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Glycaemic, blood pressure and low-density lipoprotein-cholesterol control among patients with diabetes mellitus in a specialised clinic in Botswana

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3 **Glycaemic, blood pressure and low-density lipoprotein-cholesterol control among patients with**
4 **diabetes mellitus in a specialised clinic in Botswana**
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Abstract

Objective: Control of glycaemic, hypertension and low-density lipoprotein-cholesterol (LDL-C) among type 2 diabetes mellitus (T2DM) patients is vital for the prevention of cardiovascular diseases (CVD). This study aimed to determine the prevalence of optimal glycaemic, hypertension, and LDL-C control among ambulant patients with type 2 diabetes mellitus in Botswana. Also, the research aimed at assessing factors associated with optimal glycaemic, hypertension and LDL-C therapeutic goals.

Design: A cross-sectional study.

Setting: A specialised public outpatient clinic in Gaborone, Botswana.

Participants: 500 T2DM patients on treatment for 3 or more months between August 2017 and February 2018.

Primary outcome measure: the prevalence of optimal glycaemic (HbA1c <7 %), hypertension (blood pressure <140/90mmHg) and LDL-C (<1.8mmol/L) control.

Results: The proportions of patients meeting optimal targets were 32.3% for glycaemic, 54.2% for hypertension, and 20.4% for LDL-C. Optimal glycaemic control was positively associated with age \geq 50 years (AOR 5.79; 95% CI 1.08 to 31.14) but was inversely associated with an increase in diabetes duration (AOR 0.91; 95%CI 0.85 – 0.98). Patients on angiotensin converting inhibitors (ACEIs) had lower odds of optimal hypertension control (AOR 0.35; 95% CI 0.14 – 0.85) than those without ACEIs. Being female (AOR 0.21; 95% CI 0.08 – 0.58) was inversely associated with an optimal LDL-C control.

Conclusion: T2DM in Gaborone, Botswana, presented with poor control of recommended glycaemic, hypertension and LDL-C control. These finds calls for urgent individual and health systems interventions to address key determinants of the recommended therapeutic targets among patients with diabetes in this setting.

Keywords: Glycemic, hypertension, LDL, Diabetes Mellitus, Botswana

Strengths and limitations of this study

- The study provides data on the three critical therapeutic targets in patients with diabetes in a specialised public clinic in Botswana.
- We explored factors associated with glycaemic, hypertension and LDL-C control.
- The study was limited to one public clinic and may not be generalised to other facilities in the country. However, the findings can be used as a basis for future studies and efforts to improve diabetes care in Botswana and different settings in Africa.
- The cross-sectional design limited the assessment of the temporal relationship between factors associated with poor control of glycaemic, LDL-C and hypertension.

Introduction

Diabetes mellitus and related cardiovascular complications are growing public health concerns worldwide^{1,2}. There are approximately 16 million people with diabetes in Africa, and this number is projected to increase to 41 million by 2045 due to rapid urbanisation, lifestyles changes and nutrition transition in the continent¹. This increase in prevalence and incidence of diabetes is attributable to type 2 diabetes, which is associated with multiple co-morbidities such as obesity and hypertension requiring chronic care and catastrophic health expenditure¹. Diabetes and associated comorbidities are known to increase patients risk to the development of cardiovascular diseases (CVD), which are responsible for about 70% of diabetes-related deaths³. The risk to the development of CVD is higher in people with suboptimal glycaemic, hypertension and LDL-C control^{3,4}. A reduction of HbA1c to control targets along with optimal hypertension control and the use of statins to lower LDL-C levels have been shown to improve long-term outcomes including reducing mortality among patients with diabetes⁵⁻⁸. Achieving these targets remains a challenge in most settings, especially in settings with limited access to standard diabetes care⁹⁻¹³. Only a minority of patients with diabetes in Africa achieve optimal therapeutic targets, leaving the majority of patients at high risk of diabetes-related complications¹². This is a public health concern because the current total health expenditure in most sub-Saharan Africa countries remains far below the 15% recommended in the Abuja declaration and the rising cost of managing diabetes complications will further make the health system goals unattainable^{1,14}. Thus, this study aimed to evaluate the current prevalence of glycaemic, hypertension, and LDL-C control among ambulant patients with type 2 diabetes mellitus in Botswana to provide future guidance. The study also assessed for factors associated with attainment of glycaemic, blood pressure and LDL-C therapeutic patients in these patients.

Methods

Study design and participants

We conducted a cross-sectional study of outpatients with established type 2 diabetes attending a specialised public diabetes clinic in Gaborone, Botswana between August 2017 and February 2018. The clinic has been operational since 2011 as a referral centre for health facilities in Gaborone and nearby towns. Patients were eligible to participate in the study if they were aged ≥ 18 years and had received care from the clinic for at least three months. Systematic random sampling was used to select ten patients from a list of about 80 patients who attended the clinic every day. We picked the first patient from the first eight people in the queue. Then every eighth individual was approached until the daily target of 10 patients was reached. Participants provided informed consent before enrolment. The study was granted the ethical approval by the Ministry of Health (HPDME: 13/18/1 VOL XI) and Princess Marina Hospital (PMH 5/79,317-1-2017) Institutional Review Boards.

Data collection and procedures

Patient information was collected using an interviewer-administered questionnaire and through reviews of medical charts and electronic records. The information included: demography (age, gender, occupation, marital status, and education), diabetes duration, history of hypertension, and medications for diabetes, hypertension and lipid disorders. We performed anthropometry (weight, height, waist and hip circumferences) and blood pressure measurements at enrolment. We conducted three blood pressure measurements after 10 minutes of rest, and the mean of the three measurements was recorded¹⁵. Moreover, we documented a blood pressure reading from each patient's prior visit. Patients' serum creatinine, LDL-C and HbA1c, and urine dipstick for proteinuria results over the past six months were abstracted from electronic medical records.

Definitions of the key outcomes and exposure variables

We calculated diabetes duration as the date of enrolment into the study minus the date of diabetes diagnosis. A patient was considered hypertensive by self-reported hypertension and the use of blood pressure-lowering medications or has sustained blood pressure $\geq 140/90$ mmHg during the previous visit and at enrolment¹⁶⁻¹⁸. Optimal glycaemic control was defined as HbA1c $< 7\%$ ^{16 18}. For patients who were on lipid-lowering medications, optimal LDL-C control was LDL cholesterol level < 1.8 mmol/L¹⁶. We calculated patients' eGFR by the Modification of Diet in Renal Disease (MDRD) formula, and an eGFR < 60.0 mL/min/1.73m² defined Chronic Kidney Disease (CKD)^{2 19 20}. Body mass index (BMI) was calculated as weight in kilograms divided by the square of height in meters. We considered underweight as BMI < 18.5 kg/m², normal weight as BMI of 18.5 –24.9 kg/m², overweight as BMI of 25.0 –29.9 kg/m², and obesity as BMI ≥ 30 kg/m²²¹. Waist-hip ratio (WHR) was calculated as waist circumference (WC) in centimetres divided by hip circumference (HC) in centimetres, and WHR ≥ 0.85 for women and ≥ 0.90 for men were classified as high²¹.

Statistical analysis

Analyses were performed using Stata, Version 14 (Stata Corp, College Station, TX). We used percentages to summarise categorical variables. Means and standard deviation (SD) or medians and interquartile range (IQR) were used to summarise continuous variables. Pearson's χ^2 or Fisher's exact tests were used to assessing statistical differences by gender for the categorical variables, while the Student's t-test or the Mann-Whitney U test were used for the continuous ones. Bivariate logistic regression was used to explore factors associated with each primary outcome - glycaemic, hypertension and LDL-C control. We further performed three multivariate logistic regression models for each of the three outcomes. The independent variables selected for multivariate models were those displaying a p-value < 0.2 at the univariate analysis level in addition to those considered clinically meaningful (age and gender). We described results as

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3 crude odds ratio (OR), adjusted odds ratio (AOR), and their corresponding 95% confidence intervals
4 (95% CI). A 2-tailed p-value < 0.05 was considered statistically significant.
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6 **Results**

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8 We included 500 patients with type 2 diabetes in the study, of which 330 (66%) were females. The mean
9 (SD) age was 58.9 (12.2) years and 78.2% were aged > 50 years. The median (IQR) diabetes duration
10 was 6 (2 – 13) years. There were a high percentage of patients with hypertension (80.8%), overweight
11 (31.1%) and obesity (50.2%). Table 1 summarises the patients' characteristics by gender. Female patients
12 tended to be older (60.8 vs 55.4 years, p<0.001, obese (56.7% vs 37.1%, p<0.001), have higher WHR
13 (95.2% vs 55.9%), be hypertensive (86.1% vs 70.6%, p<0.001), and have a higher mean total cholesterol
14 (4.5mmol/L vs 4.3mmol/L, p=0.030 and LDL-C (2.9mmol/L vs 2.6mmol/L, p=0.006) than male patients.
15 Urine dipstick was positive for protein in 10.2% of patients, mostly males (15.9% vs 7.3%, P = 0.003).
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22 **[Table 1: Clinical and socio-demographic characteristics of patients with type 2 diabetes at a**
23 **specialised Diabetes clinic in Gaborone (N= 500)]**
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27 **Optimal glycaemic control**

28 The mean (SD) HbA1c was 8.4 (2.4) % overall, 8.6 (2.7) % for female and 8.0 (1.6) % for male patients
29 (p=0.199) [Table 1]. The proportion of the patients receiving oral hypoglycaemic agents alone was
30 54.2%; 30% were treated with oral hypoglycaemic agents combined with insulin; 13.6% insulin alone and
31 2.2% were on a diet alone. Of all the 218 patients on insulin, 184 (84.4%) were on premix insulin. Of all
32 the 421 patients on oral hypoglycaemic agents, 411(97.6%) patients were on metformin, and 194 (46.1%)
33 patients were on a sulphonylurea. Compared to patients on other anti-diabetic medications, those on
34 insulin injections were more likely to be males. We noted optimal glycaemic control in 159 (32.3%)
35 patients, whose mean (SD) HbA1c was 6.1 (0.6) %. Age over 50 years was associated with optimal
36 glycaemic control (AOR 5.79; 95%CI 1.08 – 31.14). On the other hand, an increase in diabetes duration
37 was associated with lower odds of optimal glycaemic control (AOR 0.91; 95%CI 0.85 – 0.98). There was
38 no association between gender, anthropometry, diabetes medications or education on the prevalence of
39 optimal glycaemic control [Table 2].
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49 **[Table 2: Factors associated with optimal glycaemic control among patients with type 2 diabetes at**
50 **a specialised diabetes clinic in Gaborone (N= 500)]**
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Optimal hypertension control

There were 404 (80.8%) hypertensive patients [Table 1]. Age \geq 50 years (AOR 4.954; 95%CI 2.811– 8.732), increased WHR (AOR 3.874; 95%CI 1.723–8.709), eGFR (AOR 0.982; 95% 0.975 – 0.989), and a long diabetes duration (AOR= 1.068, 95%CI: 1.019– 1.119) were associated with hypertension. Seventeen (4.2%) hypertensive patients did not receive any antihypertensive medication. Of the 389 patients who received antihypertensive medicines, 219 (56.3%) received calcium channel blockers, 189 (48.6%) were treated with thiazides diuretics, 183(47.0%) with angiotensin-converting enzyme inhibitors (ACE inhibitors), and 74(19.0%) with angiotensin receptor blockers (ARBs). The proportions of patients receiving β - and α -blockers were 22.4% and 5.4% respectively.

Of the 389 patients who received antihypertensive medicines, optimal hypertension control was noted in 211 (54.2%) patients. Patients on ACE inhibitors had lower odds of optimal hypertension control, compared to those who were not treated by ACE inhibitors (AOR 0.35; 95% CI 0.14 – 0.85). There was no association between gender, anthropometry, or education on the prevalence of optimal hypertension control [Table 3].

[Table 3: Factors associated with optimal hypertension control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)]

Optimal LDL-C control

A total of 225 (45%) patients were receiving lipid-lowering drugs, mostly (96.4%) atorvastatin. Of these, 147 (65.3%) patients had LDL-C measurements available. Only 30 (20.4%) achieved the optimal LDL-C control target. Females were less likely to achieve optimal LDL-C control as compared to males, (AOR 0.211; 95% CI 0.077 – 0.578). There was no association of age, anthropometry, or education on the prevalence of optimal LDL-C control [Table 4].

[Table 4: Factors associated with optimal LDL-C control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)]

Discussion

This outpatient cross-sectional study showed a low proportion of patients with optimal control of glycaemic, hypertension, and LDL-C among Type 2 patients attending a diabetes clinic in Botswana. In the multivariate analysis, duration of diabetes and age above 50 years were significantly associated with the presence of an optimal glycaemic control. Being on ACEIs was inversely related to optimal

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3 hypertension control compared to those who were not using ACEIs. Females were less likely to attain
4 optimal LDL-C control than males.

