

Management Practice and Contributing Risk Factors for Chronic Complications Among Type 2 Diabetes Mellitus Adult Patients in Follow-Up at a Tertiary Care Teaching Hospital

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Background: Diabetes Mellitus is a complex, chronic disease that requires a coordinated management practice beyond blood glucose control. The disease causes chronic complications that affect the quality of the life of patients, place major pressure on the health-care system and cause a rise in diabetes-related mortality.

Objective: To determine the prevalence of chronic diabetes mellitus complications, related risk factors, and management practice among adult type 2 diabetes mellitus outpatients at Tikur Anbessa Specialized Hospital (TASH).

Methods: A cross-sectional study design was carried out from July to September 2018. The pre-tested data abstraction format was used to gather demographic and clinical information. We also used a balance of weight measurement, upright placed meter for height measurement, waist circumference measurement meter and BP equipment. Statistical analysis was accomplished using Statistical Package for the Social Sciences (SPSS) 25[®] software. The significance level for statistics was set at $p < 0.05$.

Results: In this study, 320 patients were involved. Of these, about 57% were female and had a mean age of 58 ± 11.2 years. About 85% of the study participants had comorbidity and 42.5% had complications. Hypertension and neuropathy were the most common comorbidity and complication, respectively. The usage of vascular preventive medication among study participants was 74.7% and 55.3% for statins and ASAs, respectively. Participants in the study who had disease duration of 5–10 years (AOR=3.50, 95% CI: 1.19–10.28) and fifteen and above (AOR= 3.59, 95% CI: 1.36–9.49) were at higher risk of diabetes complication as compared to less than five years.

Conclusion: The prevalence of chronic complications was high among adult T2DM outpatients. The duration of disease and the number of medications used were the factors associated with chronic complications. The use of vascular preventive medications was low among study participants.

Keywords: type 2 DM, diabetes complication, risk factors, vascular-preventive medications

Introduction

Type 2 diabetes mellitus (T2DM) is one of the most common metabolic disorders majorly affecting the adult population. It accounts for 90–95% of all diabetes cases.^{1–3} Long-term elevations in blood glucose levels contribute to many complications in various organs, including the kidneys, skin, nerves, heart, and blood vessels.⁴ This rise in blood glucose, particularly after diagnosis, is mainly due to

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a lack of compliance with the management plan that involves lifestyle modification and pharmaceutical interventions.⁵

One of the main impairment occurring from hyperglycemia is damage to the vasculature. It occurs either at small (microvascular complications) or large blood vessels (macrovascular complications). Diabetic retinopathy is the most common microvascular complication, followed by diabetic nephropathy and neuropathy. All macro-vascular complications arise from the development of atherosclerosis, which gradually causes the narrowing of arterial walls.^{6,7}

The diabetes complication is highly prevalent in developing countries such as Sub-Saharan Africans (SSAs), along with an increase in attributing factors.^{3,8} According to the study conducted in Sub-Saharan countries taking the sample from Nigeria, Ghana, and Kenya, hypertension (71%), erectile dysfunction (35%), hyperlipidemia (34%), cataract (32%), obesity (27%), retinopathy (15%), and impaired renal function (13%) were the most commonly reported complications/comorbidities in T2DM patients. The most common risk factors for the development of complications and comorbidity were duration of T2DM, higher body mass index (BMI), and older age.⁹

A local study conducted at Nedjo general and Nekemte referral hospitals showed that at least one complication of diabetes mellitus occurred in about 1/3rd of the study participants. The study also showed that the duration of diabetes, patient age, DM regimen, family history of diabetes, and the presence of comorbid medical conditions were positively associated with the occurrence of diabetes complications. In this study, hypertension (41.5%) was the most reported chronic complications in diabetes patients, followed by visual impairment (20.7%) and nephropathy (15.9%).¹⁰

A better understanding of the burden of diabetes complication among diabetes patients and factors associated with the disease requires important information. It is also necessary recognizing the management strategies practiced as treatment and prevention to prevent immediate and long-term complications, and enhance the patient's quality of life.⁵ Hence, the study was aimed to assess T2DM complications, their management practice, and risk factors attributed to the existence of complications.

