### **Research Article**

### Assessment of glycemic control and correlation of HbA1c level with lipid profile and duration of diabetes in type 2 diabetes mellitus patients

Birendra Kumar Jha<sup>1\*</sup>, Raman Mishra<sup>2</sup>, Jitendra Kumar Singh<sup>3</sup>, Satyam Prakash<sup>4</sup>, Kamlesh Kumar Yadav<sup>5</sup>

#### Author's Affiliations

<sup>1,4</sup>Department of Biochemistry, Janaki Medical College, Tribhuwan University, Janakpurdham, Madhesh Province, Nepal

<sup>2</sup>Department of Internal Medicine, Janaki Health Care & amp; Teaching Hospital, Janakpurdham, Dhanusha, Madhesh Province, Nepal

<sup>2</sup>Department of Internal Medicine, Janaki Medical College, Teaching Hospital, Tribhuwan University, Janakpurdham, Madhesh Province, Nepal <sup>3</sup>Department of Community Medicine, Janaki

Medical College, Tribhuwan University, Janakpurdham, Madhesh Province, Nepal <sup>5</sup>Department of Clinical pathology, Janaki Health Care and Teaching Hospital, Janakpurdham, Dhanusha, Madhesh Province, Nepal

### Correspondence to:

Dr. Birendra Kumar Jha Department of Biochemistry, Janaki Medical College, Tribhuwan University, Janakpurdham, Madhesh Province, Nepal Email: <u>akshatjmc@yahoo.com</u>

#### ABSTRACT

**Background & Objectives:** The prevalence of Type 2 diabetes mellitus, or Type 2 DM, is rising globally. A periodic estimation of HbA1c has been

recommended to track the state of glycemic control of Type 2 DM patients, in order to control the diabetes associated disorders. The main consequence and cause of death in Type 2 diabetes has been identified as cardiovascular disease (CVD).Optimization of glycemic status along with decreasing CVD risk factors are critical to preventing morbidity and mortality in patients with Type 2 DM. Since there is a lack of data on these CVD risk factors, in Madhesh Province, Nepal, we aimed to evaluate the glycemic status, estimate the lipid profile, and ascertain their correlation, including duration of Type 2 DM among patients in tertiary care hospitals in Madhesh Province, Nepal.

**Materials and Methods**: The tertiary care hospital based cross-sectional study was conducted on 139 Type 2 DM patients, in Madhesh Province, Nepal. In order to gather information of the participants, a systematic questionnaire was introduced. A physical and clinical examination was performed, and an appropriate volume of blood was drawn with prior written consent. The lipid profile parameters, HbA1C, Fasting and post prandial serum glucose were estimated.

**Results**: In 84 (60.3%) and 106 (76.3%) cases, there was an increased fasting and postprandial blood sugar. The mean±SD HbA1c level was 8.40±0.8%. More than half (52.5%) of the patients Jha, BK, et al.

had poor glycemic control, while 22.3% had inadequate glycemic control, in reference to HbA1c >8% and 7 to 8%, respectively. A substantial increased level of LDL-C, triglycerides and total cholesterol was observed in 122 patients and 85.6% had reduced HDL-C. A significant strong positive association was found between HbA1c level and total cholesterol and triglycerides, in contrast, no discernible association with duration of disease observed.

**Conclusion:** Glycemic control was inadequate. Triglycerides and total cholesterol were significantly positively correlated with HbA1c, but not with HDL-C, LDL-C, or the course of diabetes.