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6 Only 32.3% of our participants achieved optimal glycaemic control. Similar findings have also been seen
7 among patients with diabetes in Africa.^{10 12 22 23}. The proportion of patients with optimal glycaemic
8 control in specialised diabetes care centers across six sub-Saharan African countries was reported to be
9 29%¹², despite using a lower cut-off value of 6.5%. Similarly, only 7-20% of patients have been said to
10 have attained optimal glycaemic control (HbA1c level < 7%) in South African diabetic clinics^{11 23}.
11 Suboptimal glycaemic is, therefore, a big concern not only in Botswana but also in other African
12 countries. In most of the findings, the majority of the patients have HbA1c > 8%, well above the
13 recommended target (<7%) required to avoid the development of microvascular and macrovascular
14 complications^{11 12}. This could explain the four and tenfold prevalence of sight-threatening diabetic
15 retinopathy and proliferative retinopathy found in African populations compared to European prevalence
16 is²⁴. Despite this poor glycaemic control level, few of our patients were on insulin suggesting clinical
17 inertia in response to low glycaemic control²⁵. Similar to other studies in Sub-Saharan Africa, the
18 likelihood of attaining optimal glycaemic control decreased as the duration of diabetes increased²⁶. There
19 is evidence of a progressive loss of beta-cell function with increasing diabetes duration⁶. Insulin
20 production progressively declines over time, leading to sub-optimal glycaemic control unless higher
21 dosages or additional agents are initiated⁶. Comparable to reports from other studies, our older patients
22 were more likely to achieve optimal glycaemic control than young ones^{27 28}. It is possible that young
23 patients are less likely to be compliant with medication and lifestyle modification as compared to their
24 older counterparts. However, we need to research this further before making any concrete statements
25 and instigating pertinent quality improvement programmes.

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27 We also found a high prevalence (80.8%) of hypertension among patients attending our specialised
28 diabetes clinic. A decade ago, the prevalence of hypertension among patients with diabetes in this setting
29 was 61.2%²⁹. Our findings may suggest an increasing burden of hypertension as seen globally, but also a
30 reflection of the improvement in the screening and diagnosis of hypertension over the past few years³⁰.
31 We are aware that comparable high frequencies of hypertension have been reported in other African
32 studies¹². Consistent with previous studies, patients with hypertension were older, more obese, and had
33 declining GFR and longer diabetes duration^{10 31}. Thiazide diuretics; calcium channel blockers and ACE
34 inhibitors were the three most used anti-hypertensive agents. This finding is in line with previous studies
35 and guidelines recommendations suggesting that thiazide diuretics and calcium channels blockers are
36 more effective than other anti-hypertensives in the black population^{16 17 31}. Optimal hypertension control
37 was observed in only 54.2% of the patients on anti-hypertensives, suggesting a need for efforts to improve
38 identification and control of hypertension. This low rate is a concern given the increased mortality if

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3 hypertension is not controlled⁵. Having said this, the proportion of patients with optimal hypertension
4 control in our population was superior to several studies in Africa, notwithstanding, the variation of the
5 definitions of optimal hypertension control across these studies^{10 12 13 31 32}. In the present study, the use of
6 ACE inhibitors was inversely associated with optimal hypertension control. Although ACEIs are
7 indicated for patients with diabetes and proteinuria, they are associated with a clinically significant lesser
8 reduction in both systolic and diastolic blood pressure in blacks. This could partly explain suboptimal
9 hypertension control among our majority black patients. There appeared to be no influence of gender on
10 hypertension control similar to other studies³³, which is encouraging as a recent systematic review found
11 that males in low and middle income countries (LMICs) were more likely to be non-adherent to their
12 medications^{34 35}.

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19 Less than half of the patients (45%) were on lipid-lowering drugs, mostly statins. This is not surprising as
20 the use of lipid-lowering drugs in Africa has been reported in as low as 3 – 13% of patients with diabetes
21 due to the limited access to these drugs as well as lack of facilities for monitoring lipid profile while on
22 treatment^{12 35 36}. It is, however, a concern as the reduced use of statins will increase mortality rates in
23 patients with diabetes⁷. In some countries, the issue of co-payments limits the prescription of expensive
24 medications like statins³⁷. Medicines are provided free of charge in Botswana; consequently, concerns
25 with medication adherence due to co-payments should not be a problem^{37 38}. Although the lack of co-
26 payments has led to the more excellent use of statins in Botswana than in other African countries, we
27 would expect the rate of statin prescriptions to mirror the high rates seen in Western countries³⁹.
28 Education and adherence to guidelines will possibly improve the prescription and use of statins by the
29 majority of patients with diabetes according to treatment guidelines^{16 18}. Even when statins were used, an
30 undesirably small proportion of our patients achieved guideline-recommended LDL-C target level. Sub-
31 optimal LDL-C control rates have also been reported across different settings, even in developed
32 countries^{8 12 40 41}. This is disappointing as achieving LDL-C reduction is said to be associated with the
33 highest cardiovascular risk reduction than hypertension and HbA1c reduction⁸. Inadequate patients'
34 adherence and possibly clinicians' under-dosage of statin for fear of potential side effects are some of the
35 factors that possibly explain sub-optimal LDL-C control in our patients⁸. Besides, clinicians may be
36 unaware of the current LDL-C as well as those of HbA1c and hypertension therapeutic goals^{16 18}.
37 Irrespective of the reason, there is an urgent need to instigate measures to meet guideline-recommended
38 therapeutic, and we have started to address this in our clinics. As reported in other studies, women were
39 less likely than men to achieve optimal LDL-C control⁴⁰. Although the reason for this gender difference is
40 not apparent, this information is significant for clinicians to pay attention to the management of women
41 with diabetes in Botswana and other African countries.
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Conclusion

In conclusion, there was a high prevalence of suboptimal attainment of recommended glycaemic, hypertension and LDL-C targets for patients with diabetes in the study settings. These findings call for urgent individual and health systems interventions to address the factors associated with sub-optimal control of the cardiovascular risk factors among patients with type 2 diabetes. This will be the subject of future initiatives and research in our clinic given the growing prevalence of patients with type 2 diabetes in Botswana.

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Competing Interests statement

Dr. Mwita has nothing to disclose.

Dr. Msafiri has nothing to disclose.

Dr. Omech has nothing to disclose.

Dr. Botsile has nothing to disclose.

Dr. Oyewo has nothing to disclose.

Dr. Godman has nothing to disclose.

Dr. Tshikuka has nothing to disclose.

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Author contributions

J.C.M was involved in the conception of the study, data analysis, drafting and revising the manuscript. B.O was involved in the conception of the study and critical revision of the manuscript. A.O, M.M participated in the conception of the study, data collection and critical revision of the manuscript. J.M.F participated in data analysis and manuscript review. E.B performed data entry and management, and critical revision of the manuscript. O.J.M, B.G, and J.G.T were involved in interpretation of the data and manuscript revision. All the authors granted a final approval of the version to be published; and agreed to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

Data sharing statement

The dataset analysed during the current study is available from the corresponding author on reasonable request, through the Princess Marina Hospital and Ministry of Health of Botswana ethical approval.

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Table 1: Clinical and socio-demographic characteristics of patients with type 2 diabetes at a specialised Diabetes clinic in Gaborone (N= 500)

Characteristics	All (N=500)	Males(n= 170)	Females(n= 330)	P value
Mean age (SD), years	58.9 ± 12.2	55.4 ± 12.6	60.8 ± 11.6	<0.001
Age <50 years n (%)	109(21.80)	57 (33.5)	52 (15.8)	<0.001
Age ≥ 50 years n (%)	391 (78.20)	113 (66.5)	278(84.2)	
Diabetes duration, median, IQR, years	6 (2 – 13)	6.5 (2 – 14)	6 (2.5 – 13)	0.927
HbA1c mean (SD), %	8.4 (2.4)	8.6 (2.7)	8.4 (2.4)	0.199
Diabetes treatment				
Diet alone n (%)	11(2.2)	3(1.8)	8 (2.4)	0.001
Insulin alone n (%)	68 (13.6)	38(22.3)	30 (9.1)	
OHA alone n (%)	271(54.2)	82(48.2)	189 (57.3)	
Insulin and OHA	150 (30.0)	47(27.7)	103(31.2)	
BMI, mean (SD) kg/m ²	30.5 ± 6.0	28.7 ± 5.2	31.4 ± 6.2	<0.001
Normal weight n (%)	93 (18.7)	46(27.1)	49 (14.9)	<0.001
Overweight n (%)	155(31.1)	61(36.3)	94 (28.5)	
Obese n (%)	250(50.2)	63 (37.1)	187(56.7)	
Marital status				
Living alone n (%)	266 (53.20)	55 (32.4)	211(63.9)	<0.001
Living with a partner n (%)	234(46.8)	115 (67.6)	119(36.01)	
Education status				
No formal education, n (%)	77(15.4)	26(15.3)	51 (15.5)	<0.001
Primary School, n (%)	229(45.8)	56 (32.9)	173 (52.4)	
Secondary school, n (%)	131(26.2)	55(32.4)	76 (23.0)	
College/University, n (%)	63(12.6)	33 (19.4)	30 (9.1)	
WC, mean (SD), cm	103.3 ±12.5	101.7 ± 11.9	104.0 ± 12.7	0.049
WHR				
Low WHR n (%)	75 (44.1)	16(4.9)		<0.001
High WHR n (%)	95 (55.9)	314(95.2)		
CKD n (%)	54 (10.8)	24 (14.1)	30 (9.1)	0.086
eGFR, Median, IQR (ml/min/1.73m ²)	112.5(84.3–138.1)	113.1(80.8 –139.5)	112.3(84.4 –137.9)	0.737
Total cholesterol mean (SD), mmol/L	4.4(1.1)	4.3(1.2)	4.5(1.1)	0.030
LDL-C mean (SD), mmol/L	2.8(1.0)	2.6(1.0)	2.9(1.0)	0.006
Macroproteinuria n (%)	51(10.20)	27 (15.9)	24(7.27)	0.003
Hypertension n (%)	404 (80.80)	120 (70.59)	284(86.06)	<0.001
Dyslipidaemia n (%)	358 (71.60)	114 (67.06)	244(73.94)	0.106

Legend: DM - Diabetes mellitus; WC- waist circumference; BMI- Body Mass Index; SD- standard deviation; IQR – interquartile range; eGFR- estimated glomerular filtration rate; LDL-C – Low-density lipoprotein; WHR – Waist-hip ratio, HbA1c- Haemoglobin A1c; CKD- Chronic kidney disease; OHA – oral hypoglycaemic agent

Table 2: Factors associated with optimal glycaemic control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)

Characteristic	Unadjusted OR(95% CI)	P value	Adjusted OR (95%CI)	P value
Age group				
> 50 years	1		1	
≤ 50 years	2.16(1.29 – 3.61)	0.003	5.79 (1.08 – 31.14)	0.041
Diabetes duration, years	0.97(0.95 – 0.99)	0.040	0.909 (0.85 – 0.98)	0.011
Diabetes treatment				
Insulin alone(yes)	0.80(0.45– 1.43)	0.450		
OHA alone(yes)	2.785 (1.862 – 4.167)	<0.001	0.90 (0.46 – 1.74)	0.745
Insulin and OHA(yes)	0.185 (0.124 – 0.356)	<0.001	0.34 (0.07 – 1.70)	0.188
BMI	0.99(0.96 – 1.02)	0.427		
Normal weight	1			
Overweight	1.10 (0.64– 1.90)	0.721		
Obese	0.83(0.50 – 1.39)	0.476		
Gender				
Men	1			
Women	0.92(0.62 – 1.36)	0.663	0.42 (0.14 – 1.25)	0.120
Education status				
None	1	1		
Primary School	1.58(0.88 – 2.81)	0.124		
Secondary school	1.14 (0.60 – 2.16)	0.687		
College/University	1.00(0.47 – 2.13)	0.996		
Marital status				
Living alone	1			
Living with a partner	0.93 (0.64 – 1.36)	0.703		
WHR	0.92 (0.13 – 6.58)	0.937		
Low WHR	1			
High WHR	0.92(0.57 – 1.50)	0.745		
Weight, kg	0.99 (0.98 – 1.01)	0.298		
eGFR, (ml/min/1.73m ²)	1.00 (0.99 – 1.00)	0.074	1.00(0.99 – 1.01)	0.766
CKD (yes)	1.053 (0.578 – 1.920)	0.866		
Macroproteinuria(yes)	1.16(0.63 – 2.14)	0.624		
Optimal hypertension control(yes)	2.53 (1.63 – 3.93)	<0.001	1.61(0.63 – 4.13)	0.322
Optimal LDL-C control(yes)	2.10 (0.90 – 4.88)	0.086	2.20 (0.64 – 7.57)	0.209

Legend: Legend: BMI- Body Mass Index; CKD- Chronic kidney disease; DM - Diabetes mellitus; eGFR- estimated glomerular filtration rate; IQR – interquartile range; LDL-C – Low-density lipoprotein; OHA – oral hypoglycaemic agent; SD- standard deviation; WHR – Waist-hip ratio.