Methods

Study Design and Population

Hospital-based cross-sectional study design was applied from July to September 2018 for a total of 3 months at

the diabetes clinic of Tikur Anbessa Specialized Hospital (TASH), Addis Ababa, Ethiopia, which. TASH is the largest referral hospital in the country that provides a wide-range health-care service for more than 500,000 patients per year. Diabetes clinic is one of the specialty clinics that provide an outpatient service for different endocrine disorders. Adult T2DM patients attend the clinic every Monday and Wednesday based on their appointment. Per annum, more than 6000 patients were attending the diabetes clinic with an average of about 250 patients per week.

All adult type 2 diabetes patients (Age >18years) attending diabetes follow-up clinics of TASH who were diagnosed as DM for at least three months and on at least one medication were included in the study. Severely sick patients who need immediate medical treatment and patients who refuse to join the study were excluded from the study.

Sample Size and Sampling Technique

A total of 423 sample size were calculated based on single proportion formula assuming a prevalence (p) of 50% and 10% contingency (for non-response rate) as there was no research performed on this topic in the setup or the country. During the study period of three months, patients who had an appointment made the sampling frame and the sampling fraction was calculated. The average weekly attender (~250 patients) was used to find a constant (k). During the study period of three months, the clinic provides service for 24 days, and based on this, an average of about 18 patients were sampled every constant (k=7) each day in a systematic random sampling technique. Then, participant's medical card numbers (ID) were taken every seventh interval for comprehensive chart review and patients' interview.

Data Collection and Management

Demographic (age, gender, marital status, education, residence, and occupation status) and clinical (comorbidity and complication, duration of the disease, type of medications, source of medication, lifestyle, family history, and physical activity) information was collected using the pre-tested data abstraction format. We had also used weight-measuring balance, height measuring upright positioned meter, waist circumference measuring meter, and blood pressure measuring BP apparatus as data collection tool. For this study, two nurses and two pharmacists were recruited for data collection. They engaged in one-day training to familiarize them with the

instrument for data collection as well as how to address possible problems encountered during data collection.

A pre-test was performed on five percent of the sample at the diabetes clinic of TASH before data collection started. Based on the results obtained from the pre-test, an amendment was made on the assessment tools and ways of assessment. The training was provided for the data collectors. The principal investigator (PI) was closely supervising the data collection on a daily basis. At the end of each data collection day, the PI checked the completeness of the filled questionnaire and recorded information to ensure its accuracy. After data collection was completed the PI carefully cleaned and entered the data into SPSS. The data analysis was performed with the consultation of a biostatistician.

Data Analysis

The collected data were evaluated, cleaned, entered, and analyzed using SPSS version 25. Descriptive statistics were computed as frequency, percent, mean and standard deviation (SD) for continuous and categorical variables. Logistic regression tests were performed to determine the relationship of independent variables with diabetes complications in type 2 diabetes patients and to examine possible confounders. Binary logistic regression was performed with p -value <0.20 and those variables having significant p -value were moved into multiple logistic regression to control the influence of the confounding variables. A 95% CI and the statistical significance level were determined at a p -value of <0.05 for all variables.

Ethical Considerations

The study was conducted after ethical approval was acquired from the College of Health Sciences of Addis Ababa University Institution Review Board (Ref. No.: 002/17/SPharma). Guidelines outlined in the Declaration of Helsinki were met. Written informed consent was also obtained from participants before collecting the required data. The study result did not intend to include participants' identifiers. The raw data were not made available to anyone, other than the research team.

Results

Socio-Demographic Characteristics

In the current study, 320 adult type 2 diabetes patients had included in the final analysis. Approximately 57% of them

were female, and the majority (52.8%) were in the age range of 50 and 65 years old with a mean age of 58 (SD, 11.2) years. Nearly 3/4th of the study participants were married and got their medication freely (Table 1).

Clinical Characteristics

In this study, about 85% of study participants had comorbidity, and 42.5% had complications. Hypertension and neuropathy were the most common comorbidity and complication, respectively (Table 2). Among study participants,

Table 1 Socio-Demographic Characteristics of Adult Type 2 Diabetes Patients on Follow-Up at Tikur Anbessa Specialized Hospital, Addis Ababa, Ethiopia (N=320)