**Keywords:** Diabetes Mellitus, Glycemic Control, HbA1C, Lipid profile, Madhesh Province

### **INTRODUCTION**

The hallmark of type 2 diabetes mellitus (Type 2 DM) is characterized by the persistent hyperglycemia, which can be caused by a variety of factors, including abnormalities in insulin secretion, action, or combo of each [1]. Type 2 DM is growing progressively and thought to be the consequences of food customs, sedentary lifestyles, physical inactivity and obesity [2]. According to the WHO estimates, 422 million adults worldwide had diabetes in 2023. Type 2 DM is the most prevalent type making for around 90% of all cases of the diabetes [3]. In addition to its abnormal metabolism of carbohydrates, Type 2 DM has also been found connected to defective metabolism of both lipids and proteins, resulted by insulin resistance or insufficiency [4]. According to the International Diabetes Federation (IDF), there were approximately 90 million adults in south Asian countries (aged 20-79) who had diabetes. It is projected that by the year 2045, the figure will have surged by 69% to 152 million. In South East Asia, there are 74.2 million Type 2 DM patients, of these ,74.2 million are in India, 13.1 million are in Bangladesh, 1.4 million are in Sri Lanka, and 1.1 million are in Nepal [5].

HbA1c, which depicts the average glycemic over a few months, is a highly reliable indicator of diabetes problems. Consequently, it has been advised to measure HbA1c every three months to track the state of glycemic control diabetes patients in [1]. Cardiovascular disease (CVD), which is the disease of heart and blood vessels, is the foremost cause of death in people living with diabetes. It has been shown that the risk of stroke doubles in individuals with Type 2 DM. Numerous studies have found a substantial positive correlation between glycemic control and CVD. According to reports, around 60% of all deaths in individuals with diabetes are attributed by CVD [6]. Atypical lipid profile parameters are thought to be the most significant risk factors contributing to the development of CVD, aside from metabolic syndrome [7]. LDL cholesterol levels rise among those with Type 2 DM partly because of a reduction in insulin levels accompanied by a decrease in the amount of the LDL receptor [8]. Dyslipidemia is defined as elevated triglycerides, total cholesterol, and low density lipoprotein cholesterol (LDL-C) along with decreased HDL-C. This condition has been noticed to be strongly associated with the progression of atherosclerosis [9]. Cardiovascular disease morbidity and mortality have been affected by central obesity, hypertension, dyslipidemia, and longterm poor glycemic control [10]. Research has shown a significant association between the duration of diabetes and lipid profile parameters, as well as HbA1c levels [2]. Previous published scientific literatures depicts that the Type 2 DM prevalence is trending upward in Nepal. Just 6.3% of Jha, BK, et al.

people were reported to have diabetes in 2000; by 2007, that number had risen to 9.5%. The pooled prevalence of Type 2 diabetes was 8.4%, as per a 2014 study [11-13]. Despite the fact that Type 2 DM is increasing in Nepal, the evidence regarding glycemic control and its correlation with the duration of diabetes and serum lipid profile is limited. Therefore, we aimed to evaluate the level of glycemic control and the correlation between HbA1c and lipid profile as well as the duration of diabetes in patients with Type 2 DM, visiting tertiary healthcare center in the Madhesh Province of Nepal.

### MATERIALS AND METHODS

A hospital-based cross-sectional study was carried out among known Type 2 DM patients, who were receiving outpatient medical care at Janaki Medical College Teaching Hospital (JMCTH) and Janaki Health Care and Teaching Hospital (JHCTH), Janakpurdham, Madhesh Province, Nepal. The study was carried out between 1st October 2023, and 15th February 2024. Sample size was determined to be 120 based on an 8.5% anticipated prevalence of Type 2 DM at 95% confidence interval, using Cochran's sample size calculation formula; however, we could recruit up to 139 individuals who meet the eligibility requirements.

Participant's eligibility criteria: An individual with known Type 2 DM (as per WHO criteria: fasting plasma glucose level≥126mg/dl, post prandial plasma glucose level ≥200mg/dl, HbA1c ≥6.6 or random plasma glucose level  $\geq 200 \text{mg/dl}$ ) either diagnosed for first time or who had followed up was included. Patients on steroids, contraceptive pills, statin and nephrotic diagnosed with syndrome. hypothyroidism,

familial

hyperlipoproteinemia and chronic kidney disease were excluded, on the basis of information obtained from the patients.