Table 3: Factors associated with optimal hypertension control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)

Characteristic	Unadjusted OR (95% CI)	P value	Adjusted OR (95% CI)	P value
Age group				
< 50 years	1			
≥50 years	0.885(0.483 – 1.619)	0.691		
Diabetes duration, years	0.972 (0.947 – 0.997)	0.023	0.975 (0.923 – 1.029)	0.352
Diabetes treatment				
Insulin alone(yes)	0.446 (0.243 – 0.819)	0.009	1.150 (0.304 – 4.435)	0.837
OHA alone(yes)	1.940 (1.295 – 2.908)	0.001	1.370 (0.512 – 3.661)	0.531
Insulin and OHA(yes)	0.588 (0.403 – 0.949)	0.028	1.15(0.304 – 4.374)	0.837
BMI, kg/m ²				
Normal weight	1			
Overweight	0.786 (0.422 – 1.465)	0.449		
Obese	0.774(0.435 – 1.378)	0.384		
Gender				
Men	1			
Women	1.425(0.915 – 2.219)	0.117	0.968 (0.359 – 2.608)	0.949
Marital				
Living alone	1			
Living with partner	0.675 (0.614 –1.371)	0.668		
Education status				
No formal education	1			
Primary School	1.028(0.578 – 1.827)	0.09		
Secondary school	1.078(0.556 – 2.091)	0.22		
College/University	0.887(0.410 – 1.919)	0.761		
WHR	1.293 (0.139 – 11.997)	0.821		
Low WHR	1			
High WHR	1.177 (0.653 – 2.121)	0.589		
eGFR, (ml/min/1.73m ²)	1.004 (0.999 – 0 1.009)	0.139	1.004(0.994 – 1.015)	0.412
Serum Cholesterol	0.964 (0.804 – 1.156)	0.693		
Macroproteinuria	0.484(0.255 –0.921)	0.027	0.360(0.072 1.800) . .35961	0.213
Antihypertensive				
CCB(yes)	0.729(0.718 – 1.607)	0.729		
Thiazides(yes)	1.583(1.058 – 2.369)	0.026	1.444 (0.615 – 3.391)	0.399
ACEI(yes)	0.636 (0.425 – 0.952)	0.028	0.348 (0.142 – 0.849)	0.020
ARB(yes)	0.947(0.568 – 1.578)	0.834		
Alpha blocker(yes)	0.243(0.087 – 0.677)	0.007	0.756 (0.136 –4.195)	0.749
Beta blocker(yes)	0.702 (0.434 –1.135)	0.149	0.514(0.193 – 1.374)	0.184
Optimal Glycaemic control(yes)	2.531 (1.631 – 3.928)	<0.001	1.921 (0.706 –5.228)	0.201
Optimal LDL-C control(yes)	0.554 (0.225 – 1.363)	0.199	0.754 (0.245 – 2.320)	0.623

Legend: ACEI – Angiotensin-converting enzyme inhibitors; ARB – Angiotensin Receptor Blockers; BMI- Body Mass Index; CCB- Calcium Channel Blockers; DM - Diabetes mellitus; eGFR- estimated glomerular filtration rate; HbA1c- Haemoglobin A1c; IQR – interquartile range; LDL-C – Low-density lipoprotein; OHA – oral hypoglycaemic agent; SD- standard deviation; WHR – Waist-hip ratio.

Table 4: Factors associated with optimal LDL-C control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)

Characteristic	Unadjusted OR (95% CI)	P value	Adjusted OR (95% CI)	P value
Age group				
<50 years	1			
≥ 50 years	0.559 (0.208 – 1.503)	0.249		
Diabetes duration, years	1.016(0.968 – 1.064)	0.525		
Diabetes treatment				
Insulin alone(yes)	1.578 (0.559 – 4.458)	0.389		
OHA alone(yes)	0.95 (0.426 – 2.119)	0.9		
Insulin and OHA(yes)	0.825 (0.345 – 1.968)	0.664		
BMI	0.998 (0.933 –1.068)	0.951		
Normal weight	1			
Overweight	1.429 (0.344 – 5.940)	0.624		
Obese	1.269(0.329 – 4.889)	0.730		
Gender				
Men	1			
Women	0.2 (0.086 – 0.467)	<0.001	0.211 (0.077 – 0.578)	0.002
Education status				
No formal education	1	1		
Primary School	0.793 (0.294 – 2.140)	0.647		
Secondary school	0.284 (0.068 – 1.198)	0.087		
College/University	0.786(0.199 – 3.106)	0.731		
Marital status				
Living alone	1			
Living with a partner	.986 (0.441– 2.204)	0.973		
HbA1c	1.001(0.850 – 0.198)	0.917		
Low WHR	1			
High WHR	0.307(0.105 – 0.891)	0.030	1.699(0.361 – 7.982)	0.502
Macroproteinuria	0.688 (0.144 – 3.286)	0.64		
eGFR, (ml/min/1.73m ²)	0.999(0.991 – 1.008)	0.878		
CKD (yes)	1.8756(0.727 – 4.833)	0.193		
Hypertension (yes)	1.125 (0.299 – 4.232)	0.862		

Legend: Legend: BMI- Body Mass Index; CKD- Chronic kidney disease; DM - Diabetes mellitus; eGFR- estimated glomerular filtration rate; IQR – interquartile range; LDL-C – Low-density lipoprotein; OHA – oral hypoglycaemic agent; SD- standard deviation; WHR–Waist-hip ratio

STROBE 2007 (v4) Statement—Checklist of items that should be included in reports of *cross-sectional studies*

Section/Topic	Item #	Recommendation	Reported on page #
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract	1
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	1
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	2
Objectives	3	State specific objectives, including any prespecified hypotheses	2
Methods			
Study design	4	Present key elements of study design early in the paper	2
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	2
Participants	6	(a) Give the eligibility criteria, and the sources and methods of selection of participants	2
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	2
Data sources/ measurement	8*	For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group	2
Bias	9	Describe any efforts to address potential sources of bias	2
Study size	10	Explain how the study size was arrived at	2
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why	3
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	3
		(b) Describe any methods used to examine subgroups and interactions	3
		(c) Explain how missing data were addressed	
		(d) If applicable, describe analytical methods taking account of sampling strategy	
		(e) Describe any sensitivity analyses	
Results			3

Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed (b) Give reasons for non-participation at each stage (c) Consider use of a flow diagram	3
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders (b) Indicate number of participants with missing data for each variable of interest	4
Outcome data	15*	Report numbers of outcome events or summary measures	4,5
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and why they were included (b) Report category boundaries when continuous variables were categorized (c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	4,5
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	
Discussion			
Key results	18	Summarise key results with reference to study objectives	5
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	1
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	8
Generalisability	21	Discuss the generalisability (external validity) of the study results	8
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	8

*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at <http://www.plosmedicine.org/>, Annals of Internal Medicine at <http://www.annals.org/>, and Epidemiology at <http://www.epidem.com/>). Information on the STROBE Initiative is available at www.strobe-statement.org.

BMJ Open

Glycaemic, blood pressure and low-density lipoprotein-cholesterol control among patients with diabetes mellitus in a specialised clinic in Botswana: a cross-sectional study

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Secondary Subject Heading:	Cardiovascular medicine
Keywords:	Glycemic, Botswana, LDL, Hypertension < CARDIOLOGY, Diabetes Mellitus, control

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3 **Glycaemic, blood pressure and low-density lipoprotein-cholesterol control among patients with**
4 **diabetes mellitus in a specialised clinic in Botswana: a cross-sectional study**
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Abstract

Objective: Control of glycaemic, hypertension and low-density lipoprotein-cholesterol (LDL-C) among type 2 diabetes mellitus (T2DM) patients is vital for the prevention of cardiovascular diseases (CVD). The current study was an audit of glycaemic, hypertension, and LDL-C control among ambulant patients with type 2 diabetes mellitus in Botswana. Also, the study aimed at assessing factors associated with attaining optimal glycaemic, hypertension and LDL-C therapeutic goals.

Design: A cross-sectional study.

Setting: A specialised public diabetes clinic in Gaborone, Botswana.

Participants: Type 2 diabetes mellitus patients who had attended the clinic for \geq three months between August 2017 and February 2018.

Primary outcome measure: The proportion of patients with optimal glycaemic (HbA1c $<7\%$), hypertension (blood pressure $<140/90$ mmHg) and LDL-C (<1.8 mmol/L) control.

Results: The proportions of patients meeting optimal targets were 32.3% for glycaemic, 54.2% for hypertension, and 20.4% for LDL-C. Optimal glycaemic control was positively associated with age ≥ 50 years (AOR 5.79; 95% CI 1.08 - 31.14) but was inversely associated with an increase in diabetes duration (AOR 0.91; 95%CI 0.85 - 0.98). Being on an angiotensin-converting-enzyme inhibitor (ACE inhibitor) was inversely associated with optimal hypertension control (AOR 0.35; 95% CI 0.14 - 0.85). Being female (AOR 0.24; 95% CI (0.09 - 0.59) was inversely associated with optimal LDL-C control.

Conclusion: Patients with Type 2 diabetes mellitus in Gaborone, Botswana, presented with suboptimal control of recommended glycaemic, hypertension and LDL-C targets. These findings call for urgent individual and health systems interventions to address key determinants of the recommended therapeutic targets among patients with diabetes in this setting.

Keywords: Glycaemic, hypertension, LDL-C, cholesterol, control Diabetes Mellitus, Botswana

Strengths and limitations of this study

- The first study to objectively assess the three critical therapeutic targets in patients with diabetes in Botswana.
- The study was done in one specialised public diabetes clinic and may not be generalised to other facilities in the country. The clinic is, however, a leading diabetes care facility in the country; hence the results may predict the status in other facilities.
- The cross-sectional design limited the assessment of the temporal relationship between factors associated with poor control of glycaemia, LDL-C and hypertension.
- Systematic random sampling and incomplete data in some participants may not have yielded a representative sample of our clinic enrollees.

Introduction

Diabetes mellitus and related cardiovascular complications are growing public health concerns worldwide^{1,2}. There are approximately 16 million people with diabetes in Africa, and this number is projected to increase to 41 million by 2045 due to rapid urbanisation, lifestyles changes and nutrition transition in the continent¹. This increase in prevalence and incidence of diabetes is attributable to type 2 diabetes, which is associated with multiple comorbidities such as obesity and hypertension requiring chronic care and catastrophic health expenditure¹. Diabetes and associated comorbidities are known to increase patients' risk of developing cardiovascular diseases (CVD), which are responsible for approximately 70% of diabetes-related deaths³. The risk to the development of CVD is higher in people with suboptimal glycaemic, hypertension and LDL-C control^{3,4}. A reduction of HbA1c to control targets along with optimal hypertension control and the use of statins to lower LDL-C levels have been shown to improve long-term outcomes including reducing mortality among patients with diabetes⁵⁻⁸. Achieving these targets remains a challenge in most settings, especially those with limited access to standard diabetes care⁹⁻¹³. Only a minority of patients with diabetes in Africa achieves optimal therapeutic targets, leaving the majority of patients at high risk of diabetes-related complications¹². Suboptimal treatment to recommended targets is a public health concern because the current total health expenditure in most sub-Saharan African countries remains far below the 15% recommended in the Abuja declaration¹⁴. The rising cost of managing diabetes complications will further make health system goals unattainable¹. Thus, this study was an audit of glycaemic, hypertension, and LDL-C control among ambulant patients with type 2 diabetes mellitus in Botswana. The study also assessed factors associated with attainment of glycaemic, blood pressure and LDL-C therapeutic targets in these patients.

Methods

Study design and participants

We conducted a cross-sectional study of outpatients with established type 2 diabetes attending a specialised public diabetes clinic in Gaborone, Botswana between August 2017 and February 2018. The clinic has been operational since 2011 as a referral centre for health facilities in Gaborone and nearby towns. Eligible patients were those aged ≥ 18 years and had received care from the clinic for at least three months. Systematic random sampling was used to select patients from a list of patients who attended the clinic every day. We randomly picked the first patient from the first eight people in the queue. Subsequently, every eighth individual was enrolled until either the daily target of 10 patients was reached or the clinic came to an end. As there was a daily variation of the number of clinic attendees, the number of our daily enrolments varied as well. Participants provided informed consent before enrolment. The Ministry of Health (HPDME: 13/18/1 VOL XI) and Princess Marina Hospital (PMH 5/79,317-1-2017) Institutional Review Boards granted ethical approval for the study.

Data collection and procedures

Patient information was collected using an interviewer-administered questionnaire and through reviews of medical charts and electronic records. The information included: demographic data (age, gender, occupation, marital status, and education), diabetes duration, history of hypertension, and medications for diabetes, hypertension and lipid disorders. We performed anthropometry (weight, height, waist and hip circumferences) and blood pressure measurements at enrolment. We conducted three blood pressure measurements after 10 minutes of rest, and the mean of the three measurements was recorded¹⁵. Moreover, we documented blood pressure readings from each patient's prior visit. Patients' serum creatinine, LDL-C and HbA1c, and urine dipstick for proteinuria results over the past six months were abstracted from the electronic medical records.