Variables	Categories	N (%)
Age (years)	Mean $\pm$ SD	58.1 $\pm$ 11.2
	≤ 35	11(3.4)
	35–50	70(21.9)
	50–65	169(52.8)
	>65	70(21.9)
Sex	Male	139(43.4)
	Female	181(56.6)
Marital status	Married	250(78.1)
	Single	23(7.2)
	Divorced	18(5.6)
	Widowed	29(9.1)
Education	No formal education	47(14.7)
	Primary(1–8)	59(18.4)
	Secondary(9–12)	97(30.3)
	College/university	117(36.6)
Residence	Addis Ababa	278(86.9)
	Out of Addis Ababa	42(13.1)
Occupation	Employed	85(26.6)
	Unemployed	44(13.8)
	Private	41(12.8)
	Student	7(2.2)
	Others*	143(44.7)
Source of med	Buying	74(23.1)
	Free	246(76.9)
Smoking	Non smoker	312(97.5)
	X-smoker	3(0.9)
	Smoker	5(1.6)
Alcohol	Yes	41(12.8)
	No	279(87.2)
Physical activity	Yes	274(85.6)
	No	46(14.4)

Note: *Retired.

Abbreviation: SD, standard deviation of mean.

Table 2 Clinical Characteristics of Adult T2DM Patients on Follow-Up at Tikur Anbessa Specialized Hospital, Addis Ababa, Ethiopia (N=320)

Variables	Categories	N (%)
Body mass index	Normal	107(33.4)
	Under weight	2(0.6)
	Over weight	91(28.4)
	Obese	37(11.6)
Blood pressure	Uncontrolled	96(30.0)
	Controlled	106(33.1)
Hyperglycemia	Yes	221(69.1)
	No	99(30.1)
Hypoglycemia	Yes	39(12.2)
	No	281(87.8)
Lipid control	Controlled	300(93.8)
	Poorly controlled	20(6.3)
Number of comorbid	No	48(15.0)
	1–2	217(67.8)
	≥3	55(17.2)
Type of comorbidity	Hypertension	221(69.1)
	Dyslipidemia	90(28.1)
	Ischemic Heart Disease	50(15.6)
	Chronic Kidney Disease	23(7.2)
	HIV/AIDS	13(4.1)
	Asthma	9(2.8)
	Others*	91(28.4)
Number of complication	No	184(57.5)
	1–2	127(39.7)
	≥3	9(2.8)
Type of complications	Neuropathy	116(36.3)
	Nephropathy	26(8.1)
	Retinopathy	21(6.6)
	Diabetes foot ulcer	4(1.3)
	Others**	13(4.1)
Number of medication	Mean ± SD	4.97 ± 2.05
	<5	131(40.9)
	≥5	188(58.8)
Duration of disease (years)	Mean ± SD	13.83 ± 8.87
	<5	35(10.9)
	5–10	42(13.1)
	10–15	51(15.9)
	≥15	99(30.9)

Notes: *Thyroid disorders, osteoarthritis, psychotic disorder, infection, cancer, seizure, obesity. **Peripheral arterial disease, autonomic gastroparesis; skin infection, sexual dysfunction.

Metformin + Insulin was the most utilized antidiabetes medication, while ACEIs/ARBs were the most utilized medication for other than diabetes (Figure 1).

Factors Associated with Diabetes Complications

Sociodemographic and clinical characteristics of patients were subjected to logistic regression to determine a predictor of complications in T2DM patients. From there, the number of medications taken and the duration of the disease had significantly related to the occurrence of complications in type 2 diabetes patients (Table 3). Patients who take five and above medication were more than three times (AOR= 3.08, 95% CI: 1.62–5.85) risk of developing complications. Study participants who had a disease duration of 5–10 years (AOR=3.50, 95% CI: 1.19–10.28) and fifteen and above (AOR= 3.59, 95% CI: 1.358–9.493) were also more than three times at risk of diabetes complication as compared to those who have below five years.

Discussion

T2DM is one of the most common metabolic diseases that have an outcome of chronic complications upon progress, which impair the wellbeing of patients, challenge the health service providing system, and increase diabetic mortality.¹¹ In Ethiopia, the annual health expenditure to treat diabetes and prevent complications among adults (20–79 years) was estimated to range from \$ 80–200 million in 2015.¹² Thus, understanding the prevalence and identifying associated risk factors of type 2 DM complication was important for improved management and prevention of the disease.^{9,13} Therefore, the objective of this study was to determine the prevalence of chronic complications, identifying related factors, and its management practice among adult T2DM out-patients. Our data indicate that most participants in the study had comorbidity, which could raise the risk of complications. The duration of disease and the number of medications was the factor related to the occurrence of T2DM complications.

