An evaluation of adherence to anti-diabetic medications among type 2 diabetic patients in a Sudanese outpatient clinic

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Abstract

Introduction: adherence to anti-diabetic medication is a known cornerstone in the management of type 2 diabetic patients. We sought to assess the factors associated with adherence to medications among type 2 diabetic patients being followed up in a Sudanese outpatient clinic. Methods: this cross-sectional study conducted among 102 patients with type 2 diabetes attending an outpatient clinic in Omdurman, Sudan during the period from June to December 2017. Participants were interviewed using a structured questionnaire to collect demographic data, number, and type of medications, polypharmacy, medications side effects, financial problems and education regarding drug used. The study of participants’ adherence to anti-diabetic medications was assessed using a validated questionnaire asking the patients about the percent and self-rating of adherence (Excellent, very good, good, fair and poor). The Statistical Package for Social Sciences (SPSS) was used to compare the adherent patients and their counterparts. A P-value < 0.05 was considered significant. Results: the study results summarized the following: participants (70.6% women), their mean age was (59.62±9.91) years and nearly 60.8% were housewives, their glycated hemoglobin (mean± SD) was about 10.16±3.14, 37.3%, it implies that the patients were non-adherent to medications. In addition, other groups of patients with medication but non-adherence were younger ones (55.94±9.94 vs. 61.81±9.36, P=0.04) and had shown inadequate glycemic control (11.33±3.05vs. 9.47±3.04, P=0.04), however, this group of patients has reported more drug-related side effects (57.8% vs. 28.1%) because they were taking more drugs compared to their counterparts( F=4.115, P=0.047). The present study found no statistically significant differences in the following factors such as sex, occupation, education level, financial problems and insulin use. Conclusion: in conclusion, the study revealed that adherence to anti-diabetic medications was sub-optimal among Sudanese type 2 diabetic patients and was associated with higher glycated hemoglobin seen among younger age groups. Besides the above, overdosing of medications and their side effects were evident.
**Introduction**

Diabetes mellitus is the most common endocrine disease worldwide; it is a serious major health problem. Currently, 285 million were affected, and the projection for the year 2030 is 438 million [1]. According to the World Health Organization estimation, half-million Sudanese people were affected by diabetes mellitus in the year 2000 and this trend is expected to reach one million in the year 2030 [2]. The management of type 2 diabetes includes the adoption of a healthy lifestyle, a friendly diet and pharmacotherapy. Adherence to management plan is an essential component of diabetes holistic care. Adherence is defined as the extent to which a person’s behavior regarding diets, healthy lifestyles, or taking medications or behavior changes coincide with medical and health advice [3]. The lack of adherence to drugs according to doctor prescription has consistently shown to be associated with poor glycemic control and microvascular complications [4, 5] among diabetic patients. Patients’ non-adherence to therapeutic strategies is a common serious health concern worldwide. Medication non-adherence could be related to healthcare system factors, patients’ factors or due to medications. Non-adherent to the prescribed medications is a principal factor in poor glycemic control and microvascular complications including neuropathy, nephropathy and retinopathy with deleterious consequences on the patients and the community as a whole [6]. Polypharmacy, which is the use of simultaneous multiple drugs by the same patient for one or more disease has become a huge problem even in the developed countries [7]. The term polypharmacy, defined as taking five medications or more was shown to increase non-adherence to medications, which lead to a propensity to falls and increase meditational side effects [8, 9]. Previous studies have evaluated the adherence to anti-diabetic medication and polypharmacy in Sudan; however, due to the introduction of new anti-diabetic medications with varying doses and schedules may further affect adherence to medicines behavior, change in lifestyles and culture, which are essential factors to evaluate this critical health problem regularly. Based on the above fact, this survey was designed to assess the adherence to anti-diabetic medications among patients with type 2 diabetes in Sudan.

