

## **Glycemic control among patients with Type 2 diabetes mellitus: in relation to medication adherence and life style**

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### **Abstract**

Type 2 diabetes mellitus (DM) is a chronic metabolic disorder in which prevalence has been increasing steadily all over the world. Lifestyle management is a fundamental aspect of diabetes care and it includes nutrition, physical activity, smoking cessation, and psychosocial care. Good glycaemic control among type 2 diabetes mellitus patients involves interplay of self-management measures including physical activity and diet, in addition to medication adherence. One of the major contributing factors is poor medication adherence in type 2 diabetes mellitus which is well documented to be very common and is associated with inadequate glycemic control; increased morbidity and mortality; and increased costs of outpatient care, emergency room visits, hospitalization, and managing complications of diabetes.

Observational cross sectional study, conducted at different major health care facilities in northern Jordan, recruited more than 300 adult patients who had been on the same antidiabetic treatment regimen for a minimum of 3 months. Patients' data, physical measurements, medications and disease history, dietary carbohydrates and lipids intake data were obtained using pre-validated questionnaires. Fasting blood samples were collected and assessed for HbA1c.

A total of 300 type2 diabetes mellitus patients were enrolled in this study, 63.7% of them were females and 36.3 % were males. The average age of patients was 60.1 years. About 52 (17.3%) of them were smokers, most of them were males (30%). Medication adherence showed that more than two-thirds (69.4%) of the study samples were highly adherent to their medication. Medication adherence was associated significantly with HbA1c level. Only 41.4% of the patients had diet counseling and 58.6% followed a diet plan, both of them were none significantly associated with HbA1c level.

Glycemic control in type2 diabetes mellitus is not just a matter of proper medications use and dosages, but other factors may contribute to poor glycemic control.

**Key words:** Glycemic control, HbA1c, medication adherence, type 2 diabetes mellitus.

### **Introduction:**

Type 2 diabetes mellitus (DM) is a chronic metabolic disorder in which prevalence has been increasing steadily all over the world; with nearly 200 million adults affected along with a 5-fold increased risk of death in developed countries<sup>(14,29)</sup>. Type 2 diabetes mellitus was firstly described as a component of metabolic syndrome in 1988<sup>(30)</sup>. It is characterized by hyperglycemia, insulin resistance, and relative insulin deficiency<sup>(21)</sup>.

Complex biological elements that influence the development and course of the disease in high-risk populations are important but often insufficient to implement effective prevention and treatment plans. Multiple factors must be addressed in routine diabetes clinical care<sup>(8)</sup>.

The current DM management therapeutic regimen includes medications, diet, physical activity, adherence; and recently the standards of medical care in diabetes has been updated by American diabetic association (ADA) to address psychosocial issues in all aspects of care including self-management, mental health, communication, complications, co morbidities, and life-stage considerations<sup>(3)</sup>.

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Lifestyle management is a fundamental aspect of diabetes care which includes nutrition, physical activity, smoking cessation, and psychosocial care<sup>(39)</sup>. Lifestyle, ethnicity, and age, are other factors partly contributing to high prevalence of DM. A sedentary lifestyle, in addition to consumption of high-fat and high-carbohydrate foods, lead to increased body weight and obesity . Approximately, 90% to 95% of all diagnosed cases of DM have type 2 DM, most of them are overweight<sup>(4)</sup>. Advancing age is another factor contributing to the increased incidence and prevalence of type 2 DM<sup>(5)</sup> and mostly occurs in developing countries where the majority of patients are aged between 45 and 64 years<sup>(41)</sup>.

Physical activity plays an important role in preventing type 2 diabetes. A healthful diet and a moderate exercise program resulting in a 5 to 7 percent weight loss can delay and possibly prevent Type 2 diabetes.<sup>(11)</sup> Aerobic exercise improves glycemic control, blood pressure and triglycerides level<sup>(25)</sup>, which lead to reduced cardiovascular mortality and total mortality in patients with type 2 DM<sup>(37)</sup> and exercise for at least 4 hours per week reduces CVD risk by 40% in women with type 2 DM<sup>(12)</sup>.

Cigarette smoking is an independent modifiable risk factor for development of diabetes<sup>(23,44)</sup>, and smokers are more likely to develop diabetes compared to non-smokers<sup>(42)</sup>.

The medication adherence which is defined as the extent to which a person's behavior agrees with the agreed medication regimen from a health care provider<sup>(22)</sup>. At least 45% of patients with type 2 DM fail to achieve adequate glycemic control (HbA1c <7%). One of the major contributing factors is poor medication adherence in type 2 DM which is well documented to be very common and is associated with inadequate glycemic control; increased morbidity and mortality, increased costs of outpatient care, emergency room visits, hospitalization, and managing complications of diabetes<sup>(33)</sup>.