In this study, about 42% of study participants had at least one complication. This finding is consistent with other local research conducted in Wollo (48.6%)¹⁴ and Jimma (52.5%)¹⁵ Ethiopia, Chin (52%),¹³ and Taiwan (52.6%).¹⁶ This high prevalence of complications among T2DM patients could be associate with the advanced age of study participants (mean= 58 (SD, 11.2)) and the presence of many comorbidities (85%). Our findings were lower than the study conducted in the United Arab Emirates (UAE) (83.5%).¹⁷ Some of the disparity realized in the occurrence of complications across different countries could also be due to differences in medical care,

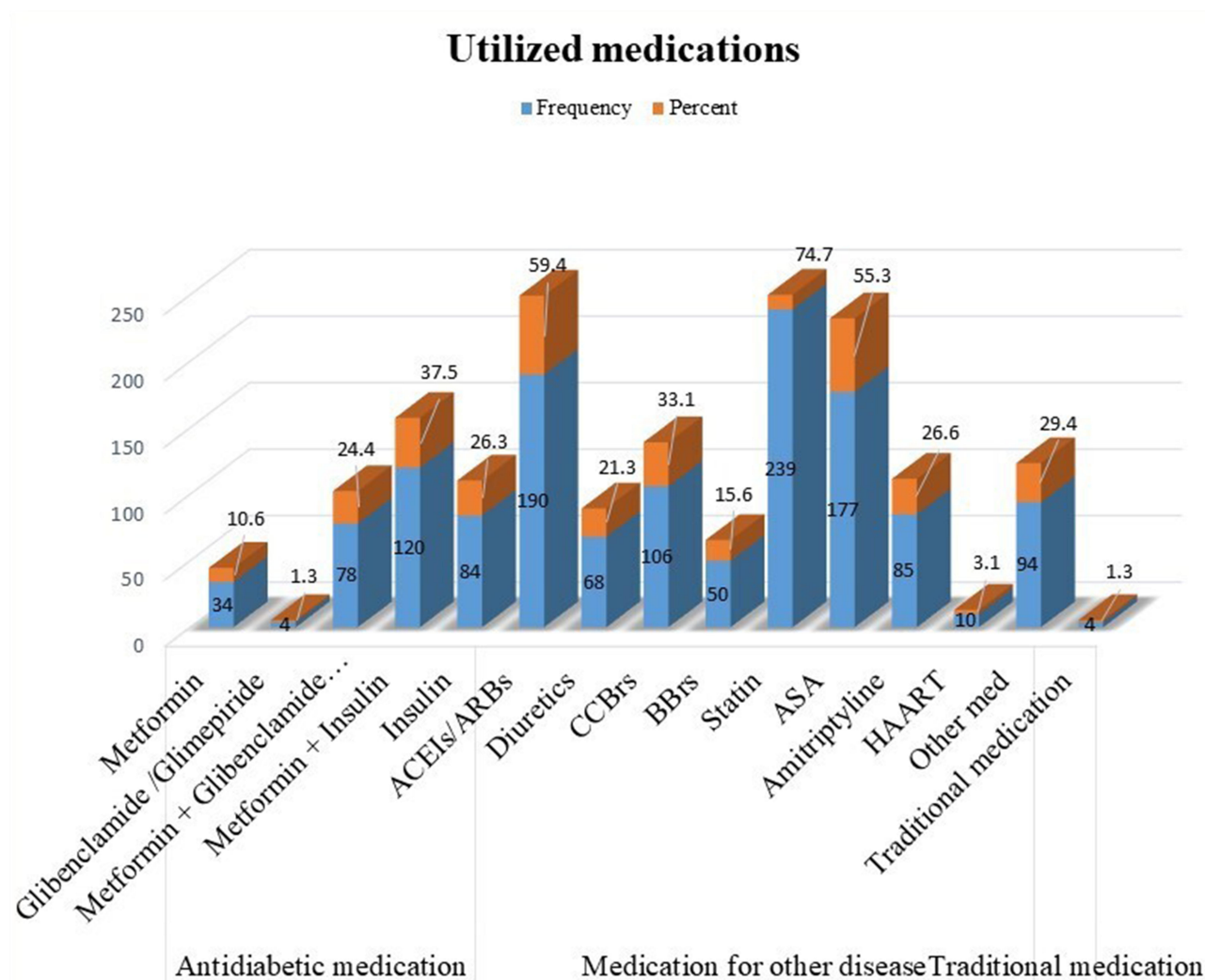


Figure 1 Prescribed medications among adult type 2 diabetic patients on follow-up at Tikur Anbessa Specialized Hospital, Addis Ababa, Ethiopia.