**Methods**

This cross-sectional study conducted at a randomly chosen diabetes clinic from seven clinics in Omdurman, Sudan during the period from June to September 2017. One hundred and two consecutive patients with the diagnosis of type 2 diabetes (according to American Diabetes Association Guidelines) [10] were invited to sign a written informed consent then interviewed using a structured questionnaire to collect age, sex, occupation, their educational level, time since the diagnosis of diabetes, number and type of medications taken by the patients over the period, history of microvascular and macrovascular complications of diabetes if any, financial issues with drugs, and medications side effects, etc. were also included. With the above background, the adherence to anti-diabetic medicines was assessed using the following questions: during the past month what percent of the time did you take all your diabetes medications as your doctor prescribed? And on average, how would you rate your ability to take all your diabetes medications as your doctor prescribed (excellent, very good, good, fair, poor, and very poor)? The short medication adherence questionnaire had been previously validated for use among patients with diabetes [11]. Additionally, the days covered with medication in the past week were reported. On the other hand, non-adherence to anti-diabetic medications was defined as taking less than 80% of the prescribed treatment [12]. A blood sample was taken for the glycated hemoglobin (HbA1c) to assess the degree of glycemic control using the reagent from utilizing a glycol hemoglobin reagent set from HB1C Siemens Healthcare Diagnostics Newark, DE 19714, USA. Patients on five drugs or more were considered as polypharmacy [8]. This study obtained ethical committee approval from the Faculty of Medicine, University of Tabuk to conduct this cohort research. The Statistical Package for Social Sciences (SPSS, Version 20, Chicago) was used for data analysis, the Chi-square test, t-test, and One Way-ANOVA were also used to compare patients who were adherent to medications with their counterparts, a P-value of < 0.05 was considered significant.

**Results**

Out of 102 patients with type 2 diabetes (70.6% women), their ages ranged from 28-82 years with a mean of (59.62±9.91 years). Nearly 25.5% were not educated but 29.4% of them had primary education, while 21.6% had a high school education, however, the majority
of the patients were housewives and 17.6% were laborer, 15.7% were employed and 5.9% were teachers. Table 1 shows the good glycemic control was evident only in 23.5% of patients. The participants mean HbA1c% was (10.16±3.14), the meantime since the diagnosis of diabetes was (11.90±8.72), the mean number of medications was (4.96±1.35) and the neck circumference was (34.27±9.26cm) (Table 2).

In the present study, 88.2% were on metformin, 78.4% were taking sulphonylureas, 9.8% were prescribed insulin and 2% were on pioglitazone. It is interesting to note that 92.2% of patients were taking regular vitamins, 35.3% were on regular proton pump inhibitors, 11.8% were on regular non-steroidal anti-inflammatory drugs, while 62.7% were taking five medications or more (polypharmacy). Table 3 illustrates the prescription among patients with type 2 diabetes. The current data showed that 62.7% of patients were adherent to medications, 43.1% rated their adherence as excellent, 31.4% were very good, 21.6% as good, while 3.9% rated themselves as fair. The days covered with medications were seven in 72.5% of patients, six in 15.7%, five in 2%, 5.9% covered four days, while 2% covered two days. Regarding the negative attitudes of patients regarding medications, sixty-two percent were not adequately educated regarding medication, financial issues were reported by 39.2%, while 37.3% reported medications side effects (Table 4).

In the present study, results revealed that patients who were adherent to medications were a mature adult than the non-adherent younger patients (61.81±9.36 vs. 55.94±9.94), P-value=0.040 and these patients had better glycemic control (HbA1c,9.47±3.04 vs. 11.33±3.05) with a significant statistical difference, P=0.040 compared to their counterparts. Besides the above, no significant statistical difference was found regarding the duration of the diagnosis of diabetes (11.93±9.33 vs. 11.84±7.84), P-value=0.970 Table 5.

The present study showed that adherence was lower among patients who experienced medication side effects (28.1% vs. 57.8%), P-value=0.035, no differences were evident regarding insulin use (6.2% vs. 15.7%), P-value=0.286, sex (31.2% vs. 26.3% of males and 68.8% vs. 73.7% of women), P-value=0.761 and financial issues (37.5% vs. 36.8%), P-value=0.963. Table 6 illustrated the comparison between adherence and non-adherence. In the current study, medication adherence was lower among patients taking higher medications (P-value=0.047, F=4.115, no differences in medications adherence was evident regarding occupation and level of education (P-values=0.483 and0.057, F=0.499 and 3.801 respectively) (Table 7).

**Discussion**

The present study showed that 62.7% of patients with type 2 diabetes were adherent to the prescribed medications with 74% reported their adherence as excellent and very good, and 88.2% had six or more days /week covered with drugs. The present findings were similar to a study conducted in India [6] and reported adherence in 54.6%. Higher rates (74.9%) were reported in Ethiopia [13] and Nepal [14] (97.3%), differences in sample size and the measures used could partially explain the discrepancies between the studies. In the present study, females’ dominance was obvious, the majority were housewives, and more than half received only primary education, a study published in Turkey [15] concluded similar findings.