### **Aims of study:**

To examine the role of lifestyle risk factors (diet, exercise and smoking ) association with poor glycemic control in type 2 Diabetes mellitus and assess patients medication adherence.

### **Patients and materials:**

This is a cross-sectional observational study conducted during the period of April 2017 to July 2017. The is three hundred fasting patients visiting the outpatient clinics in northern Jordan; at Princess Bassama Ambulatory Care Clinics for follow-up on diabetes management were enrolled. The laboratory work was performed at JUST Medical Health Center lab, during the same period of samples collection.

The Inclusion criteria were:

- Adult patients (Age  $\geq$  18 years).
- Patient with DM type 2 treated with hypoglycemic medications for a minimum of 6 months on the same DM regimen, and participating patients willing to complete the study surveys.

Exclusion criteria were:

- recent major trauma.
- recent surgery, and hospitalization within 12 week, or acute infection requiring antibiotic use.
- Any significant change in the usual diet within the past month.
- Pregnancy, breast feeding or postpartum within 6 months.
- Acute coronary syndrome within 12 weeks.
- Patients unable to provide an informed consent.
- Patients who were not fasting.

The study procedures were explained to eligible patients and an informed consent was obtained prior to enrollment and interview.

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A validated questionnaire was utilized to assess the adherence to hypoglycemic treatment regimen.

Medication adherence level categorized into: High, medium and low, according to Morisky medication adherence scale<sup>(34,26)</sup>. Dietary carbohydrates, lipids intake data, and life-style activity were recorded. Physical activity was classified into: light exercise includes activities that not causing sweat breaking or produce shortness of breath less than 3.0 METS (less than 3.5 calories per minute) (One MET is defined as the energy cost of sitting quietly and is equivalent to a caloric consumption of 1kcal/kg/hour). Moderate exercise is an exercise which causing break out in a light to moderate sweat or makes it difficult to carry on a long conversation. Less 3.0-6.0 METS (3.5 – 7 calories per minute). Vigorous exercise causing sweat profusely and makes it difficult to carry on a conversation of any length. (Greater than 6.0 METS (more than 7 calories per minute)<sup>(9)</sup>).

Smoking survey was estimated using a pre-validated survey slightly modified to dietary habits in the Jordanian population. Blood samples were obtained after an overnight fasting period (minimum of 10 hours) and 5 ml of patient's whole blood was obtained in EDTA test tube for the HbA1c measurement.

Data were statistically analyzed using SPSS version 22 software program. General linear model (GLM) was utilized to assess the effect of the predictors on attaining target blood glucose level. All statistical tests were considered significant when P-value of <0.05.

### Results:

A total of 300 type 2 DM patients were enrolled in this study, 63.7% of them were females and 36.3 % were males. The average age of patients was 60.1 years. About 52 (17.3%) of them were smokers, most of them were males 33(30%).

**Table 1:** Association of age and gender with HbA1C goal attainment:

➤ Age (years)	Elderly patients	Non -elderly	P value
HbA1c target	8.23 ± 1.89	9.7± 2.46	0.03*
❖ Gender	Male	Female	P value
HbA1c (%)	25%	32%	0.028*

➤ Significant association between the subjects' age and attaining HbA1C target. General linear model (GLM): \* Significant P-value of <0.05.  
 ❖ Significant association for one gender with lower HbA1C control. General linear model (GLM):  
 \* Significant P-value of <0.05.

**Table 2:** Association of life style (Diet counseling, Physical activity, Smoking)

Factors	All patients (n=300)	Male ( n=109)	Female (n=191)	P value
<b>Diet counseling</b>	126 (41.4%)	45 (40.2%)	81 (42.2%)	0.067
<b>Meal plan</b>	178 (58.6%)	66 (58.9%)	112 (58.3)	0.132
<b>Physical activity</b>	123 (41.1%)	39 (34.8%)	85 (44.8%)	0.737
Light	161 (53.0%)	65 (58.0%)	96 (50.0%)	
Moderate	18 (5.9%)	8 (7.1%)	10 (5.2%)	
<b>Smoking</b>	52 (17.3%)	33 (30.3%)* (63.5%)**	19 (10%)* (36.5)**	<b>0.004*</b>

Data represented as number with percentage, statistically significant - p value <0.05  
 \*Calculated from the total male (109) and female (191)  
 \*\* Calculated from the total of male and female smoking (52)

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Medication adherence showed that more than two-thirds (69.4%) of the study sample were highly adherent to their medication, medication adherence was significantly associated with HBA1C goal attainment (P = 0.035).