**Procedure:** The study was carried out from 25th of September 2023 to 15th of March 2024. Participants were enrolled using consecutive sampling, until the desired sample size was reached. To gather data on participants, a standardized questionnaire with anthropometric and sociodemographic parameters was employed. . Physical and circumference. clinical (waist Hip circumference, height, weight and blood pressure) examination were conducted by the trend medical health professionals (interns, medical officer) for all participants. A 5ml of fasting venous blood samples were collected, maintaining aseptic condition. Sterile gloves were worn while using a sterile disposable 5 ml syringe to draw blood, and methylated spirit was applied as an antiseptic around the puncture site.

Fasting blood sugar, total cholesterol, LDL-C, HDL-C, triglycerides were estimated using fully automated Accent 200 analyzer. For post prandial blood sugar, blood sample was collected at 2 hours of 75 gram glucose drink. HbA1C was estimated by fully automated D-10 (Bio-RAD), at the laboratory of JHCTH.

**Statistical analysis:** The data were entered in Excel sheet and analyzed by SPSS version 20. Frequencies and percentage were used for categorical variables. Central tendency and dispersion were used for numerical variables. The p value of less than 0.05 was considered as statistically significant.

**Ethical consideration:** The Ethical Review Board (ERB) of the Nepal Health Research Council (Ref. 463/2023) provided the ethical Jha, BK, et al.

approval for this study. Written as well as verbal consent were obtained from all the enrolled patients. All the enrolled participants were assured about the confidentiality.

### RESULTS

Out of total 139 participants, a nearly equal number of males and females participated

(males: 69, or 49.6%, females: 70, or 50.4%)).

The majority of study participants (42, or 30.2%) belonged to the 51–60 year age group, with a mean±2SD age of 53±12.2 years. A majority of study participants (117, or 84.2%) were Hindu, and nearly all of them were married (94.2%). In terms of education, a majority of study subjects (82, or 59%) had just completed their primary schooling. The

Table 1: So	ocio-demograp	hic charact	eristics of	studv sul	biects (r	n=139)
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Variables	Category	Frequency (%)	
Age in years		53.13±12.248	
(mean±2SD)]			
	≤30	7(5.0)	
	31-40	19(13.7)	
Age group (vears)	41-50	32(23.0)	
Age group (years)	51-60	42(30.2)	
	61-70	31(22.3)	
	71-80	8(5.8)	
Conder	Male	69(49.6)	
Genuer	Female	70(50.4)	
	Unmarried	1(0.7)	
Marital status	Married	131(94.2)	
	Widowed	7(5.0)	
Religion	Hindu	117(84.2)	
Kengion	Muslim	22(15.8)	
	Primary	82(59.0)	
	Vocational training	2(1.4)	
Educational level	Secondary	24(17.3)	
Buutational level	Graduation/Masters	14(10.1)	
	Middle school	8(5.8)	
	Certificate level/diploma	9(6.5)	
	Professional	26(18.7)	
	Semi-professional	9(6.5)	
Occupation	Farmer/shopkeeper	79(56.8)	
o o o a participa	/clerk	(00.05)	
	Skilled manpower	13(9.4)	
	Unemployment	12(8.6)	
	≥29166	66(47.5)	
	14584-29166	14(10.1)	
Monthly income	10937-14583	25(18.0)	
monthly income	7291-10936	15(10.8)	
	4374-7290	14(10.1)	
	1473-4373	5(3.6)	

next most common level of education was secondary (24, or 17.3%), followed by master's or graduation-level education (10.1%).

Farmers, shopkeepers, and clerks accounted for more than half of the study subjects (79, or 56.8%), followed by professionals (26, or 18.7%). Approximately 47.5 percent of the study subjects (66, or 47.5%) had a monthly household income of  $\geq$ 29166 (Table 1).

Table 2 indicates the mean±SD duration of

33, or 23.7%, were overweight, and 32, or 23%, were obese. Majority of the study subjects (98, or 70.5%) were non-smoker and 102, or 73.4% did not consume alcohol. However, 23 or 16.5% was current smoker and 15, or 10.8% was current alcohol consumer. Regarding comorbidities, 49, or 35.3% of study participants had hypertension and 7, or 5% have been diagnosed with dyslipidemia, table 2.

