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Adherence to Treatment in Diabetic Patients in Lebanon

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Abstract. Non-adherence to prescribed treatment represents an important dilemma that deserves special attention especially when dealing with chronic and critical diseases as in diabetes mellitus. The objective of this study was to evaluate the level of adherence in Lebanese adult outpatients suffering from diabetes mellitus, where they may present other illness and to identify the possible predictors of non-adherence in this patient group. A questionnaire was administered face-to-face to a sample of Lebanese adults visiting two pharmacies and two endocrinologist clinics. The questionnaire covered the socio-demographics, lifestyle and health status of the patients, regimen characteristics, drug affordability, relationship with healthcare provider, and patients' attitudes and beliefs. The level of adherence was assessed using the 8-item Morisky Medication Adherence Scale. Statistical analysis was performed using SPSS version 17. Univariate associations between drug adherence and patient and drugs characteristics were tested using chi-square test. Out of the 130 patients included in this study, 64.6% were classified as adherent and 35.4% were not. In the univariate analyses, statistically significant predictors of low adherence included smoking status ($p=0.012$), level of stress and anxiety ($p=0.009$), monthly cost in LL ($p=0.049$), changing treatment plan a lot recently ($p=0.010$), taking any acute drugs ($p=0.014$), forgetting to take their pills ($p<0.001$), taking their medicine over the past two weeks ($p<0.001$), cutting back or stop taking their medication without telling their doctor because they felt worse when they took it ($p<0.001$), forgetting to bring along their medications during traveling ($p<0.001$), stop taking their medicine when they feel that the disease is under control ($p<0.001$), feeling disturbed taking medication everyday ($p<0.001$), difficulty remembering to take all their medications ($p<0.001$). As forgetfulness being the most important reason of patient non adherence, there should be a cooperative work between healthcare providers in order to overcome such a problem where they should create methods that help in educating patients and improving their adherence.

Keywords. Adherence, Treatment, Diabetic Patients, Morisky scale

Introduction

Diabetes mellitus (DM) is a growing health problem with a continuously increasing prevalence and high levels of morbidity and mortality. The International Diabetes Federation has predicted that there will be 380 million individuals with diabetes in 2025 compared with an estimated 240 million individuals in 2007, with a higher disease burden in low- and middle-income nations [1]. Effective and successful glucose control requires the use of effective medications over time. The morbidity and mortality resulting from micro- and macro-vascular complications of type 2 DM, place a considerable financial burden on individual patients and

on society [2, 3]. Although there are many factors that affect glycemic control among patients with diabetes, it is known to be improved by adherence to DM medications, and treatment utility is limited by lack of adherence [4–6].

The definition of the terms adherence, compliance, and persistence is a controversial issue among scientists, but in the recent literature, the terms ‘adherence’ and ‘compliance’ are becoming interchangeable [7, 8]. However, medication compliance initially referred to the extent to which a patient acts in accordance with the prescribed interval and dose of a dosing regimen [7, 8]. It is measured over a period of time and reported as a percentage. On the other hand, persistence refers to the duration of time from initiation to discontinuation of therapy in which the patient continues to take any amount of the medication [7, 8]. Today, the term ‘compliance’ is used less frequently because it implies that only the patients is responsible for the medical treatment [7, 8]. The term ‘adherence’ has now replaced ‘compliance’, because it includes the responsibility of the caregivers. Adherence has been defined as ‘the active, voluntary, and collaborative involvement of the patient in a mutually acceptable course of behavior to produce a therapeutic result [7, 8].

The problem with drug non-adherence doesn’t only lie in the failure to achieve treatment goals. Non-adherence has been associated with several other consequences. Non-adherence is associated with increased mortality, increased risk of hospitalization, and significantly increased medical costs [9, 10].

To improve medication adherence, the multifactorial causes of decreased adherence must be understood. The WHO classifies these factors into 5 categories: socioeconomic factors, factors associated with the health care team and system in place, disease-related factors, therapy-related factors, and patient-related factors [11].

Lots of studies targeted the issue of adherence predictors, seeking a deeper understanding of adherence barriers. Patient related characteristics include socio-demographics which were shown to affect adherence in some studies, while other studies rejected this theory. Those include sex, age, race, educational level, and marital and employment statuses [12].

Some studies showed that females adhere better to their medications; some others suggest that men adhere better, and other studies denied any gender difference [12]. Many studies addressed age as a predictor which resulted in adherence studies targeting certain age groups; discussing the issue of adherence in the elderly or in teenagers, as each of these age groups report individual barriers to adherence [13]. Other patient related factors include patient’s lifestyle. As previously mentioned, adherence to drug therapy is an indicator of an overall healthy behavior [14]. It is also an indicator of better health awareness, drug knowledge, and a positive attitude towards the treatment. In fact, recent literature shows a great interest in patient beliefs and many studies suggest that it is the most important and powerful predictor of non-adherence [14]. The fear of side-effects, little drug knowledge, the perception of overmedication, and low perceived health status were found to be critical barriers [15].