Notes: Others medication: metformin + sitagliptin, Proton pump inhibitors, carbamazepine, gabapentin, tramadol, propylthiouracil, Non-steroidal anti-inflammatory drugs.

Abbreviations: ACEIs/ARBs, angiotensin-converting enzyme inhibitors/angiotensin II receptor blockers; CCBrs, calcium channel blockers; BBrs, beta-blockers; HAART, highly active antiretroviral therapy.

health service resources, diagnostic guidelines applied, and definitions used in clinical practice.

Neuropathy was the most commonly reported complication that had a negative influence on the quality of life in patients with diabetes.¹¹ In the current study neuropathy was the most common observed complication followed by nephropathy while in some studies nephropathy was the most common followed by neuropathy.^{18–20} In agreement with our findings, the systematic review of research conducted in North Africa showed that chronic diabetes complications prevalence ranged from 6.7% to 46.3% for nephropathy and 21.9% to 60% for neuropathy.²¹ Similarly, in a multinational study conducted at 28 countries in different continents such as Asia, Africa, Europe,

and South America, neuropathy was the most common microvascular complication across the study regions (25% to 83%).²²

Diabetes comorbidities are also disorders that impact people with diabetes compared to those people without diabetes.²³ In the current study, about 85% of study participants had at least one comorbidity. Among these, hypertension (69%) was the most common prevalent comorbidity followed by dyslipidemia (28%). This finding is consistent with other studies that reported hypertension as a common comorbid disease.^{9,15,17,20} The high proportion of hypertension, in specific, has a prominent impact on the prevalence of microvascular and macro-vascular complications.

Table 3 Bivariate and Multivariate Analysis of Factors Associated with Diabetes Complications Among Adult T2DM Patients on Follow-Up at Tikur Anbessa Specialized Hospital, Addis Ababa, Ethiopia

Variables	Categories	Complication		Odds Ratio (95% CI)	
		Yes (N (%))	No (N (%))	COR	AOR
Age (years)	≤35	10(3.1)	1(0.3)	1.00	
	35–50	47(14.7)	23(7.2)	4.89(0.59–40.58)	0.94(0.08–11.51)
	50–65	97(30.3)	72(22.5)	7.42(0.93–59.30)	1.27(0.11–14.98)
	>65	30(9.4)	40(12.5)	13.33(1.62–109.91)	1.13(0.09–14.27)
Blood pressure	Controlled	53(26.2)	53(26.2)		
	Uncontrolled	55(27.2)	41(20.3)	0.75(0.43–1.30)	0.69(0.33–1.44)
Hyperglycemic	Yes	127(39.7)	94(29.4)		
	No	57(17.8)	42(13.1)	1.00(0.62–1.62)	0.81(0.36–1.81)
Lipid	Controlled	173(54.1)	127(39.7)		
	Poorly controlled	11(3.4)	9(2.8)	0.90(0.36–2.23)	1.20(0.26–5.53)
Diabetes regimen	Metformin	27(8.4)	7(2.2)		
	Glibenclamide/Glimepiride	3(0.9)	1(0.3)	1.29(0.12–14.33)	1.03(0.07–16.48)
	Metformin+ Glibenclamide/Glimepiride	52(16.3)	26(8.1)	1.93(0.74–5.01)	1.26(0.33–4.91)
	Metformin+ Insulin	62(19.4)	58(42.6)	3.61(1.46–8.92)	2.83(0.74–10.88)
	Insulin	40(12.5)	44(13.8)	4.24(1.67–10.81)	2.68(0.68–10.62)
Statins	Yes	124(38.8)	115(35.9)	1.00	
	No	60(18.8)	21(6.6)	0.63(0.18–2.19)	0.53(0.17–1.71)
Aspirin	Yes	94(29.4)	83(25.9)	1.00	
	No	90(28.1)	53(16.6)	1.75(0.58–5.24)	1.35(0.53–3.44)
ACEI/ARB	Yes	101(31.6)	89(27.8)	1.00	
	No	83(25.9)	47(14.7)	0.86(0.24–3.07)	1.07(0.74–3.08)
Number of medication	<5	92(28.8)	39(12.2)	1.00	
	≥5	91(28.5)	97(30.4)	2.52(1.57–4.03)	3.08(1.62–5.85)
Duration of disease (years)	<5	28(12.3)	7(3.1)	1.00	
	5–10	21(9.3)	21(9.3)	4.00(1.43–11.16)	3.50(1.19–10.28)
	10–15	31(13.7)	20(8.8)	2.58(0.95–7.02)	2.47(0.85–7.18)
	≥15	52(22.9)	47(20.7)	3.62(1.45–9.05)	3.59(1.36–9.49)
Number of comorbidity	No	31(9.7)	17(5.3)	1.00	
	1–2	131(40.9)	86(26.9)	1.20(0.62–2.30)	0.50(0.20–1.23)
	≥3	22(6.9)	33(10.3)	2.74(1.23–6.09)	1.20(0.41–3.54)