Definitions of the key outcomes and exposure variables

We calculated the diabetes duration as the date of enrolment into the study minus the date of diabetes diagnosis. A patient was considered hypertensive by self-reported hypertension and the use of blood pressure-lowering medications or had sustained blood pressure $\geq 140/90$ mmHg during the previous visit and at enrolment¹⁶⁻¹⁸. Optimal glycaemic control was defined as HbA1c $< 7\%$ ^{16 18}. For patients who were on lipid-lowering medications, optimal LDL-C control was LDL cholesterol level < 1.8 mmol/L¹⁶. We calculated patients' eGFR by the Modification of Diet in Renal Disease (MDRD) formula, and an eGFR < 60.0 mL/min/1.73m² defined Chronic Kidney Disease (CKD)^{2 19 20}. Body mass index (BMI) was calculated as weight in kilograms divided by the square of height in meters. We considered underweight as BMI < 18.5 kg/m², normal weight as BMI of 18.5 –24.9 kg/m², overweight as BMI of 25.0 –29.9 kg/m², and obesity as BMI ≥ 30 kg/m²²¹. Waist-hip ratio (WHR) was calculated as waist circumference (WC) in centimetres divided by hip circumference (HC) in centimetres and classified as high when WHR was ≥ 0.85 and ≥ 0.90 for women and men respectively²¹.

Patient and Public Involvement

We did not directly involve patients in the design, recruitment to and conduct of the study. However, the development of the research question and outcome measures were informed by patients' priorities, experience, and preferences. These were realised during the regular diabetes support group meetings where the authors of this study interact with patients and their families. Investigators working at the clinic will discuss the study findings with colleagues and provide them with critical results for sharing with patients (study participants). In close collaboration with the patient support group, the investigation team will summarize the results in plain language for a large poster and place it in a waiting room.

Statistical analysis

We performed analyses using Stata, Version 14 (Stata Corp, College Station, TX). We used percentages to summarise categorical variables. Means and standard deviation (SD) or medians and interquartile range (IQR) were used to summarise continuous variables. Pearson's χ^2 or Fisher's exact tests were used to assessing statistical differences by gender for the categorical variables, while the Student's t-test or the Mann-Whitney U test were used for the continuous ones. Bivariate logistic regression was used to explore factors associated with each primary outcome - glycaemic, hypertension and LDL-C control. We further performed three multivariate logistic regression models for each of the three outcomes. The independent variables selected for multivariate models were those displaying a p-value < 0.2 at the univariate analysis level in addition to those considered clinically meaningful (age and gender). We described results as crude odds ratio (OR), adjusted odds ratio (AOR), and their corresponding 95% confidence intervals (95% CI). We used the Hosmer-Lemeshow goodness-of-fit test to assess how well the data fit the model²². A 2-tailed p-value < 0.05 was considered statistically significant.

Results

The response rate was 97%, as only 17 (3.4%) of the approached participants declined participation because of time constraints. We included 500 patients with type 2 diabetes in the study, of which 330 (66%) were females. The mean (SD) age was 58.9 (12.2) years, and 78.2% were aged > 50 years. The median (IQR) diabetes duration was 6 (2 - 13) years. There was a high percentage of patients with hypertension (80.8%), overweight (31.1%) and obesity (50.2%). Table 1 summarises the patients' characteristics by gender. Female patients tended to be older (60.8 vs 55.4 years, $p < 0.001$), obese (56.7% vs 37.1%, $p < 0.001$), have higher WHR (95.2% vs 55.9%), be hypertensive (86.1% vs 70.6%, $p < 0.001$), and have a higher mean total cholesterol (4.5mmol/L vs 4.3mmol/L, $p = 0.030$) and LDL-C (2.9 mmol/L vs 2.6 mmol/L, $p = 0.006$) than male patients. Urine dipstick was positive for protein in 10.2% of patients, mostly males (15.9% vs 7.3%, $p = 0.003$).

[Table 1: Clinical and socio-demographic characteristics of patients with type 2 diabetes at a specialised Diabetes clinic in Gaborone (N= 500)]

Optimal glycaemic control

The mean (SD) HbA1c was 8.4 % (2.4) overall, 8.6% (2.7) for female and 8.0% (1.6) for male patients ($p=0.199$) [Table 1]. The proportion of the patients receiving oral hypoglycaemic agents alone was 54.2%; 30% were on oral hypoglycaemic agents combined with insulin; 13.6% on insulin alone and 2.2% were on a diet alone. Of the 218 patients on insulin, 184 (84.4%) were on premix insulin. Of the 421 patients on oral hypoglycaemic agents, 411(97.6%) patients were on metformin, and 194 (46.1%) patients were on a sulphonylurea. Compared to patients on other anti-diabetic medications, those on insulin injections were

more likely to be males. We noted optimal glycaemic control in 159 (32.3%) patients, whose mean HbA1c was 6.1 %. Age over 50 years was associated with optimal glycaemic control (AOR 5.79; 95%CI 1.08 - 31.14). On the other hand, an increase in diabetes duration was inversely associated with optimal glycaemic control (AOR 0.91; 95%CI 0.85 - 0.98). There was no association between gender, anthropometry, diabetes medications or education on the level of glycaemic control [Table 2].

[Table 2: Factors associated with optimal glycaemic control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)]

Optimal hypertension control

There were 404 (80.8%) hypertensive patients [Table 1]. Age \geq 50 years (AOR 4.95; 95%CI 2.81- 8.73), increased WHR (AOR 3.87; 95%CI 1.72 - 8.71), eGFR (AOR 0.98; 95% CI 0.97 - 0.99), and a long diabetes duration (AOR= 1.07, 95% CI 1.02 - 1.12) were associated with hypertension. Seventeen (4.2%) hypertensive patients did not receive any antihypertensive medication. Of the 389 patients who received antihypertensive medications, 219 (56.3%) received calcium channel blockers, 189 (48.6%) were treated with thiazides diuretics, 183(47.0%) with angiotensin-converting-enzyme inhibitors (ACE inhibitors), and 74(19.0%) with Angiotensin II receptor blockers (ARBs). The proportions of patients receiving β - and α -blockers were 22.4% and 5.4% respectively.

Of the 389 patients who received antihypertensive medicines, optimal hypertension control was noted in 211 (54.2%) patients. Patients on ACE inhibitors were less likely to attain optimal hypertension control compared to those who were not on ACE inhibitors (AOR 0.24; 95% CI (0.09 - 0.59). There was no association between gender, anthropometry, or education on the level of hypertension control [Table 3].

[Table 3: Factors associated with optimal hypertension control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)]

Optimal LDL-C control

A total of 225 (45%) patients were receiving lipid-lowering drugs, mostly (96.4%) atorvastatin. Of these, 147 (65.3%) patients had LDL-C measurements available. Only 30 (20.4%) achieved the optimal LDL-C control target. Females were less likely to achieve optimal LDL-C control as compared to males, (0.24; 95% CI (0.09 - 0.59). There was no association of age, anthropometry, or education on the level of LDL-C control [Table 4].

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3 **[Table 4: Factors associated with optimal LDL-C control among patients with type 2 diabetes at a**
4 **specialised diabetes clinic in Gaborone (N= 500)]**
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7 **Discussion**
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9 This outpatient cross-sectional study showed a low proportion of patients with optimal control of glycaemic,
10 hypertension, and LDL-C among Type 2 patients attending a diabetes clinic in Botswana. In the multivariate
11 analysis, duration of diabetes and age above 50 years were significantly associated with optimal glycaemic
12 control. Being on ACE inhibitors was inversely related to optimal hypertension control. Females were less
13 likely to attain optimal LDL-C levels than males.
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16 Only 32.3% of our participants achieved optimal glycaemic control. Similarly, low levels of glycaemic
17 control have also been seen among patients with diabetes in Africa^{10 12 23 24}. The proportion of patients
18 with optimal glycaemic control (HbA1c < 6.5%) in specialised diabetes care centres across six sub-
19 Saharan African countries was reported to be 29%¹². Similarly, only 7-20% of patients attained optimal
20 glycaemic control (HbA1c level < 7%) in South African diabetic clinics^{11 24}.
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23 Consequently, suboptimal glycaemic levels are an apparent concern not only in Botswana but also in other
24 African countries. In most studies, the majority of the patients have HbA1c > 8%, well above the
25 recommended target (<7%) required to avoid the development of microvascular and macrovascular
26 complications^{11 12}. This suboptimal glycaemic control could explain the four and tenfold prevalence of
27 sight-threatening diabetic retinopathy and proliferative retinopathy respectively found among African
28 populations compared to their European counterparts²⁵. Despite poor glycaemic control level, only a
29 few of our patients were on insulin, suggesting clinical inertia of our clinicians in response to low
30 glycaemic control²⁶. We will be investigating this further given concerns with the lack of glycaemic control
31 in our patients. Similar to other studies in Sub-Saharan Africa, the likelihood of attaining optimal
32 glycaemic control decreased as the duration of diabetes increased²⁷. There is evidence of a progressive
33 loss of beta-cell function with increasing diabetes duration⁶. Insulin production progressively declines
34 over time, leading to suboptimal glycaemic control unless higher dosages or additional agents are initiated
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51 We also found a high prevalence (80.8%) of hypertension among patients attending our specialised diabetes
52 clinic. A decade ago, the prevalence of hypertension among patients with diabetes in this setting was 61.2%
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3 the improvement in the screening and diagnosis of hypertension over the past few years³¹. We are aware
4 of the comparable high frequencies of hypertension in other African studies¹². Consistent with previous
5 studies, patients with hypertension were older, more obese, and had declining GFR and longer diabetes
6 duration^{10 32}. Thiazide diuretics, calcium channel blockers and ACE inhibitors were the three most
7 prescribed antihypertensive agents. This finding is in line with the available evidence recommending
8 thiazide diuretics and calcium channels blockers as the most effective antihypertensives in the black
9 population^{16 17 32}. Optimal hypertension control was observed in only 54.2% of the patients on
10 antihypertensives, suggesting an urgent need for initiatives to improve the identification and control of
11 hypertension. This low control level is a concern given the increased mortality if hypertension is not
12 controlled⁵. Having said this, the proportion of patients with optimal hypertension control in our population
13 was superior to several studies in Africa, notwithstanding the variation of the definitions of optimal
14 hypertension control across these studies^{10 12 13 32 33}. However, there is no room for complacency. In the
15 present study, the use of ACE inhibitors was inversely associated with optimal hypertension control.
16 Although ACE inhibitors are indicated for patients with diabetes and proteinuria, they have a clinically
17 significant lesser reduction in both systolic and diastolic blood pressure in the black population. This could
18 partly explain suboptimal hypertension control among predominantly black patients in our study. There
19 appeared to be no influence of gender on hypertension control similar to other studies³⁴, which is
20 encouraging as a recent systematic review found that males in low and middle income countries (LMICs)
21 are more likely to be non-adherent to their medications^{35 36}.

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34 Less than half of the patients (45%) were on lipid-lowering drugs, mostly statins. This is not surprising as
35 the prescription of lipid-lowering medications in Africa is as low as 3 - 13% in patients with diabetes due
36 to the limited access to these drugs as well as lack of facilities for monitoring lipid profiles while patients
37 are on treatment^{12 36 37}. It is, however, a concern as the reduced use of statins will increase mortality rates
38 in patients with diabetes⁷. In some countries, the issue of co-payments limits the prescription of expensive
39 medications like statins³⁸. However, this is not an issue in Botswana where medications are provided free
40 of charge to patients. Although the lack of co-payments might have led to a higher prescription of statins
41 in Botswana than in other African countries, we would expect the rate of statin prescriptions to mirror the
42 high rates seen in Western countries³⁹. Education and adherence to guidelines will possibly improve the
43 prescription and use of statins among the majority of patients with diabetes according to treatment
44 guidelines^{16 18}. Even when statins were used, an undesirably small proportion of our patients achieved
45 guideline-recommended LDL-C target level. Suboptimal LDL-C control rates are also frequent across
46 different settings, even in developed countries^{8 12 40 41}. This is disappointing as achieving LDL-C reduction
47 is associated with the highest cardiovascular risk reduction than hypertension and HbA1c reduction⁸.
48 Inadequate patients' adherence and possibly clinicians' under-dosage of statin for fear of potential side
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3 effects are some of the factors that possibly explain suboptimal LDL-C control in our patients⁸. Besides,
4 clinicians may be unaware of the current LDL-C as well as those of HbA1c and hypertension therapeutic
5 goals^{16 18}. Irrespective of the reason, there is an urgent need to instigate measures to meet guideline-
6 recommended therapeutic goals, and we have started to address this in our clinic. As reported in other
7 studies, women were less likely than men to achieve optimal LDL-C control⁴⁰. Although the reason for this
8 gender difference is not apparent, this information is significant for clinicians to pay attention to the
9 management of women with diabetes in Botswana and other African countries.
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14 **Conclusion**

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16 In conclusion, there was suboptimal glycaemic, hypertension and LDL-C control among patients with
17 diabetes in our setting. These findings call for urgent individual and health systems interventions to address
18 the factors associated with suboptimal control of the cardiovascular risk factors among patients with type 2
19 diabetes. This will be the subject of future initiatives and research in our clinic given the growing prevalence
20 of patients with type 2 diabetes in Botswana.
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25
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28