In the current study, nearly two-thirds of patients were not educated about the medications, 37.3% were concerned about drugs side effects, and 39.2% reported financial issues regarding the medicines to use, previous researchers [6] reported similar results regarding barriers to medications adherence. A recent large study [16] conducted in five European countries observed the agreement regarding metformin as the first-line treatment among patients with type 2 diabetes and differences in the second line medications attributed to different national guidelines. The current data showed the most frequently prescribed medications were metformin followed by sulphonylureas in accordance with Acharya et al. [17] who conducted a survey in India and found metformin and glimepiride were the most common medications. The absence of DPP-4 inhibitors and Glucagon-Like Peptide agonists was due to either unavailability or cost.

There is a trend towards increasing combination of statins, aspirin and hypertensive medications among patients with type 2 diabetes [18]. In the present study, more than half of the patients were using these drugs. Previous literature reported that polypharmacy is associated with medication non-adherence, medications side effects, and dementia, in the present study nearly two-thirds of the patients were on five or more drugs in agreement with Patel et al. [19] who reported polypharmacy and hyper pharmacy in 59% and 31% respectively. It is interesting to note that
92.2% of the study sample were on regular vitamin supplementation despite the possible harmful effects of regular multivitamins supplementations as fat-soluble vitamins could lead to toxicities due to accumulation in the body. The American Diabetes Association [20] recommended to test serum vitamin B12 and replace if deficient. The present data showed that, 35.3% were on regular proton pump inhibitors in spite of the strong recommendation against their use for more than eight weeks, and their association with Clostridium difficile infections and dementia [21] and 11.9% were taking non-steroidal anti-inflammatory drugs on regular base in spite of the danger of lethal side effects. The pharmacists and the prescribing physicians may need proper communication and counseling of the patients and use of the emerging computer system to optimize the prescription. In the current study, medication non-adherence was higher among patients with poor glycemic control in line with Jemal et al. [13] who found an association between non-adherence and poor glycemic control. The relationship between medications side effects and non-adherence to medications is obvious. In the current study, non-adherent patients reported a higher rate of drugs side effects.

The present results found lower medication adherence among patients with polypharmacy in accordance with a previous study [22] which showed a relationship with non-adherence and the complexity of prescription, another study [23] showed adherence to anti-diabetic medication was better among those taking ≥ four drugs than those on lesser drugs and attributed this to the age and level of education in their sample in contradiction to the present findings. In the present study, no differences in adherence were observed regarding sex, occupation, level of education, duration of diabetes, financial problems and insulin use in agreement with Patel et al. [20] who concluded similar results. Similar results were observed by Serap and Bayram [15] regarding the association of sex, financial problems, and duration of diabetes with medication adherence, however, the relationship of medications adherence to the level of education was in contradiction to the present findings. In the current survey, non-adherence was common among the young age group and those with higher glycated hemoglobin and in agreement with a population-based study [24] conducted in France, but Tabannick and Fidel [25] recommended that independent variables should not be strongly related to each other. The American Diabetes Association recent recommendations of incorporating agents proven to reduce cardiovascular mortality and major cardiovascular events and the equipoise gave to much second-line therapy may encourage the developing countries to consider the use of new classes of medications in spite of the high cost. Thus, medication class shifting might be a useful method to improve medication non-adherence.

Medication therapy management and better communication of the patients and healthcare providers are highly relevant for therapy optimization and reducing non-adherence and approaching the glycated hemoglobin targets. Further larger multi-center studies to assess adherence to an individual class of medications are highly recommended. The study limitations were the small sample of the research, the reliance on a self-reported questionnaire which could lead to overestimation of the adherence and the fact that the study was conducted at a single diabetes clinic.

**Conclusion**

In conclusion, the adherence to anti-diabetic medications was sub-optimal and common among young patients with poor glycemic control, those on five or more drugs and those who exposed to medications side effects, no difference between adherent and non-adherent patients regarding sex, occupation, level of education, the duration since the diagnosis of diabetes, insulin use, and financial problems.

**What is known about this topic**

- Polypharmacy is common among patients with type 2 diabetes;
- Inadequate adherence to anti-diabetic medications compromise safety and treatment effectiveness.

**What this study adds**

- Non-adherence to anti-diabetic medications was common among young type-2 diabetic patients, these patients need more effort regarding the importance of medication adherence;
- Patients who observed medications side effects were non-adherent. Health education regarding antidiabetic side effects may be briefed or needed before the prescription was given. Also, sharing the decision with the patients is vital to enhance medication adherence. Involvement of the pharmacists in the education of patients with type 2 diabetes patients will bring more benefits and of great help to achieve the greater outcome.
Competing interests

The author declares no competing interests.

Authors’ contributions

HOM conceived and designed the study, conducted research, data acquisition, interpretation and manuscript drafting.