Note: Bolds: statistical significant (P-value <0.05).

Abbreviations: AOR, adjusted odds ratio; CI, confidence interval; COR, crude odds ratio; ACEIs/ARBs, angiotensin converting enzyme inhibitors/angiotensin II receptor blockers.

Poor glycemic control is a significant factor that contributes to the development of chronic complications related to diabetes.²⁴ Approximately 69% of the blood glucose levels of the study participants were in the hyperglycemia range based on the standard treatment guideline reference. These require appropriate management action since sufficient metabolic management either prevents or delays the occurrence of chronic complications.²⁵ Justification for the difficulty in attaining target blood

glucose levels could include the presence of polypharmacy that affect medication adherence, fears of medication adverse effect such as dyspepsia, hypoglycemia, and weight gain, lack of access to therapy and unable to follow dietary approaches to stop hypertension (DASH).

In the current study, about 59% of study participants took five and above medication for the management of their disease condition. Regarding medications, metformin + insulin was the most utilized medication, followed by

insulin for the management of diabetes. Metformin was the most utilized medication alone or in combination with other antidiabetic medications. It is the most recommended medication to reduce the development of chronic complications and comorbidity in diabetes patients.²³

In this study, the usage of vascular preventive modality was suboptimal, including those patients with vascular complications. Our findings were consistent with a multinational study conducted in twenty-eight countries across seven geographical states (South America; North Africa, South, and East Asia, Russia, Middle East/Gulf, and China).²²

In this study, the occurrence of chronic complications in T2DM patients was closely associated with the number of medications used and disease duration. For type 2 diabetes patients, polypharmacy is a common problem which is known to affect a patient's quality of life.²⁶ In the current study, the risk of developing diabetes complications was more than three times (AOR= 3.08, 95% CI: 1.62–5.85) in patients who took five or more medications. The current finding was consistent with the systematic review and meta-analysis that showed polypharmacy was strongly associated with the increase of all-cause mortality and macrovascular complications among type 2 diabetes patients.²⁷

The prevalence of the identified chronic diabetes complications was also found to be positively associated with the duration of disease, regardless of the patients' age. The study participants who had disease duration of 5–10 years (AOR=3.50, 95% CI: 1.19–10.28) and fifteen and above years (AOR= 3.59, 95% CI: 1.358–9.493) were more than three times at risk to develop diabetes complication compared to those who had less than five years disease duration. This finding is in agreement with the results of other studies done elsewhere.^{13,15,22,28}

Limitation

Though the study showed important findings of chronic complications of T2DM, its risk factor, and management practice in TASH, it has its own limitations. Firstly, the cross-sectional study design could not reveal the condition of the actual patient. Secondly, the study participants were patients who had a multi-disease burden having following up in tertiary referral healthcare, resulting in an over-consideration of the actual prevalence of the diabetes complication. Therefore, it may be somewhat improper to generalize the findings of this study to the entire population of T2DM patients.

Conclusion

The prevalence of complications among T2DM patients was found to be high in TASH. The number of medications and the duration of the disease was strongly associated with the development of chronic diabetes complications. Vascular preventive regimen in patients with or without complication is suboptimal in the current study. Appropriate actions should be taken to build up medical therapy and lifestyle management to overcome amendable risk factors for complications in order to reduce morbidity and mortality.

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Author Contributions

Both authors contributed significantly to the reported work, in the concept and design of the research, execution, data collection, analysis, and interpretation; engaged in the drafting, revision or critical review of the paper; gave final approval of the version to be published; agreed on the journal to which the paper was submitted and to take responsibility and accountability for the contents of the article.

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