29 **Competing Interests statement**

30 Dr Mwita has nothing to disclose.

31 Dr Msafiri has nothing to disclose.

32 Dr Omech has nothing to disclose.

33 Dr Botsile has nothing to disclose.

34 Dr Oyewo has nothing to disclose.

35 Dr Godman has nothing to disclose.

36 Dr Tshikuka has nothing to disclose.
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48 **Author contributions**

49 JCM was involved in the conception of the study, data analysis, drafting and revising the manuscript. BO
50 was involved in the design of the study and critical revision of the manuscript. AO, MM participated in the
51 design of the study, data collection and critical revision of the manuscript. JMF participated in data analysis
52 and manuscript review. EB performed data entry and management, and critical revision of the manuscript.
53 OJM, BG, and JGT were involved in the interpretation of the data and manuscript revision. All the authors
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3 granted final approval of the version to be published; and agreed to be accountable for all aspects of the
4 work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately
5 investigated and resolved.
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8 9 **Data sharing statement**

10 The dataset analysed during the current study is available from the corresponding author on reasonable
11 request, through the Princess Marina Hospital and Ministry of Health of Botswana ethical approval.
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Table 1: Clinical and socio-demographic characteristics of patients with type 2 diabetes at a specialised Diabetes clinic in Gaborone (N= 500)

Characteristics	All (N=500)	Males (n= 170)	Females (n= 330)	p-value
Mean age (SD), years	58.9 (12.2)	55.4 (12.6)	60.8(11.6)	<0.001
Age <50 years n (%)	109(21.80)	57 (33.5)	52 (15.8)	<0.001
Age ≥ 50 years n (%)	391 (78.20)	113 (66.5)	278(84.2)	
Diabetes duration, median, IQR, years	6 (2 - 13)	6.5 (2 - 14)	6 (2.5 - 13)	0.927
HbA1c mean (SD), %	8.4 (2.4)	8.6 (2.7)	8.4 (2.4)	0.199
Diabetes treatment				
Diet alone n (%)	11(2.2)	3(1.8)	8 (2.4)	0.001
Insulin alone n (%)	68 (13.6)	38(22.3)	30 (9.1)	
OHA alone n (%)	271(54.2)	82(48.2)	189 (57.3)	
Insulin and OHA	150 (30.0)	47(27.7)	103(31.2)	
BMI, mean (SD) kg/m ²	30.5(6.0)	28.7(5.2)	31.4(6.2)	<0.001
Normal weight n (%)	93 (18.7)	46(27.1)	49 (14.9)	<0.001
Overweight n (%)	155(31.1)	61(36.3)	94 (28.5)	
Obese n (%)	250(50.2)	63 (37.1)	187(56.7)	
Marital status				
Living alone n (%)	266 (53.20)	55 (32.4)	211(63.9)	<0.001
Living with a partner n (%)	234(46.8)	115 (67.6)	119(36.01)	
Education status				
No formal education, n (%)	77(15.4)	26(15.3)	51 (15.5)	<0.001
Primary School, n (%)	229(45.8)	56 (32.9)	173 (52.4)	
Secondary school, n (%)	131(26.2)	55(32.4)	76 (23.0)	
College/University, n (%)	63(12.6)	33 (19.4)	30 (9.1)	
WC, mean (SD), cm	103.3(12.5)	101.7 (11.9)	104.0 (12.7)	0.049
WHR	0.94(0.10)	0.97(0.09)	0.93(0.10)	<0.001
Low WHR n (%)	75 (44.1)	16(4.9)		<0.001
High WHR n (%)	95 (55.9)	314(95.2)		
CKD n (%)	54 (10.8)	24 (14.1)	30 (9.1)	0.086
eGFR, Median, IQR (ml/min/1.73m ²)	112.5(84.3-138.1)	113.1(80.8 -139.5)	112.3(84.4 - 137.9)	0.737
Cholesterol, mean (SD), mmol/L	4.4(1.1)	4.3(1.2)	4.5(1.1)	0.030
LDL-C mean (SD), mmol/L	2.8(1.0)	2.6(1.0)	2.9(1.0)	0.006
Macroproteinuria n (%)	51(10.20)	27 (15.9)	24(7.27)	0.003
Hypertension n (%)	404 (80.80)	120 (70.59)	284(86.06)	<0.001
Dyslipidaemia n (%)	358 (71.60)	114 (67.06)	244(73.94)	0.106

Legend: DM - Diabetes mellitus; WC- waist circumference; BMI- Body Mass Index; SD- standard deviation; IQR – interquartile range; eGFR- estimated glomerular filtration rate; LDL-C – Low-density lipoprotein; WHR – Waist-hip ratio, HbA1c- Haemoglobin A1c; CKD- Chronic kidney disease; OHA – oral hypoglycaemic agent

Table 2: Factors associated with optimal glycaemic control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)

Characteristic	Unadjusted OR (95% CI)	p-value	Adjusted OR (95%CI)	p-value
Age group				
> 50 years	1			
≤ 50 years	2.16(1.29 - 3.61)	0.003	5.79 (1.08 - 31.14)	0.041
Diabetes duration, years	0.97(0.95 - 0.99)	0.040	0.91 (0.85 - 0.98)	0.011
Use of Insulin alone				
No	1			
Yes	0.80(0.45- 1.43)	0.590		
Use of OHA alone				
No	1			
Yes	2.785 (1.862 - 4.167)	<0.001	0.90 (0.46 - 1.74)	0.745
Use of Insulin plus OHA				
No	1			
Yes	0.185 (0.124 - 0.356)	<0.001	0.34 (0.07 - 1.70)	0.188
BMI	0.99(0.96 - 1.02)	0.427	–	–
Normal weight	1	–	–	–
Overweight	1.10 (0.64 - 1.90)	0.721	–	–
Obese	0.83(0.50 - 1.39)	0.476	–	–
Gender			–	–
Men	1	–	–	–
Women	0.92(0.62 - 1.36)	0.663	0.42 (0.14 - 1.25)	0.120
Education status				
None	1	1	–	–
Primary School	1.58(0.88 - 2.81)	0.124	–	–
Secondary school	1.14 (0.60 - 2.16)	0.687	–	–
College/University	1.00(0.47 - 2.13)	0.996	–	–
Marital status				
Living alone	1	–	–	–
Living with a partner	0.93 (0.64 - 1.36)	0.703	–	–
WHR	0.92 (0.13 - 6.58)	0.937	–	–
Low WHR	1	–	–	–
High WHR	0.92(0.57 - 1.50)	0.745	–	–
Weight, kg	0.99 (0.98 - 1.01)	0.298	–	–
eGFR, (ml/min/1.73m ²)	1.00 (0.99 - 1.00)	0.074	1.00(0.99 - 1.01)	0.766
CKD				
No	1			
Yes	1.053 (0.578 - 1.920)	0.866	–	–
Macroproteinuria				
No	1		–	
Yes	1.16(0.63 - 2.14)	0.624	–	–
Optimal hypertension control				
No	1		–	
Yes	2.53 (1.63 - 3.93)	<0.001	1.61(0.63 - 4.13)	0.322
Optimal LDL-C control				
No	1			
Yes	2.10 (0.90 - 4.88)	0.086	2.20 (0.64 - 7.57)	0.209

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Legend: Legend: BMI- Body Mass Index; CKD- Chronic kidney disease; DM - Diabetes mellitus; eGFR- estimated glomerular filtration rate; IQR – interquartile range; LDL-C – Low-density lipoprotein; OHA – oral hypoglycaemic agent; SD- standard deviation; WHR – Waist-hip ratio.

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Table 3: Factors associated with optimal hypertension control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)

Characteristic	Unadjusted OR (95% CI)	p-value	Adjusted OR (95% CI)	p-value
Age group				
< 50 years	1	–	–	–
≥50 years	0.89(0.48 - 1.62)	0.691	–	–
Diabetes duration, years	0.97(0.94 - 0.99)	0.023	0.98 (0.92 - 1.03)	0.352
Use of Insulin alone				
No	1	–	–	–
Yes	0.45 (0.24 - 0.82)	0.009	1.15 (0.30 - 4.44)	0.837
Use of OHA alone				
No	1	–	–	–
Yes	1.94 (1.30 - 2.91)	0.001	1.37 (0.51 - 3.66)	0.531
Use of Insulin plus OHA				
No	1	–	–	–
Yes	0.59 (0.40 - 0.95)	0.028*	–	–
BMI, kg/m ²	0.98 (0.94 - 1.01)	0.149	0.93(0.86 - 1.01)	
Normal weight	1	–	–	–
Overweight	0.79 (0.42 - 1.47)	0.449	–	–
Obese	0.77(0.44 -1.38)	0.384	–	–
Gender				
Men	1	–	–	–
Women	1.43(0.92 - 2.22)	0.117	0.97 (0.36 - 2.61)	0.949
Marital				
Living alone	1	–	–	–
Living with a partner	0.68 (0.61 - 1.37)	0.67	–	–
Education status				
No formal education	1	–	–	–
Primary School	1.03(0.58 - 1.83)	0.09	–	–
Secondary school	1.08(0.56 - 2.09)	0.22	–	–
College/University	0.89(0.41 - 1.92)	0.761	–	–
WHR	1.29 (0.14 - 12.00)	0.821	–	–
Low WHR	1	–	–	–
High WHR	1.18 (0.65 - 2.12)	0.589	–	–
eGFR _c (ml/min/1.73m ²)	1.00 (1.00 - 1.01)	0.139	1.00(0.99 -1.02)	0.412
Serum Cholesterol	0.964 (0.80 - 1.16)	0.693	–	–
Macroproteinuria				
No	1	–	–	–
Yes	0.48(0.26 - 0.92)	0.027	0.36(0.07 - 1.80)	0.213
Use of CCB				
No	1	–	–	–
Yes	0.73(0.72 - 1.61)	0.729	–	–
Use of Thiazides				
No	1	–	–	–
Yes	1.58(1.06 - 2.37)	0.026	1.44(0.62 - 3.39)	0.399
Use of ACE INHIBITORS				
No	1	–	–	–
Yes	0.64 (0.43 - 0.95)	0.028	0.35(0.14 - 0.85)	0.020
Use of ARB				
No	1	–	–	–

	Yes	0.95(0.57 - 1.58)	0.834	–	–
Alpha blocker					
	No	1	–	–	–
	Yes	0.24(0.09 - 0.68)	0.007	0.76 (0.14 - 4.20)	0.749
Beta-blocker					
	No	1	–	–	–
	Yes	0.70 (0.43 -1.14)	0.149	0.51(0.19 - 1.37)	0.184
Optimal Glycaemic control					
	No	1	–	–	–
	Yes	2.53 (1.63- 3.93)	<0.001	1.92 (0.71 - 5.23)	0.201
Optimal LDL-C control					
	No	1	–	–	–
	Yes	0.55 (0.23 - 1.36)	0.199	0.75 (0.25 - 2.32)	0.623

Legend: * omitted because of collinearity. ACE INHIBITORS – angiotensin-converting-enzyme inhibitor; ARB – Angiotensin II receptor blocker; BMI- Body Mass Index; CCB- Calcium Channel Blockers; DM - Diabetes mellitus; eGFR- estimated glomerular filtration rate; HbA1c- Haemoglobin A1c; IQR – interquartile range; LDL-C – Low-density lipoprotein; OHA – oral hypoglycaemic agent; SD- standard deviation; WHR – Waist-hip ratio.

Table 4: Factors associated with optimal LDL-C control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)

Characteristic	Unadjusted OR (95% CI)	p-value	Adjusted OR (95% CI)	p-value
Age group				
<50 years	1		–	–
≥ 50 years	0.56 (0.21 - 1.50)	0.249	–	–
Diabetes duration, years	1.02(0. 97 - 1.06)	0.525		–
Use of Insulin alone				
No	1	–	–	–
Yes	1.58 (0.56 - 4.46)	0.389	–	–
Use of OHA alone				
No	1	–	–	–
Yes	0.95 (0.43 - 2.12)	0.900	–	–
Use of Insulin plus OHA				
No	1	–	–	–
Yes	0.83 (0.35 - 1.97)	0.664	–	–
BMI	1.00 (0.93 -1.07)	0.951		–
Normal weight	1	–	–	–
Overweight	1.43 (0.34 - 5.94)	0.624	–	–
Obese	1.27(0.33 - 4.89)	0.730	–	–
Gender				
Men	1		–	–
Women	0.2 (0.09 - 0.47)	<0.001	0.24(0.09- 0.59)	0.002
Education status				
No formal education	1	1	–	–
Primary School	0.79 (0. 29 - 2.14)	0.647	–	–
Secondary school	0.28 (0.07 - 1.20)	0.087	–	–
College/University	0.79(0.20 - 3.11)	0.731	–	–
Marital status				
Living alone	1	–	–	–
Living with a partner	0.99 (0.44 - 2.20)	0.973	–	–
HbA1c	1.00 (0.85 - 0.20)	0.917		–
WHR	7.59(0.19 - 303.60)	0.281	–	–
Low WHR	1			
High WHR	0.31(0.11 – 0.89)	0.030	0.64(0.20 - 2.10)	0.463
Macroproteinuria				
No		–	–	–
Yes	0.69 (0.14 - 3.27)	0.64	–	–
eGFR, (ml/min/1.73m ²)	1.00(0.99 - 1.01)	0.878	–	–
CKD				
No	1	–	–	–
Yes	1.88(0. 73 - 4.83)	0.193	1.67 (0.61 - 4.58)	0.321
Hypertension				
No		–	–	–
Yes	1.13 (0.30 - 4.23)	0.86	–	–