Effective strategies to improve adherence in patients with chronic disease is a relatively recent subject in drug research. Most of the studies focused on identifying barriers rather than testing methods of improving adherence. Some effective measures in improving drug compliance target the drug regimen. Reducing drug cost [10], decreasing the frequency of administration [8, 16, 17], the use of fixed dose combinations [8, 18], and the use of generic drugs [19, 20] were found to improve adherence levels.

As this subject was considered to have a highly important role in improving treatment goals, the objective of this study was to evaluate the level of adherence in Lebanese adult

outpatients suffering from diabetes mellitus, where they may present other illness and to identify the possible predictors of non-adherence in this patient group.

Patients and Methods

Sample

This is a transversal prospective pilot study assessing adherence in type II diabetic Lebanese outpatients. A convenience sample of patients was randomly selected from those visiting two pharmacies: AL CHIFAA pharmacy (Bekaa) and FARDOUN pharmacy (South Lebanon) and two endocrinologist clinics: Dr.Mazen Koteish (msharafeyye) and Dr.Ihab Fardoun (Tyre). The objectives of the study were explained to individual patient. Patients were asked if they have been taking any antidiabetic medications for more than three months. Only those who were interested and who gave their voluntary informed oral consent to participate in the study were enrolled. Patients were assured of their anonymity and confidentiality of responses. Data were collected from the 1st of February till the 30th of March 2014.

Inclusion criteria

Eligible participants were adult outpatients (≥ 25 years). The patients should be diagnosed and treated for type II diabetes mellitus. They should be taking their medication(s) for 3 months or more.

Our sample was including patients treated for type II diabetes mellitus where they may present other chronic co morbidities.

Type of study

This is a transversal prospective pilot study assessing adherence in type II diabetes mellitus Lebanese outpatients. Data were acquired through a structured questionnaire filled by an interviewer. The questionnaire included many sections. The possible reasons for medication non-adherence were included in different sections of the questionnaire and were chosen following an extensive review of literature; it contained some of the intentional and unintentional barriers or reasons for non-adherence among patients taking chronic drugs.

Questionnaire

The questionnaire consists of eight sections (Annex):

- The first section obtained information on socio-demographic data including age, sex, occupation, educational and marital status, monthly income, and the presence of a care provider at home.
- The second section obtained information about lifestyle data: smoking status (non smoker, ex smoker or smoker), sports (doing or not), stress level (none, sometimes or all the time), and memory status (forgetfulness: never, sometimes or all the time).
- The third section assessed the patient's health follow-up and diet adherence.
- The fourth section was concerned with the drug regimen: the number of drugs patient takes daily, the number of drugs the patient recalls their names, the duration since the administration of the diabetic drug, drug cost, medical insurance (yes or no), having difficulty paying for their medications (yes or no), previous side-effects (yes or no), reaction towards such side-effects (told the doctor or stopped taking the drug), administration of acute drugs (yes or no), and whether the drug regimen has changed a lot recently (yes or no).
- The fifth section assessed the patient's relationship with the healthcare providers: the physician and the pharmacist. The patients were asked whether the healthcare providers explained the regimen and disease and the number of providers they consult.

- The sixth section provides information about the patients' attitudes, behaviors, knowledge, and patients' opinions on some of the likely reasons for their non-adherence.
- The seventh section assessed patient adherence using 8-MMMAS (8-item modified Morisky medication adherence scale).
- The last section provides information about measuring HbA1C, duration between two measurements and the value of last measurement.

Scale validation

Patient adherence was assessed through self-report using 8-MMMAS (8-item modified Morisky medication adherence scale) in the seventh section. The scale is a widely used and validated method to assess patient adherence/non-adherence to drug regimen. It includes questions about intentional and non-intentional non-adherence. The Arabic version was used in the questionnaires. The scale consists of 8 questions about intended and non-intended drug non-adherence. The first 7 questions are dichotomous (yes/no) while the last question is a 5-item Likert scale. It was dichotomized for the statistical analysis so the final score was 8. A score between 0-5 represents non-adherence and 6 through highest represents adherence. An addition for all the eight scores was done for each patient. As the score increase, it indicates an increase in adherence for the patient.

Morisky questions and its scores

Question	Yes	No
1- Do you sometimes forget to take your pills?	0	1
2- People sometimes miss taking their medications for reasons other than forgetting. Thinking over the past two weeks, were there any days when you did not take your medicine?	0	1
3- Have you ever cut back or stopped taking your medication without telling your doctor, because you felt worse when you took it?	0	1
4- When you travel or leave home, do you sometimes forget to bring along your medications?	0	1
5- Did you forget to take your medications yesterday?	0	1
6- When you feel like your disease is under control, do you sometimes stop taking your medicine?	0	1
7- Taking medication everyday is a real inconvenience for some people. Do you ever feel hassled about sticking to your treatment plan?	0	1
	Sometimes/Often/Always	Never/Rarely
8- How often do you have difficulty remembering to take all your medications?	0	1

Sample size

The study was done on a convenience sample, as this is a pilot study.