An elevated level of fasting and post prandial blood sugar was observed in 84, or 60% and

Variables	Category	Frequency (%)	
DM duration (mean±SD)		5.4±4.6	
Anti-diabetic medication			
Oral (hypoglycemic tablet)		133(95.6)	
Insulin		6(4.3)	
	1-5	89(65.44)	
Duration of DM (years)	6-10	36(25.89)	
	≥11	14(10.0)	
	Under weight (<18.5kg)	12(8.6)	
BMI $(ka/m^2)$	Healthy weight (18.5-24.9kg)	62(44.6)	
	Over weight (25.0-29.9kg)	33(23.7)	
	Obese (≥30kg)	32(23.0)	
Smoking status	Non smoker	98(70.5)	
Shioking status	Past smoker (>1year)	18(12.9)	
	Current smoker	23(16.5)	
Alcohol consumption status	Non consumer	102(73.4)	
Alconol consumption status	Past consumer	19(13.7)	
	Current consumer	15(10.8)	
Comorhidities	Dyslipidemia	7(5)	
Comoi Diulties	Hypertension	49(35.3)	

(DM diabetes mellitus. BMI body mass index)

Type 2 DM was 5.4±4.6 years .The majority of the study subjects (89, or 65.44%) belong to the group with a Type 2 DM duration of 1-5 years. Among 139 subjects, 133 or 95.6%, were on oral hypoglycemic drugs, while the remaining people were taking insulin. Of them, 62, or 44.6%, were at a healthy weight, 106, or 76.3%, respectively, among study subjects. The mean±2SD HbA1c level was 8.40±0.8%. More than half (52.5%) of the study subjects had poor (>8%), 22.3% had inadequate and the remaining had good glycemic control as shown in Figure 1.





Figure 1: Glycemic control index among study subjects

Parameters	Category	Frequency
HbA1c (mean±SD)		8.40±0.8
	<110	55(39.6)
Fasting blood sugar (mg/dl)	≥110	84(60.4)
Post prandial blood sugar	<140	33(23.7)
(mg/dl)	≥1140	106(76.3)
Total cholostorol (mg/dl)	<200	122(87.8)
Total cholester of (hig/ df)	≥200	17(12.2)
Triglycoridos (mg/dl)	<150	96(69.1)
(ing/ui)	≥150	43(30.9)
HDL C (mg/dl)	M≥40, F≥150	20(14.3)
IIDE-C (IIIg/ul)	M<40, F<150	119(85.6)
	<130	24(17.3)
LDL-C (mg/dl)	≥130	115(82.7)

 Table 3: Biochemical parameters of study subjects (n=139)

(HDL-C high density lipoprotein cholesterol, LDL-C low density lipoprotein cholesterol)

Table 2 and 3 depicts that the prevalence of current alcohol users (15, or 10.8%) and smokers (23, or 16.5%) was found to be almost equal. A substantial increase in LDL-C (115 or 82.7%), triglycerides (96 or 69.1%),

and total cholesterol (122 or 87.8%) was observed in with Type 2 diabetes. In contrast, 119 participants, or 85.6%, had reduced HDL-C levels. In addition, 49, or 35% of total had hypertension.

Table 4 signifies significant strong positive association was found between the HbA1c level and total cholesterol (r=0.215, p=0.05) and triglycerides (r=0.233, p=0.01). Nonetheless, there was no discernible association with the length of diabetes observed. rate. According to this study, the age group of 51–60 years was found as the most affected by Type 2 DM (42 30.2%), followed by the 61–70 years age group (31 22.3%). Previous national and international studies have shown that an important risk factor influencing the elevated risk of pre-diabetes

Biochemical	Statistical test	HbA1c	Duration of DM
parameters			
Total cholesterol	Spearman correlation	0.215*	-0.110
	Sig.(2-tailed)	0.011	0.197
Triglycerides	Spearman correlation	0.233**	-0.154
	Sig.(2-tailed)	0.006	0.071
HDL-C	Spearman correlation	0.134	-0.19
	Sig.(2-tailed)	0.117	0.826
LDL-C	Spearman correlation	0.134	-0.107
	Sig.(2-tailed)	0.116	0.209