Legend: Legend: BMI- Body Mass Index; CKD- Chronic kidney disease; DM - Diabetes mellitus; eGFR- estimated glomerular filtration rate; IQR – interquartile range; LDL-C – Low-density lipoprotein; OHA – oral hypoglycaemic agent; SD- standard deviation; WHR–Waist-hip ratio

STROBE 2007 (v4) Statement—Checklist of items that should be included in reports of *cross-sectional studies*

Section/Topic	Item #	Recommendation	Reported on page #
Title and abstract	1	(a) Indicate the study’s design with a commonly used term in the title or the abstract	1
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	1
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	2
Objectives	3	State specific objectives, including any prespecified hypotheses	2
Methods			
Study design	4	Present key elements of study design early in the paper	2
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	2
Participants	6	(a) Give the eligibility criteria, and the sources and methods of selection of participants	2
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	2
Data sources/ measurement	8*	For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group	2
Bias	9	Describe any efforts to address potential sources of bias	2
Study size	10	Explain how the study size was arrived at	2
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why	3
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	3
		(b) Describe any methods used to examine subgroups and interactions	3
		(c) Explain how missing data were addressed	
		(d) If applicable, describe analytical methods taking account of sampling strategy	
		(e) Describe any sensitivity analyses	
Results			3

Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed (b) Give reasons for non-participation at each stage (c) Consider use of a flow diagram	3
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders (b) Indicate number of participants with missing data for each variable of interest	4
Outcome data	15*	Report numbers of outcome events or summary measures	4,5
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and why they were included (b) Report category boundaries when continuous variables were categorized (c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	4,5
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	
Discussion			
Key results	18	Summarise key results with reference to study objectives	5
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	1
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	8
Generalisability	21	Discuss the generalisability (external validity) of the study results	8
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	8

*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at <http://www.plosmedicine.org/>, Annals of Internal Medicine at <http://www.annals.org/>, and Epidemiology at <http://www.epidem.com/>). Information on the STROBE Initiative is available at www.strobe-statement.org.

BMJ Open

Glycaemic, blood pressure and low-density lipoprotein-cholesterol control among patients with diabetes mellitus in a specialised clinic in Botswana: a cross-sectional study

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Secondary Subject Heading:	Cardiovascular medicine
Keywords:	Glycemic, Botswana, LDL, Hypertension < CARDIOLOGY, Diabetes Mellitus, control

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3 **Glycaemic, blood pressure and low-density lipoprotein-cholesterol control among patients with**
4 **diabetes mellitus in a specialised clinic in Botswana: a cross-sectional study**

5
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Abstract

Objective: Control of glycaemic, hypertension and low-density lipoprotein-cholesterol (LDL-C) among type 2 diabetes mellitus (T2DM) patients is vital for the prevention of cardiovascular diseases (CVD). The current study was an audit of glycaemic, hypertension, and LDL-C control among ambulant patients with type 2 diabetes mellitus in Botswana. Also, the study aimed at assessing factors associated with attaining optimal glycaemic, hypertension and LDL-C therapeutic goals.

Design: A cross-sectional study.

Setting: A specialised public diabetes clinic in Gaborone, Botswana.

Participants: Type 2 diabetes mellitus patients who had attended the clinic for \geq three months between August 2017 and February 2018.

Primary outcome measure: The proportion of patients with optimal glycaemic (HbA1c $<7\%$), hypertension (blood pressure $<140/90$ mmHg) and LDL-C (<1.8 mmol/L) control.

Results: The proportions of patients meeting optimal targets were 32.3% for glycaemic, 54.2% for hypertension, and 20.4% for LDL-C. Optimal glycaemic control was positively associated with age ≥ 50 years (AOR 5.79; 95% CI 1.08 - 31.14) but was inversely associated with an increase in diabetes duration (AOR 0.91; 95%CI 0.85 - 0.98). Being on an angiotensin-converting-enzyme inhibitor (ACE inhibitor) was inversely associated with optimal hypertension control (AOR 0.35; 95% CI 0.14 - 0.85). Being female (AOR 0.24; 95% CI (0.09 - 0.59) was inversely associated with optimal LDL-C control.

Conclusion: Patients with Type 2 diabetes mellitus in Gaborone, Botswana, presented with suboptimal control of recommended glycaemic, hypertension and LDL-C targets. These findings call for urgent individual and health systems interventions to address key determinants of the recommended therapeutic targets among patients with diabetes in this setting.

Keywords: Glycaemic, hypertension, LDL-C, cholesterol, control Diabetes Mellitus, Botswana

Strengths and limitations of this study

- The first study to objectively assess the three critical therapeutic targets in patients with diabetes in Botswana.
- The study was undertaken in one specialised public diabetes clinic and the findings may not be generalised to other facilities in the country..
- The cross-sectional design limited the assessment of the temporal relationship between factors associated with poor control of glycaemia, LDL-C and hypertension.
- Systematic random sampling and incomplete data in some participants may not have yielded a representative sample of our clinic enrolees.

Introduction

Diabetes mellitus and related cardiovascular complications are growing public health concerns worldwide^{1,2}. There are approximately 16 million people with diabetes in Africa, and this number is projected to increase to 41 million by 2045 due to rapid urbanisation, lifestyles changes and nutrition transition in the continent¹. This increase in prevalence and incidence of diabetes is attributable to type 2 diabetes, which is associated with multiple comorbidities such as obesity and hypertension requiring chronic care and catastrophic health expenditure¹. Diabetes and associated comorbidities are known to increase patients' risk of developing cardiovascular diseases (CVD), which are responsible for approximately 70% of diabetes-related deaths^{3,4}. The risk to the development of CVD is higher in people with suboptimal glycaemic, hypertension and LDL-C control^{3,5}. A reduction of HbA1c to control targets along with optimal hypertension control and the use of statins to lower LDL-C levels have been shown to improve long-term outcomes including reducing mortality among patients with diabetes⁶⁻⁹. Achieving these targets remains a challenge in most settings, especially those with limited access to standard diabetes care¹⁰⁻¹⁶. Only a minority of patients with diabetes in Africa achieves optimal therapeutic targets, leaving the majority of patients at high risk of diabetes-related complications^{13,15}. Suboptimal treatment to recommended targets is a public health concern because the current total health expenditure in most sub-Saharan African countries remains far below the 15% recommended in the Abuja declaration¹⁷. The rising cost of managing diabetes complications will further make health system goals unattainable¹. Thus, this study was an audit of glycaemic, hypertension, and LDL-C control among ambulant patients with type 2 diabetes mellitus in Botswana. The study also assessed factors associated with the attainment of glycaemic, blood pressure and LDL-C therapeutic targets in these patients.

Methods

Study design and participants

We conducted a cross-sectional study of outpatients with established type 2 diabetes attending a specialised public diabetes clinic in Gaborone, Botswana between August 2017 and February 2018. The clinic has been operational since 2011 as a referral centre for health facilities in Gaborone and nearby towns. Eligible patients were those aged ≥ 18 years and had received care from the clinic for at least three months. We needed a sample size of 500 to produce a two-sided 95% confidence interval with a width equal to 3.86% based on the assumptions of approximately 26.2% glycaemic control among patients with type 2 diabetes in Botswana¹⁶. Systematic random sampling was used to select patients from a list of patients who attended the clinic every day. In a recruitment day, we randomly picked the first patient from the list of the first eight clinic attendees. Subsequently, we enrolled every eighth individual until either the daily target of 10 patients was reached or the clinic came to an end. As there was a daily variation of

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3 the number of clinic attendees, the number of our daily enrolments varied as well. Participants provided
4 informed consent before enrolment. The Ministry of Health (HPDME: 13/18/1 VOL XI) and Princess
5 Marina Hospital (PMH 5/79,317-1-2017) Institutional Review Boards granted ethical approval for the
6 study.
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10 **Data collection and procedures**

11 Patient information was collected using an interviewer-administered questionnaire and through reviews of
12 medical charts and electronic records. The information included: demographic data (age, gender,
13 occupation, marital status, and education), diabetes duration, history of hypertension, and medications for
14 diabetes, hypertension and lipid disorders. We performed anthropometry (weight, height, waist and hip
15 circumferences) and blood pressure measurements at enrolment. We conducted three blood pressure
16 measurements after 10 minutes of rest, and the mean of the three measurements was recorded¹⁸. Moreover,
17 we documented blood pressure readings from each patient's previous visit. Patients' serum creatinine, LDL-
18 C and HbA1c, and urine dipstick for proteinuria results over the past six months were abstracted from the
19 electronic medical records.
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26 **Definitions of the key outcomes and exposure variables**

27 We calculated the diabetes duration as the date of enrolment into the study minus the date of a diabetes
28 diagnosis. A patient was considered hypertensive by self-reported hypertension and the use of blood
29 pressure-lowering medications or had sustained blood pressure $\geq 140/90$ mmHg during the previous visit
30 and at enrolment¹⁹⁻²¹. Optimal glycaemic control was defined as HbA1c $< 7\%$ ^{19 21}. For patients who were
31 on lipid-lowering medications, optimal LDL-C control was LDL cholesterol level < 1.8 mmol/L¹⁹. We
32 calculated patients' eGFR by the Modification of Diet in Renal Disease (MDRD) formula, and an eGFR
33 < 60.0 mL/min/1.73m² defined Chronic Kidney Disease (CKD)^{22 23}. Body mass index (BMI) was
34 calculated as weight in kilograms divided by the square of height in meters. We considered underweight as
35 BMI < 18.5 kg/m², normal weight as BMI of 18.5-24.9 kg/m², overweight as BMI of 25. -29.9 kg/m², and
36 obesity as BMI ≥ 30 kg/m²²⁴. Waist-hip ratio (WHR) was calculated as waist circumference (WC) in
37 centimetres divided by hip circumference (HC) in centimetres and classified as high when WHR was ≥ 0.85
38 and ≥ 0.90 for women and men respectively²⁴.
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48 **Patient and Public Involvement**

49 We did not directly involve patients in the design, recruitment to and conduct of the study. However, the
50 development of the research question and outcome measures were informed by patients' priorities,
51 experience, and preferences. These were realised during the regular diabetes support group meetings where
52 the authors of this study interact with patients and their families. Investigators working at the clinic will
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3 discuss the study findings with colleagues and provide them with critical results for sharing with patients
4 (study participants). In close collaboration with the patient support group, the investigation team will
5 summarise the results in plain language for a large poster and place it in a waiting room.
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8 **Statistical analysis**

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10 We performed analyses using Stata, Version 14 (Stata Corp, College Station, TX). We used percentages to
11 summarise categorical variables. Means and standard deviation (SD) or medians and interquartile range
12 (IQR) were used to summarise continuous variables. Pearson's χ^2 or Fisher's exact tests were used to
13 assessing statistical differences by gender for the categorical variables, while the Student's t-test or the
14 Mann-Whitney U test were used for the continuous ones. Bivariate logistic regression was used to explore
15 factors associated with each primary outcome - glycaemic, hypertension and LDL-C control. We further
16 performed three multivariate logistic regression models for each of the three outcomes. The independent
17 variables selected for multivariate models were those displaying a p-value < 0.2 at the univariate analysis
18 level in addition to those considered clinically meaningful (age and gender). We described results as crude
19 odds ratio (OR), adjusted odds ratio (AOR), and their corresponding 95% confidence intervals (95% CI).
20 We used the Hosmer-Lemeshow goodness-of-fit test to assess how well the data fit the model ²⁵.
21 A 2-tailed p-value < 0.05 was considered statistically significant.
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30 **Results**

31 The response rate was 97%, as only 17 (3.4%) of the approached participants declined participation because
32 of time constraints. We included 500 patients with type 2 diabetes in the study, of which 330 (66%) were
33 females. The mean (SD) age was 58.9 (12.2) years, and 78.2% were aged > 50 years. The median (IQR)
34 diabetes duration was 6 (2 - 13) years. There was a high percentage of patients with hypertension (80.8%),
35 overweight (31.1%) and obesity (50.2%). Table 1 summarises the patients' characteristics by gender.
36 Female patients tended to be older (60.8 vs 55.4 years, $p < 0.001$, obese (56.7% vs 37.1%, $p < 0.001$), have
37 higher WHR (95.2% vs 55.9%), be hypertensive (86.1% vs 70.6%, $p < 0.001$), and have a higher mean total
38 cholesterol (4.5mmol/L vs 4.3mmol/L, $p = 0.030$) and LDL-C (2.9 mmol/L vs 2.6 mmol/L, $p = 0.006$) than
39 male patients. Urine dipstick was positive for protein in 10.2% of patients, mostly males (15.9% vs 7.3%,
40 $p = 0.003$).
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49 **[Table 1: Clinical and socio-demographic characteristics of patients with type 2 diabetes at a**
50 **specialised Diabetes clinic in Gaborone (N= 500)]**
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52 **Optimal glycaemic control**