**Table 3:** Frequency of Medications adherence level and HbA1c attainment.

Adherence level	N=300	Controlled HbA1c (N= 89)	Non controlled HbA1c (N=211)
High adherence level	209 (69.7%)	64(72%)	145 (68.5%)
Medium adherence level	70 (23.4%)	18 (20%)	52 (24.9%)
Low adherence level	21 (6.9%)	7 (8%)	14 (6.6%)

**Discussion:**

The relationship between various patients’ related factors, such as age and gender and blood glucose level, revealed a statistically significant association between the subjects’ age and attaining HBA1c target. The mean HBA1c in elderly patients was  $8.23 \pm 1.89$ , while it was  $9.7 \pm 2.46$  (P = 0.03) for non-elderly patient. This finding agrees with Sanal et al(2011) who concluded that glycemic control is better at patients who are 60 years or older. Kayar et al (2017) and Al-Mukhtar et al (2012) also stated that the disease onset younger were related to poor glycemic control with higher HbA1c%, , who suppose that the poor control of blood sugar in the younger population, compared to the elderly, can be linked to the fact that young people do not pay attention to their treatments as elders.

In Our study, gender also showed a significant association with lower HBA1c control in men (25%), compared to women (32%) (P =0.028). This is inconsistent with other findings <sup>(16,19)</sup>. Sasisekhar et al (2013) found that males had better glycemic control when compared to females, while Waari et al (2018) found that sex not to be significantly associated with adherence and glycemic control.

Examination patient’s diet (if the patients received diet counseling and/or following a diet plan) showed that only 41.4% of participants had diet counseling (40% men vs. 42% women), and 58.6% of them were following a diet plan, without differences in percentage between men and women. No significant association was found between those having a diet plan and glycemic control. Otherwise; When we examined the relation between the ability of participants to follow their diet plan and glycemic control, a significant association was found with attainment of HBA1c goal (P = 0.003) which is parallel with Kayar et al (2017).

These outcomes cannot be interpreted in isolation of other factors, as most participants were obese or overweight, Yet, they highlight an important factor in poor glycemic control which is an unhealthy diet due to many reasons lack of diet awareness (more than 50%), and this is attributed to lack of medical counseling; unhealthy nutritional habits in Jordan, as many depend mainly on a carbohydrate diet and largely due to low financial income for the majority of the study population. These are the main reasons that impede patients from following a healthy diet. A recent study showed that T2DM patients’ education to enhance national habits and reduce caloric intake in association with increased physical activity for six months was associated with significant reduction in mean HbA1c from  $8.5 \pm 2.7\%$  to  $6.0 \pm 1.8\%$  <sup>(38)</sup>.

Patient life-style, diet, exercise, and patient counseling are the primary non-pharmacotherapy an intervention is recommended for treatment of type 2 DM. This study evaluated patients’ daily activity and the regular exercise. It is found that only 20% of the study samples had exercised on regular basis, and about 38.3% of them attained their target HbA1c, while 27.5% for inactive patients. These results showed a significant association between exercise and attainment of HbA1c target (P = 0.021). This is inconsistent with Kayar et al (2017). Physical activity enhances insulin sensitivity in diabetic patients in the same way as it does in non-diabetic patients <sup>(21,32,43)</sup>. Several studies established that physical exercise reduces cardiovascular risk factors and improves glycemic control in type 2 DM patients <sup>(13,25)</sup>.

Smoking is associated with glycemic control in this study’s population sample. A significant association between smoking and poor achievement of HBA1c target (P = 0.004) was observed,

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despite the small percent (17.3%) of smokers with type 2 DM in our sample, several studies support this finding. Nicotine increases the level of circulating insulin-antagonistic hormones<sup>(6,20)</sup> and decreases insulin sensitivity and insulin-mediated glucose uptake<sup>(27)</sup>. Recent studies support our finding that current smokers with diabetics have higher risk for poor glycemic control than nonsmokers<sup>(31)</sup> or those who ceased smoking<sup>(10)</sup>.

Diabetes medication adherence is important in helping patients with type 2 diabetes achieving glycemic control. Medication adherence showed that more than two-thirds (69.4%) of the study sample were highly adherent to their medication. Medication adherence was significantly associated with HbA1c goal attainment in parallel with<sup>(1,17,40)</sup>. Poor medication adherence has been reported as one of the important factors associated with inadequate glycemic control<sup>(23)</sup>. Many studies concluded that poor medication adherence to be more common factor of inadequate glycemic control, and its rate was difficult to reduce<sup>(3,7)</sup>. In contrast, a large majority of this study population, 145 (68.5%) were highly adherent to their medications and showed poor controlled HbA1c which is similar to Waari et al (2018) who concluded that a good glycemic control among type 2 diabetics is the interaction of self-management measures including physical activity and diet, in addition to adherence to medication. This excludes adherence as the main factor responsible for poor glycemic control in this population.