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### Table 1: Basic characters and adherence to anti-diabetic medications among Sudanese patients with type 2 diabetes mellitus

| Character          | N (%)       |
|--------------------|-------------|
| **Sex**            |             |
| Males              | 30 (29.4%)  |
| Females            | 72 (70.6%)  |
| **Education**      |             |
| Illiterate         | 26 (25.5%)  |
| Primary School     | 30 (29.4%)  |
| Intermediate School| 10 (9.8%)   |
| High secondary School | 22 (21.6%) |
| University         | 14 (13.8%)  |
| **Occupation**     |             |
| Housewives         | 62 (60.8%)  |
| Laborer            | 18 (17.6%)  |
| Employee           | 16 (15.7%)  |
| Teachers           | 6 (5.9%)    |
| Good glycemic control | 24 (23.5%) |

### Table 2: Patient's characters mean ±SD

| Character                        | Mean ±SD      |
|----------------------------------|---------------|
| Age years                        | 59.62±9.91    |
| HbA1c%                           | 10.16±3.14    |
| Meantime since diabetes diagnosis| 11.90±8.72    |
| Number of medications            | 4.96±1.35     |
| Neck circumference (cm)          | 34.27±9.26    |

### Table 3: Prescription pattern among patients with type 2 diabetes

| Medication                      | N (%)       |
|---------------------------------|-------------|
| Metformin                       | 90 (88.2%)  |
| Sulphonylureas                  | 80 (78.4%)  |
| Pioglitazone                    | 2 (2%)      |
| Insulin                         | 10 (9.8%)   |
| Antihypertensive                | 58 (56.9%)  |
| Statins                         | 62 (60.8%)  |
| Aspirin (low dose)              | 56 (54.9%)  |
| Proton pump inhibitors          | 36 (35.3%)  |
| Vitamins                        | 94 (92.2%)  |
| No steroidal anti-inflammatory drugs | 12 (11.8%) |
| Polypharmacy                    | 64 (62.7%)  |
Table 4: medication adherence to anti-diabetic medications among patients with type 2 diabetes

| Character | N (%) |
|-----------|-------|
| Medication adherence % | | |
| 100% | 36 (35.3%) |
| 90% | 26 (25.5%) |
| 85% | 4 (3.9%) |
| 80% | 12 (11.8%) |
| 75% | 8 (7.8%) |
| 70% | 8 (7.8%) |
| 60% | 2 (2%) |
| 50% | 2 (2%) |
| 45% | 4 (3.9%) |

Adherence self-rating

| Character | |
|----------|---------|
| Excellent | 44 (43.1%) |
| Very good | 32 (31.4%) |
| Good | 22 (21.6%) |
| Fair | 4 (3.9%) |

Number of days covered with drugs

| Character | |
|-----------|---------|
| Seven days | 74 (72.5%) |
| Six days | 16 (15.7%) |
| Five days | 2 (2%) |
| Four days | 6 (5.9%) |
| Two days | 2 (2%) |
| One day | 2 (2%) |
| Lack of knowledge about medications | 64 (62.7%) |
| Financial issues | 40 (39.2%) |
| Medications side effects | 38 (37.3%) |

Table 5: the relationship of adherence to anti-diabetic medication, mean age, duration since the diagnosis of diabetes, and HbA1c

| Character | Adherence (n=32) | Non-adherence (n=19) | P-value |
|-----------|------------------|----------------------|---------|
| Age | 61.81±9.36 | 55.94±9.94 | 0.040 |
| Duration of diabetes | 11.93±9.33 | 11.84±7.84 | 0.970 |
| HbA1c | 9.47±3.04 | 11.33±3.05 | 0.040 |

*T-test

Table 6: the relationship of adherence to anti-diabetic medication, sex, insulin use and barriers to adherence

| Character | Adherence (n=32) | Non-adherence (n=19) | P-value |
|-----------|------------------|----------------------|---------|
| Sex | | | |
| Males | 10 (31.2%) | 5 (26.3%) | 0.761 |
| Females | 22 (68.8%) | 14 (73.7%) | |
| Insulin use | 2 (6.2%) | 3 (15.7%) | 0.286 |
| Financial issues | 12 (37.5%) | 7 (36.8%) | 0.963 |
| Medications side effects | 9 (28.1%) | 11 (57.8%) | 0.035 |

*Chi-square-test
Table 7: relationship of adherence to anti-diabetic medications to the occupation, level of education, and the number of prescribed medications

| Character              | F    | P-value |
|------------------------|------|---------|
| Level of education     | 3.801| 0.057   |
| Occupation             | 0.499| 0.483   |
| Number of medications  | 4.115| 0.047   |

*One Way-ANOVA