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3 The mean (SD) HbA1c was 8.4 % (2.4) overall, 8.6% (2.7) for female and 8.0% (1.6) for male patients
4 (p=0.199) [Table 1]. The proportion of the patients receiving oral hypoglycaemic agents alone was 54.2%;
5 30% were on oral hypoglycaemic agents combined with insulin; 13.6% on insulin alone and 2.2% were on
6 a diet alone. Of the 218 patients on insulin, 184 (84.4%) were on premix insulin. Of the 421 patients on oral
7 hypoglycaemic agents, 411(97.6%) patients were on metformin, and 194 (46.1%) patients were on a
8 sulphonylurea. Compared to patients on other anti-diabetic medications, those on insulin injections were
9 more likely to be males. We noted optimal glycaemic control in 159 (32.3%) patients, whose mean) HbA1c
10 was 6.1 %. Age over 50 years was associated with optimal glycaemic control (AOR 5.79; 95%CI 1.08 -
11 31.14). On the other hand, an increase in diabetes duration was inversely associated with optimal glycaemic
12 control (AOR 0.91; 95%CI 0.85 - 0.98). There was no association between gender, anthropometry, diabetes
13 medications or education on the level of glycaemic control [Table 2].
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22 **[Table 2: Factors associated with optimal glycaemic control among patients with type 2 diabetes at**
23 **a specialised diabetes clinic in Gaborone (N= 500)]**
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27 **Optimal hypertension control**

28 There were 404 (80.8%) hypertensive patients [Table 1]. Age \geq 50 years (AOR 4.95; 95%CI 2.81-
29 8.73), increased WHR (AOR 3.87; 95%CI 1.72 - 8.71), eGFR (AOR 0.98; 95% CI 0.97 - 0.99),
30 and a long diabetes duration (AOR= 1.07, 95% CI 1.02 - 1.12) were associated with hypertension.
31 Seventeen (4.2%) hypertensive patients did not receive any antihypertensive medication. Of the
32 389 patients who received antihypertensive medications, 219 (56.3%) received calcium channel
33 blockers, 189 (48.6%) were treated with thiazides diuretics, 183(47.0%) with angiotensin-
34 converting-enzyme inhibitors (ACE inhibitors), and 74(19.0%) with Angiotensin II receptor
35 blockers (ARBs). The proportions of patients receiving β - and α -blockers were 22.4% and 5.4%
36 respectively.
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39 Of the 389 patients who received antihypertensive medicines, optimal hypertension control was
40 noted in 211 (54.2%) patients. Patients on ACE inhibitors were less likely to attain optimal
41 hypertension control compared to those who were not on ACE inhibitors (AOR 0.24; 95% CI (0.09
42 - 0.59). There was no association between gender, anthropometry, or education on the level of
43 hypertension control [Table 3].
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53 **[Table 3: Factors associated with optimal hypertension control among patients with type 2 diabetes**
54 **at a specialised diabetes clinic in Gaborone (N= 500)]**
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Optimal LDL-C control

A total of 225 (45%) patients were receiving lipid-lowering drugs, mostly (96.4%) atorvastatin. Of these, 147 (65.3%) patients had LDL-C measurements available. Only 30 (20.4%) achieved the optimal LDL-C control target. Females were less likely to achieve optimal LDL-C control as compared to males, (0.24; 95% CI (0.09 - 0.59). There was no association of age, anthropometry, or education on the level of LDL-C control [Table 4].

[Table 4: Factors associated with optimal LDL-C control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)]

Discussion

This outpatient cross-sectional study showed a low proportion of patients with optimal control of glycaemic, hypertension, and LDL-C among Type 2 patients attending a diabetes clinic in Botswana. In the multivariate analysis, duration of diabetes and age above 50 years were significantly associated with optimal glycaemic control. Being on ACE inhibitors was inversely related to optimal hypertension control. Females were less likely to attain optimal LDL-C levels than males.

Only 32.3% of our participants achieved optimal glycaemic control. Similarly, low levels of glycaemic control have also been seen among patients with diabetes in Africa^{11 13 26 27}. The proportion of patients with optimal glycaemic control (HbA1c < 6.5%) in specialised diabetes care centres across six sub-Saharan African countries was reported to be 29%¹³. Similarly, only 7-31% of patients attained optimal glycaemic control (HbA1c level < 7%) in other settings in Africa^{12 14 27}. Consequently, suboptimal glycaemic levels are an apparent concern not only in Botswana but also in other African countries. In most studies, the majority of the patients have HbA1c > 8%, well above the recommended target (<7%) required to avoid the development of microvascular and macrovascular complications^{12 13}. This suboptimal glycaemic control could explain the four and tenfold prevalence of sight-threatening diabetic retinopathy and proliferative retinopathy respectively found among African populations compared to their European counterparts²⁸. Despite poor glycaemic control level, only a few of our patients were on insulin, suggesting clinical inertia of our clinicians in response to low glycaemic control²⁹. We will be investigating this further given concerns with the lack of glycaemic control in our patients. Similar to other studies in Sub-Saharan Africa, the likelihood of attaining optimal glycaemic control decreased as the duration of diabetes increased³⁰. There is evidence of a progressive loss of beta-cell function with increasing diabetes duration⁷. Insulin production progressively declines over time, leading to suboptimal glycaemic control unless higher dosages or additional agents are initiated⁷. Comparable to reports from other studies, older patients in our study were more likely to achieve optimal glycaemic control than young ones^{31 32}. It is possible that young patients are less likely to be compliant with medication and lifestyle modification as

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3 compared to their older counterparts. However, again we need to research this further before making any
4 concrete statements and instigating pertinent quality improvement programmes.

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6 We also found a high prevalence (80.8%) of hypertension among patients attending our specialised diabetes
7 clinic. A decade ago, the prevalence of hypertension among patients with diabetes in this setting was 61.2%
8 ³³. Our findings may suggest an increasing burden of hypertension as seen globally, but also a reflection of
9 the improvement in the screening and diagnosis of hypertension over the past few years³⁴. We are aware
10 of the comparable high frequencies of hypertension in other African studies ¹³. Consistent with previous
11 studies, patients with hypertension were older, more obese, and had declining GFR and longer diabetes
12 duration^{11 35}. Thiazide diuretics, calcium channel blockers and ACE inhibitors were the three most
13 prescribed antihypertensive agents. This finding is in line with the available evidence recommending
14 thiazide diuretics and calcium channels blockers as the most effective antihypertensives in the black
15 population ^{19 20 35}. Optimal hypertension control was observed in only 54.2% of the patients on
16 antihypertensives, suggesting an urgent need for initiatives to improve the identification and control of
17 hypertension. This low control level is a concern given the increased mortality if hypertension is not
18 controlled⁶. Having said this, the proportion of patients with optimal hypertension control in our population
19 was superior to several studies in Africa, notwithstanding the variation of the definitions of optimal
20 hypertension control across these studies ^{11-14 35-37}. The proportion of type 2 diabetes with optimal
21 hypertension in Africa is often below 35%^{11 12 37}.

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23 However, there is no room for complacency. In the present study, the use of ACE inhibitors was inversely
24 associated with optimal hypertension control. Although ACE inhibitors are indicated for patients with
25 diabetes and proteinuria, they have a clinically significant lesser reduction in both systolic and diastolic
26 blood pressure in the black population. This could partly explain suboptimal hypertension control among
27 predominantly black patients in our study. There appeared to be no influence of gender on hypertension
28 control similar to other studies³⁸, which is encouraging as a recent systematic review found that males in
29 low and middle income countries (LMICs) are more likely to be non-adherent to their medications ^{39 40}.

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31 Less than half of the patients (45%) were on lipid-lowering drugs, mostly statins. This is not surprising as
32 the prescription of lipid-lowering medications in Africa is as low as 3 - 13% in patients with diabetes due
33 to the limited access to these drugs as well as lack of facilities for monitoring lipid profiles while patients
34 are on treatment, regular medication stock outs, and insufficient health professionals ^{13 40 41}. It is, however,
35 a concern as the reduced use of statins will increase mortality rates in patients with diabetes⁸. In some
36 countries, the issue of co-payments limits the prescription of expensive medications like statins⁴². However,
37 this is not an issue in Botswana where medications are provided free of charge to patients. Although the
38 lack of co-payments might have led to a higher prescription of statins in Botswana than in other African
39 countries, we would expect the rate of statin prescriptions to mirror the high rates seen in Western
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3 countries⁴³. Education and adherence to guidelines will possibly improve the prescription and use of statins
4 among the majority of patients with diabetes according to treatment guidelines^{19 21}. Even when statins were
5 used, an undesirably small proportion of our patients achieved guideline-recommended LDL-C target level.
6 Suboptimal LDL-C control rates are also frequent across different settings, even in developed countries^{9 13}
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15⁴⁴. This is disappointing as achieving LDL-C reduction is associated with the highest cardiovascular risk
reduction than hypertension and HbA1c reduction⁹. Inadequate patients' adherence and possibly clinicians'
under-dosage of statin for fear of potential side effects are some of the factors that possibly explain
suboptimal LDL-C control in our patients⁹. Besides, clinicians may be unaware of the current LDL-C as
well as those of HbA1c and hypertension therapeutic goals^{19 21}. Irrespective of the reason, there is an urgent
need to instigate measures to meet guideline-recommended therapeutic goals, and we have started to
address this in our clinic. As reported in other studies, women were less likely than men to achieve optimal
LDL-C control⁴⁴. Although the reason for this gender difference is not apparent, this information is
significant for clinicians to pay attention to the management of women with diabetes in Botswana and other
African countries.

To the best of our knowledge this first study to objectively assess the three critical therapeutic targets in
patients with diabetes in one of the few specialised diabetes clinics in Botswana. However, our findings
should be interpreted considering several limitations. First, the study was limited to one specialised public
diabetes clinic, and the findings may not be generalised to other public and private facilities in the country.
Nevertheless, being the leading specialised diabetes clinics in the country, our findings likely represent the
'best' quality of diabetes care in Botswana. Consequently, highlighted concerns are likely to be higher in
non-specialist healthcare facilities treating patients with type 2 diabetes in Botswana. Second, the study was
cross-sectional in design and therefore unable to establish a temporal relationship between the factors
associated with poor control of glycaemia, LDL-C and hypertension. Third, the results may be subject to
selection bias because of incomplete data in some participants. Another potential risk of selection bias is
the fact that the study enrolled only those patients available at the clinic during the study period. As such,
patients unable to attend the clinic or those whose appointments did not coincide with the study period did
not participate. Despite these limitations, we believe our findings are robust to help improve the care of patients
with type 2 diabetes in Botswana.

In conclusion, there was suboptimal glycaemic, hypertension and LDL-C control among patients with
diabetes in our setting. These findings call for urgent individual and health systems interventions to address
the factors associated with suboptimal control of the cardiovascular risk factors among patients with type 2
diabetes in Botswana. This will be the subject of future initiatives and research in our clinic given the
growing prevalence of patients with type 2 diabetes in Botswana.

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5

6 **Competing Interests statement**

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8 None disclosed
9

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12 collection. Also, we appreciate the other staff at the diabetes clinic for their help and the participants for
13 their time and patience to be part of the study.
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16 **Author contributions**

17 JCM was involved in the conception of the study, data analysis, drafting and revising the manuscript. BO
18 was involved in the design of the study and critical revision of the manuscript. AO, MM participated in the
19 design of the study, data collection and critical revision of the manuscript. JMF participated in data analysis
20 and manuscript review. EB performed data entry and management, and critical revision of the manuscript.
21 OJM, BG, and JGT were involved in the interpretation of the data and manuscript revision. All the authors
22 granted final approval of the version to be published; and agreed to be accountable for all aspects of the
23 work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately
24 investigated and resolved.
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32 **Data sharing statement**

33 The dataset analysed during the current study is available from the corresponding author on reasonable
34 request, through the Princess Marina Hospital and Ministry of Health of Botswana ethical approval.
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Table 1: Clinical and socio-demographic characteristics of patients with type 2 diabetes at a specialised Diabetes clinic in Gaborone (N= 500)