Table 4: Correlation of lipid profile with HbA1c and duration of DM

\*\*Correlation is significant at the 0.01 level (2-tailed)

\*Correlation is significant at the 0.05 level (2-tailed)

### DISCUSSION

Madhesh is the most populated province in Nepal according to the findings of the 2021 population census, containing 20.98% (61, 26,288) of the country's total population, with over 37,95,848 of those individuals being within the most susceptible age group of 15 to 59 years for Type 2 DM [14]. The main risk factors for developing diabetes are advanced age, ethnicity, a family history of the disease, smoking, obesity, and physical inactivity. These risk factors also raise the possibility of developing kidney disease, neuropathy, blindness, and lower-extremity amputation, which significantly raises the morbidity and mortality rates of diabetics [15,16].

Population aging has emerged as an obvious concern impacting public health as society continues to advance at an extremely rapid

and diabetes is advanced age [15-17] which is in support of our findings. Results were found to be consistent with a similar hospital-based study conducted in Nigeria, which showed that 28.7% of Type 2 DM cases were of the same age range [18]. A study form Netherlands has reported that almost 50% of all type 2 DM patients were of age >70 years [19]. Additionally, the most impacted age group of 55-64 years was reported for Ehiopians [20]. A previous similar study conducted in the Kathmandu, Nepal has reported that 95% of type 2 DM patients was of age group 41 to 80 years [21]. Furthermore, a survey found that the prevalence of pre-diabetes and diabetes among people aged 40-49 was 40.3% and 11.1%, respectively, while the comparable figures for people aged 60–69 surged to 23.9% and 47.6%, respectively [22]. The elderly are consequently more prone than the young and middle-aged to have diabetes and prediabetes, as well as to experience issues with their cardiovascular, retinal, and renal systems [23,24]. Moreover, there may be a correlation between a surge in lifestyle modification and the decreasing trend in the onset age of type 2 diabetes in developing nations like Nepal. Due to the fact that age affects both clinical and socioeconomic characteristics, age may have a significant impact on the contribution of risk factors for diabetes and pre-diabetes. Thus, it is important for healthcare institutions to create accurate diabetes prevention and control programs in Madhesh Province, it is imperative that they should provide clarity on the relationship between age and factors that influence the abnormal state of glucose metabolism.

The abnormal lipid profile that coexists with type 2 diabetes contributes to the elevated risk of cardiovascular diseases in T2DM patients. Numerous studies have shown a correlation between HbA1c and one or more lipid profile characteristics in T2DM patients, and some have proposed HbA1c as a potential biomarker for identifying T2DM patients at risk of CVD and for identifying the abnormal lipid profile of T2DM patients [25,26]. Our findings depicts statistically significant strong positive correlation was found in Type 2 DM patient's lipid profile parameters and the level of HbA1c. The importance of routinely assessing HbA1c as a diagnostic tool for monitoring glycemic control and diabetes treatment goals is widely acknowledged. This suggests that HbA1c helps to determine the risk of micro- and macrovascular issues indirectly and is directly linked to dyslipidemia in T2DM diabetic patients [27,28]. Dyslipidemia in T2DM patients is thought to be caused by insulin resistance. Increased TG levels in T2DM patients are reported to be associated with insufficient insulin secretion or function through several of mechanisms [25,29].

In our findings, one out of 2 (73, or 52.5%) of the Type 2 DM patients had reported having poor glycemic control, which was found higher than the previous hospital based study documented from the western Nepal [21]. The possible explanation for this could be that diabetes management is challenging owing to the chronic nature of the disease and its associated complications. Other potential causes of poor glycemic control may include different contributing factors such as inadequate or poor adherence to prescribed therapies, unhealthy diet, physical inactivity, rising levels of obesity, growing urbanization, changing lifestyle habits and modified sedentary lifestyle. Furthermore, the majority of healthcare facilities are located in urban regions, while rural healthcare facilities lacks in staff, infrastructure, and resources.