**Conclusion:** Certain factors were significantly associated with glycemic control in patients with type 2 DM. The medication used either the drugs or their doses were one of the most important factors associated with poor glycemic control, especially when combined with the lack of patient counseling and lack of regular and close follow up. Type 2 DM failure or inadequate control could not be explained by a single factor; otherwise those factors are simultaneously associated in a complex manner with poor glycemic control to certain extent. Glycemic control in type2 DM is not just a matter of proper medications use and dosages, but other factors may contribute to medication adherence.

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## السيطرة على نسبة السكر في الدم بين مرضى داء السكري من النوع 2:

### فيما يتعلق بالالتزام بالدواء وأسلوب الحياة

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### الملخص

داء السكري من النوع الثاني (DM) هو اضطراب استقلابي مزمن يتزايد انتشاره باطراد في جميع أنحاء العالم. تعد المعالجة بطريقة واسلوب الحياة أحد الجوانب الأساسية لرعاية مرضى السكري وتشمل التغذية والنشاط البدني والإقلاع عن التدخين والرعاية النفسية الاجتماعية. إن نمط الحياة المريح غير مستقر بالإضافة إلى استهلاك الأطعمة الغنية بالدهون والكربوهيدرات العالية، وأحجام الأجزاء الكبيرة، يؤدي إلى زيادة وزن الجسم والسمنة. وما يقرب من 90 % إلى 95 % من جميع الحالات تشخيص DM لديها النوع 2 DM ، ومعظمهم يعانون من زيادة الوزن . يؤدي النشاط البدني أيضًا دورًا مهمًا في الوقاية من داء السكري من النوع الثاني، واتباع نظام غذائي صحي، وبرنامج رياضي معتدل يؤدي إلى فقدان الوزن بنسبة تتراوح بين 5 إلى 7 بالمائة، مما قد يؤدي إلى تأخير السكري من النوع 2 وربما الوقاية منه. بالنسبة للعلاقة بين التدخين ومعدل الإصابة بالسكري ، فمن المرجح أن يصاب المدخنون بالسكري مقارنة مع غير المدخنين.

والتحكم الجيد في نسبة السكر في الدم بين مرضى النوع الثاني من مرضى الدم ينطوي على التفاعل بين تدابير المعالجة الذاتية لأسلوب الحياة بما في ذلك النشاط البدني والنظام الغذائي بالإضافة إلى الالتزام بالدواء و أحد العوامل الرئيسية المساهمة هو عدم الالتزام بالدواء في النوع 2 DM وهو ما وثق بشكل جيد ليكون شائع جداً ويرتبط مع عدم كفاية تحكم نسبة السكر في الدم و زيادة المرض والوفيات وزيادة تكاليف رعاية المرضى الخارجيين ، زيارات غرفة الطوارئ ، والاستشفاء، والتعامل مع مضاعفات مرض السكري. دراسة مقطعية مستعرضة، أجريت في مراكز رعاية صحية رئيسية مختلفة في شمال الأردن ، قامت الدراسة بتوظيف أكثر من 300 مريض بالغ كانوا في نظام المعالجة المضادة نفسها لمرض السكر لمدة لا تقل عن 3 أشهر. تم الحصول على بيانات المرضى والقياسات المادية والأدوية وتاريخ المرض والكربوهيدرات الغذائية وبيانات تناول الدهون باستخدام استبيانات تم التحقق منها مسبقًا. تم جمع عينات الدم الصائم وتقييمها لـ cHbA1c.

تم تسجيل ما مجموعه 300 مريض من النوع 2 في هذه الدراسة ، (63.7 %) منهم من الإناث و(36.3 %) من الذكور. كان متوسط عمر المرضى 60.1 سنة. حوالي 52 (17.3 %) منهم كانوا مدخنين، معظمهم من الذكور. أظهر التزام الدواء أن أكثر من الثلثين (69.4 %) من عينة الدراسة كانوا ملتزمين بشدة بأدويتهم، العمر، الجنس، الالتزام بالعقاقير، ارتبطوا بشكل كبير بمستوى HbA1C فقط 41.4 % من المرضى تلقوا مشورة غذائية و58.6 % يتبعون خطة النظام الغذائي، غير مرتبطين بشكل كبير بمستوى HbA1C السيطرة على نسبة السكر في الدم في نوع 2 DM ليس مجرد مسألة استعمال الأدوية المناسبة والجرعات، ولكن قد تساهم عوامل أخرى ، الالتزام بالدواء، على الرغم من أن دورها المهم ليس العامل الرئيسي المسؤول عن ضعف مراقبة نسبة السكر في الدم في مجتمع الدراسة.

**الكلمات المفتاحية:** التحكم في نسبة السكر في الدم ، HbA1C، الالتزام بالدواء.