Characteristics	All (N=500)	Males (n= 170)	Females (n= 330)	p-value
Mean age (SD), years	58.9 (12.2)	55.4 (12.6)	60.8(11.6)	<0.001
Age <50 years n (%)	109(21.80)	57 (33.5)	52 (15.8)	<0.001
Age ≥ 50 years n (%)	391 (78.20)	113 (66.5)	278(84.2)	
Diabetes duration, median, IQR,years	6 (2 - 13)	6.5 (2 - 14)	6 (2.5 - 13)	0.927
HbA1c mean (SD), %	8.4 (2.4)	8.6 (2.7)	8.4 (2.4)	0.199
Diabetes treatment				
Diet alone n (%)	11(2.2)	3(1.8)	8 (2.4)	
Insulin alone n (%)	68 (13.6)	38(22.3)	30 (9.1)	0.001
OHA alone n (%)	271(54.2)	82(48.2)	189 (57.3)	
Insulin and OHA	150 (30.0)	47(27.7)	103(31.2)	
BMI, mean (SD) kg/m ²	30.5(6.0)	28.7(5.2)	31.4(6.2)	<0.001
Normal weight n (%)	93 (18.7)	46(27.1)	49 (14.9)	<0.001
Overweight n (%)	155(31.1)	61(36.3)	94 (28.5)	
Obese n (%)	250(50.2)	63 (37.1)	187(56.7)	
Marital status				
Living alone n (%)	266 (53.20)	55 (32.4)	211(63.9)	<0.001
Living with a partner n (%)	234(46.8)	115 (67.6)	119(36.01)	
Education status				
No formal education, n (%)	77(15.4)	26(15.3)	51 (15.5)	<0.001
Primary School, n (%)	229(45.8)	56 (32.9)	173 (52.4)	
Secondary school, n (%)	131(26.2)	55(32.4)	76 (23.0)	
College/University, n (%)	63(12.6)	33 (19.4)	30 (9.1)	
WC, mean (SD), cm	103.3(12.5)	101.7 (11.9)	104.0 (12.7)	0.049
WHR	0.94(0.10)	0.97(0.09)	0.93(0.10)	<0.001
Low WHR n (%)	75 (44.1)	16(4.9)		<0.001
High WHR n (%)	95 (55.9)	314(95.2)		
CKD n (%)	54 (10.8)	24 (14.1)	30 (9.1)	0.086
eGFR, Median, IQR (ml/min/1.73m ²)	112.5(84.3-138.1)	113.1(80.8 -139.5)	112.3(84.4 - 137.9)	0.737
Cholesterol, mean (SD), mmol/L	4.4(1.1)	4.3(1.2)	4.5(1.1)	0.030
LDL-C mean (SD), mmol/L	2.8(1.0)	2.6(1.0)	2.9(1.0)	0.006
Proteinuria n (%)	51(10.20)	27 (15.9)	24(7.27)	0.003
Hypertension n (%)	404 (80.80)	120 (70.59)	284(86.06)	<0.001
Dyslipidaemia n (%)	358 (71.60)	114 (67.06)	244(73.94)	0.106

Legend: DM - Diabetes mellitus; WC- waist circumference; BMI- Body Mass Index; SD- standard deviation; IQR – interquartile range; eGFR- estimated glomerular filtration rate; LDL-C – Low-density lipoprotein; WHR – Waist-hip ratio, HbA1c- Haemoglobin A1c; CKD- Chronic kidney disease; OHA – oral hypoglycaemic agent

Table 2: Factors associated with optimal glycaemic control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)

Characteristic	Unadjusted OR (95% CI)	p-value	Adjusted OR (95%CI)	p-value
Age group				
> 50 years	1			
≤ 50 years	2.16(1.29 - 3.61)	0.003	5.79 (1.08 - 31.14)	0.041
Diabetes duration, years	0.97(0.95 - 0.99)	0.040	0.91 (0.85 - 0.98)	0.011
Use of Insulin alone				
No	1			
Yes	0.80(0.45- 1.43)	0.590		
Use of OHA alone				
No	1			
Yes	2.785 (1.862 - 4.167)	<0.001	0.90 (0.46 - 1.74)	0.745
Use of Insulin plus OHA				
No	1			
Yes	0.185 (0.124 - 0.356)	<0.001	0.34 (0.07 - 1.70)	0.188
BMI	0.99(0.96 - 1.02)	0.427	–	–
Normal weight	1	–	–	–
Overweight	1.10 (0.64 - 1.90)	0.721	–	–
Obese	0.83(0.50 - 1.39)	0.476	–	–
Gender				
Men	1	–	–	–
Women	0.92(0.62 - 1.36)	0.663	0.42 (0.14 - 1.25)	0.120
Education status				
None	1	1	–	–
Primary School	1.58(0.88 - 2.81)	0.124	–	–
Secondary school	1.14 (0.60 - 2.16)	0.687	–	–
College/University	1.00(0.47 - 2.13)	0.996	–	–
Marital status				
Living alone	1	–	–	–
Living with a partner	0.93 (0.64 - 1.36)	0.703	–	–
WHR	0.92 (0.13 - 6.58)	0.937	–	–
Low WHR	1	–	–	–
High WHR	0.92(0.57 - 1.50)	0.745	–	–
Weight, kg	0.99 (0.98 - 1.01)	0.298	–	–
eGFR, (ml/min/1.73m ²)	1.00 (0.99 - 1.00)	0.074	1.00(0.99 - 1.01)	0.766
CKD				
No	1			
Yes	1.053 (0.578 - 1.920)	0.866	–	–
Proteinuria				
No	1		–	–
Yes	1.16(0.63 - 2.14)	0.624	–	–
Optimal hypertension control				
No	1		–	–
Yes	2.53 (1.63 - 3.93)	<0.001	1.61(0.63 - 4.13)	0.322
Optimal LDL-C control				
No	1			
Yes	2.10 (0.90 - 4.88)	0.086	2.20 (0.64 - 7.57)	0.209

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3 **Legend:** Legend: BMI- Body Mass Index; CKD- Chronic kidney disease; DM - Diabetes mellitus; eGFR- estimated glomerular
4 filtration rate; IQR – interquartile range; LDL-C – Low-density lipoprotein; OHA – oral hypoglycaemic agent; SD- standard
5 deviation; WHR – Waist-hip ratio.
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Table 3: Factors associated with optimal hypertension control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)

Characteristic	Unadjusted OR (95% CI)	p-value	Adjusted OR (95% CI)	p-value
Age group				
< 50 years	1	–	–	–
≥50 years	0.89(0.48 - 1.62)	0.691	–	–
Diabetes duration, years	0.97(0.94 - 0.99)	0.023	0.98 (0.92 - 1.03)	0.352
Use of Insulin alone				
No	1	–	–	–
Yes	0.45 (0.24 - 0.82)	0.009	1.15 (0.30 - 4.44)	0.837
Use of OHA alone				
No	1	–	–	–
Yes	1.94 (1.30 - 2.91)	0.001	1.37 (0.51 - 3.66)	0.531
Use of Insulin plus OHA				
No	1	–	–	–
Yes	0.59 (0.40 - 0.95)	0.028*	–	–
BMI, kg/m ²	0.98 (0.94 - 1.01)	0.149	0.93(0.86 - 1.01)	
Normal weight	1	–	–	–
Overweight	0.79 (0.42 - 1.47)	0.449	–	–
Obese	0.77(0.44 -1.38)	0.384	–	–
Gender				
Men	1	–	–	–
Women	1.43(0.92 - 2.22)	0.117	0.97 (0.36 - 2.61)	0.949
Marital				
Living alone	1	–	–	–
Living with a partner	0.68 (0.61 - 1.37)	0.67	–	–
Education status				
No formal education	1	–	–	–
Primary School	1.03(0.58 - 1.83)	0.09	–	–
Secondary school	1.08(0.56 - 2.09)	0.22	–	–
College/University	0.89(0.41 - 1.92)	0.761	–	–
WHR	1.29 (0.14 - 12.00)	0.821	–	–
Low WHR	1	–	–	–
High WHR	1.18 (0.65 - 2.12)	0.589	–	–
eGFR, (ml/min/1.73m ²)	1.00 (1.00 - 1.01)	0.139	1.00(0.99 -1.02)	0.412
Total serum Cholesterol	0.964 (0.80 - 1.16)	0.693	–	–
Proteinuria				
No	1	–	–	–
Yes	0.48(0.26 - 0.92)	0.027	0.36(0.07 - 1.80)	0.213
Use of CCB				
No	1	–	–	–
Yes	0.73(0.72 - 1.61)	0.729	–	–
Use of Thiazides				
No	1	–	–	–
Yes	1.58(1.06 - 2.37)	0.026	1.44(0.62 - 3.39)	0.399
Use of ACE inhibitors				
No	1	–	–	–
Yes	0.64 (0.43 - 0.95)	0.028	0.35(0.14 - 0.85)	0.020
Use of ARB				
No	1	–	–	–
Yes	0.95(0.57 - 1.58)	0.834	–	–
Alpha blocker				

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3	No	1	–	–	–
4	Yes	0.24(0.09 - 0.68)	0.007	0.76 (0.14 - 4.20)	0.749
5	Beta-blocker				
6	No	1	–	–	–
7	Yes	0.70 (0.43 - 1.14)	0.149	0.51(0.19 - 1.37)	0.184
8	Optimal Glycaemic control				
9	No	1	–	–	–
10	Yes	2.53 (1.63- 3.93)	<0.001	1.92 (0.71 - 5.23)	0.201
11	Optimal LDL-C control				
12	No	1	–	–	–
13	Yes	0.55 (0.23 - 1.36)	0.199	0.75 (0.25 - 2.32)	0.623

Legend: * omitted because of collinearity. ACE INHIBITORS – angiotensin-converting-enzyme inhibitor; ARB – Angiotensin II receptor blocker; BMI- Body Mass Index; CCB- Calcium Channel Blockers; DM - Diabetes mellitus; eGFR- estimated glomerular filtration rate; HbA1c- Haemoglobin A1c; IQR – interquartile range; LDL-C – Low-density lipoprotein; OHA – oral hypoglycaemic agent; SD- standard deviation; WHR – Waist-hip ratio.

Table 4: Factors associated with optimal LDL-C control among patients with type 2 diabetes at a specialised diabetes clinic in Gaborone (N= 500)

Characteristic	Unadjusted OR (95% CI)	p-value	Adjusted OR (95% CI)	p-value
Age group				
<50 years	1		–	–
≥ 50 years	0.56 (0.21 - 1.50)	0.249	–	–
Diabetes duration, years	1.02(0.97 - 1.06)	0.525		–
Use of Insulin alone				
No	1	–	–	–
Yes	1.58 (0.56 - 4.46)	0.389	–	–
Use of OHA alone				
No	1	–	–	–
Yes	0.95 (0.43 - 2.12)	0.900	–	–
Use of Insulin plus OHA				
No	1	–	–	–
Yes	0.83 (0.35 - 1.97)	0.664	–	–
BMI	1.00 (0.93 -1.07)	0.951		–
Normal weight	1	–	–	–
Overweight	1.43 (0.34 - 5.94)	0.624	–	–
Obese	1.27(0.33 - 4.89)	0.730	–	–
Gender				
Men	1	–	–	–
Women	0.2 (0.09 - 0.47)	<0.001	0.24(0.09- 0.59)	0.002
Education status				
No formal education	1	1	–	–
Primary School	0.79 (0.29 - 2.14)	0.647	–	–
Secondary school	0.28 (0.07 - 1.20)	0.087	–	–
College/University	0.79(0.20 - 3.11)	0.731	–	–
Marital status				
Living alone	1	–	–	–
Living with a partner	0.99 (0.44 - 2.20)	0.973	–	–
HbA1c	1.00 (0.85 - 0.20)	0.917		–
WHR	7.59(0.19 - 303.60)	0.281	–	–
Low WHR	1	–	–	–
High WHR	0.31(0.11 – 0.89)	0.030	0.64(0.20 - 2.10)	0.463
Proteinuria				
No	1	–	–	–
Yes	0.69 (0.14 - 3.27)	0.64	–	–
eGFR, (ml/min/1.73m ²)	1.00(0.99 - 1.01)	0.878	–	–
CKD				
No	1	–	–	–
Yes	1.88(0.73 - 4.83)	0.193	1.67 (0.61 - 4.58)	0.321
Hypertension				
No	1	–	–	–
Yes	1.13 (0.30 - 4.23)	0.86	–	–

Legend: BMI- Body Mass Index; CKD- Chronic kidney disease; DM - Diabetes mellitus; eGFR- estimated glomerular filtration rate; IQR – interquartile range; LDL-C – Low-density lipoprotein; OHA – oral hypoglycaemic agent; SD- standard deviation; WHR–Waist-hip ratio

STROBE 2007 (v4) Statement—Checklist of items that should be included in reports of *cross-sectional studies*

Section/Topic	Item #	Recommendation	Reported on page #
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract	1
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	1
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	2
Objectives	3	State specific objectives, including any prespecified hypotheses	2
Methods			
Study design	4	Present key elements of study design early in the paper	2
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	2
Participants	6	(a) Give the eligibility criteria, and the sources and methods of selection of participants	2
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	2
Data sources/ measurement	8*	For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group	2
Bias	9	Describe any efforts to address potential sources of bias	2
Study size	10	Explain how the study size was arrived at	2
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why	3
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	3
		(b) Describe any methods used to examine subgroups and interactions	3
		(c) Explain how missing data were addressed	
		(d) If applicable, describe analytical methods taking account of sampling strategy	
		(e) Describe any sensitivity analyses	
Results			3

Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed (b) Give reasons for non-participation at each stage (c) Consider use of a flow diagram	3
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders (b) Indicate number of participants with missing data for each variable of interest	4
Outcome data	15*	Report numbers of outcome events or summary measures	4,5
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and why they were included (b) Report category boundaries when continuous variables were categorized (c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	4,5
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	
Discussion			
Key results	18	Summarise key results with reference to study objectives	5
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	1
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	8
Generalisability	21	Discuss the generalisability (external validity) of the study results	8
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	8

*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at <http://www.plosmedicine.org/>, Annals of Internal Medicine at <http://www.annals.org/>, and Epidemiology at <http://www.epidem.com/>). Information on the STROBE Initiative is available at www.strobe-statement.org.