when it is associated with diabetes (Spanheimer, 2001). The present investigation documented that 39.2% of patients with type-2 diabetes have history of hypertension. The present findings are also supported by a cross-sectional survey in Sindh and Baluchistan provinces (Shera *et al.*, 1995, 1999a). Results from these landmark intervention studies demonstrate that the complications of diabetes can be prevented or delayed by controlling hypertension (UKPDS, 1998, 2000).

#### *Physical and biochemical factors*

In present study, it is found that lack of physical activity, greater body mass index and greater waist to hip ratio are more common in diabetic patients with complications. Present findings are supported by a number of studies which have shown that lack of physical activity, greater body mass index and greater waist to hip ratio play a significant role in the development of complications in patients with diabetes type-2 (Wing *et al.*, 1998; Williamson *et al.*, 2000).

HbA<sub>1C</sub> levels are a valuable adjustment to glucose determinations in individuals with diabetes mellitus, providing much more reliable determinations of glucose. Numerous studies have shown that diabetes related complications may be reduced by the long-term monitoring and tight control of blood glucose level (DCCT, 1993; Tietz, 1999). In the present study 159 diabetic patients having complications show poor glycemic control (HbA<sub>1C</sub> >8%). These findings are supported by Goldstein *et al.* (1995) and guidelines given in the clinical monitoring of diabetic patients (ADA, 1998). Suomen (1995) has also reported that HbA<sub>1C</sub> >8% is associated with poor glycemic control and development of diabetic complications. In present investigation, it was found that diabetic patients with complications showed high level of cholesterol, triglycerides, LDL-cholesterol and lower levels of HDL-cholesterol. This lipid profile is an important index to assess the metabolic control in diabetic patients (Davidson, 2002). A number of epidemiological studies support the present finding that low HDL-cholesterol concentrations, high concentrations of cholesterol, triglycerides, and LDL cholesterol are associated with high risk of

developing diabetic complications and poor metabolic control. Similar findings were also reported by Garcia and McNamara (1974) and Malmberg and Ryden (1988).

Retinopathy is more common and its prevalence is strongly related to the duration of diabetes, lack of exercise, hypertension and obesity/overweight (more BMI than 25 and more waist to hip ratio). In present study retinopathy was more commonly seen (41.4%) supported by the findings of Herman *et al.* (1998), who have reported retinopathy in 42% of their study subjects, contrary to findings (Ramachandran *et al.*, 1999), 23.7% retinopathy it has been reported in Indian (Morgan *et al.*, 2000), in 30.6% in Spain (Zafar *et al.*, 2000) and 44.6% in Mexico (Paisey *et al.*, 1984).

Present findings show that retinopathy is strongly associated ( $P < 0.01$ ) with age of the patient, duration of diabetes, having history of hypertension, more body mass index and more waist to hip ratio. These findings are also supported by other workers (Kinoshita *et al.*, 1990, Esperance *et al.*, 1990; ADA, 1998a).

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