In the line with our study, similar incidence (70%) of poor glycemic control had been reported in a study carried out in the capital city of Nepal [30]. The study findings showed that 74.8%, almost three quarters of the type 2 DM patients had inadequate and poor glycemic control. Nearly the same, 73.8% of inadequate and poor glycemic control has been reported in a similar study documented from Ethiopia [20]. Furthermore, a study from India has also reported the similar incidence of poor glycemic status (urban population: 68.9% and rural population: 69.2%) conducted in self-reported Type 2 DM patients [31]. Our results, which are consistent with earlier studies carried out in Nepal, indicates that controlling and managing diabetes is a significant challenge in Nepal, including low disease awareness

among the general public, a variety of sociocultural factors, educational approaches, and a lack of programs to identify, treat, and prevent diabetes and its complications [13]. Consequently, it is vital that health care providers and policymakers should collaborate together to evaluate the growing diabetes burden in Madhesh province and develop effective preventative and control strategies.

Hyperglycemia and the duration of the disease are major risk factors for the progress micro-angiopathic-associated of various complications [32]. In uncontrolled diabetes, dysglycemia increases reactive oxygen species, and changes in signaling pathways lead to a variety of vascular dysfunctions [33]. additionally induce several It can complications in all organs, including the cardiorespiratory system [34]. A notable beneficial correlation was found in this study between the lipid profile parameter and the HbA1c level, an indicator of glycemic control. Similar finding was reported by a study done by Moss et al. [35]. Furthermore, a study done from eastern India has also reported the similar association between dyslipidemia and HbA1c level in diabetic patients [20].

However, the correlation of HbA1c and duration of diabetes was found not to be significant, which is in contrast of the previous similar study conducted in Bangladesh [36]. The positive correlation between HbA1c and duration of diabetes has been marked in a study suggesting that an incidence of complications can significantly increase in number in conjunction with elevated HbA1c levels. Fewer DM complications were found at HbA1c levels of 7% HbA1c levels [37]. It is possible that improving glycaemic control is typically

thought to reduce type 2 diabetes symptoms and may be plausible cause for no relation between duration of diabetes and HbA1c as depicted in our findings. The other probable reason may be due to the limited number of subjects in this study and bias from study subjects responding during the data collection.

Among 2 diabetes type patients, dyslipidemias is one of the modifiable risk factors for coronary heart disease. Diabetic or atherogenic dyslipidemia is characterized by a high triglyceride and low- and high-density lipoprotein profile. It is a reliable indicator of silent myocardial ischemia or coronary artery disease [38,39]. Our results showed a significant positive correlation between HbA1c and triglycerides (r=0.233, p=0.01) and HbA1c and total cholesterol (r=0.215, p=0.05), however it did not show any significant correlation with LDL-C and HDL-C. The findings of current study agree with some previous studies, reported significant correlation between HbA1C and one or more parameters of the lipid profile in Type 2 DM patients [40,41].

The similar positive correlation of HbA1c with serum triglycerides and total cholesterol but no significant correlation shown with LDL-C, documented by a study carried out in Bangladesh [36]. However, the correlation of HbA1c and duration of diabetes did not found to be significant, which is in contrast of the previous similar study conducted in Nepal [21]. The positive correlation of HbA1c with high triglycerides signifies HbA1c as a direct marker of hypertriglyceridemia and an indirect marker of risk assessment of coronary artery disease [28]. However, the study was limited in its ability to reflect the entire population of Madhesh Province,

Nepal, due to its hospital based design and implementation in only two tertiary care institutions located in the province's capital city.

### CONCLUSION

The significant number of Type 2 DM patients had inadequate and poor glycemic control while being on medication. HbA1c showed a significant positive association with triglycerides and total cholesterol, but not with HDL-C, LDL-C, or the course of diabetes. A notable positive correlation between HbA1c and lipid profile parameters suggests that, in addition to serving as a glycemic status indicator, HbA1c can also be employed as a dyslipidemia prediction in the treatment plan, helping to avert future CVD issues.

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