Data entry and statistical analysis

Data was entered and analyzed using SPSS version 17.

Descriptive statistics including frequency, mean, and standard deviation were used to summarize patients' baseline socio-demographic data and evaluate distribution of responses.

The dependent variable was the medication adherence: a dichotomized MMMAS.

Chi-square test was done. A p-value of 0.05 or less was considered to be statistically significant in the tests.

Results

One hundred and thirty patients were included in this study. The mean age was 59.39 years \pm 13.12 and ranged between 25 and 88 years. 56.2% were males and no significant age difference ($p=0.578$) was found between males (58.82 years \pm 13.71) and females (60.12 years \pm 12.47)

General characteristic of the study population

The majority of the study population was married (83,8%) and had an elementary education (43.8%). 45.4% of the study sample was unemployed. 79% reported having a daily break during their working hours. 92% of the working group has a monthly income between five hundred and one million LL. 38.5% of the studied sample had a caregiver at home (Table 1).

Table 1: General characteristics of the study population

Characteristics		
Age in years	Mean 59.39 SD 13.127 Range [27-88]	
	Frequency	Valid Percentage (%)
Sex		
Males	73	56.2
Females	57	43.8
Marital Status		
Single	8	6.2
Married	109	83.8
Divorced	6	4.6
Widowed	7	5.4
Educational Level		
Illiterate	34	26.2
Elementary	57	43.8
Intermediate	21	16.2
Secondary	10	7.7
University	8	6.2
Employment Status		
Unemployed	59	45.4
Employee	44	33.8
Self-employed	27	20.8

Working Hours		
[3-8] hours	106	81.5
[7-16] hours	24	18.5
Daily Break		
Yes	79	60.8
No	51	39.2
Monthly Income		
[500,000-1,000,000 L.L.]	92	70.8
[1,000,000-2,000,000 L.L.]	26	20
[>2,000,000 L.L.]	12	9.2
Care provider at home		
No	80	61.5
Yes	50	38.5

Healthy Lifestyle characteristics of the study population

Around half of the population were non-smokers (50.8%), had a very stressful life (25.4%), and had moderate memory problems (66.9%). Only 39.2% of the sample did sports on a regular basis, mainly walking (100%) four to seven times per week. Only 16.2% admitted not regularly monitoring important disease indicators (BP, BS, Cholesterol...). Among those who had a specific physician recommended diet (93.8%), 25% rarely adhered to this diet (Table 2).

Table 2: Healthy lifestyle characteristics of the study population

Characteristics:		
	Frequency	Valid Percentage (%)
Smoking status		
Non-smoker	66	50.8
Ex-smoker	38	29.2
Smoker	26	20.0
Regular sports		
Yes	51	39.2
No	79	60.8
Type of sports		
Walking	51	100
Missing	79	
Frequency per week		
[2-3]	21	41.17
[4-7]	30	58.82
missing	79	

Levels of stress and anxiety		
None	13	10.1
Moderate	84	64.6
Severe	33	25.4
Memory (forgetfulness)		
Never	28	21.5
Sometimes	87	66.9
All the time	15	11.5
Do you regularly measure your BP, BS, Cholesterol, etc?		
No	21	16.2
Yes	109	83.8
Did the Dr recommend a certain diet for you to follow?		
No	8	6.2
Yes	122	93.8
How often do you adhere to this diet?		
Rarely	25	20.49
Sometimes	63	51.63
Always	34	27.86
Missing	8	

Drug regimen related characteristic

On average, patients reported taking 4 medications (mean=4.33 \pm 2.10). The number of drugs per patient ranged between 1 and 13. Patients recalled approximately 4 medications (mean=4.06 \pm 1.86) of the names of their drugs. The average monthly drug cost was 136,46 LL (\pm 107,95). Patients visited 1.65 (\pm 1.03) physicians every 5.21 (\pm 3.31) months (Table 3).

Table 3: Drug regimen related Characteristics

	Mean	SD \pm	Range (Min-Max)
Number of chronic drugs	4.33	2.103	1-13
Number of drugs the patient could state their names	4.06	1.867	1-12
Duration since the administration of the first chronic drug in months	48.63	34.841	3-168
Monthly cost of drugs in LL	136,469	107,951	8,000-600,000
Number of Drs patient visits	1.65	1.032	1-7
Frequency of visits (in months)	5.21	3.308	1-12

Paying for drugs and treatment plan

Approximately half of the patients (49.2%) had a medical insurance; with 32.3% reported having a difficulty paying for their drugs. 21.5% experienced a previous drug side-effect, where 92.85% of them told their physician. Most of the patients did not experience a change in their treatment plans in the past months (77.7%). Along with their chronic medications, 81.5% used OTC painkillers or other acute drugs (Table 4).

Table 4: Paying for drugs and treatment plan

	Frequency	Valid Percentage (%)
Medical insurance		
No	64	49.2
Yes	66	50.8
Difficulty paying for drugs		
No	88	67.7
Yes	42	32.3
Ever experienced a drug related side-effect?		
No	102	78.5
Yes	28	21.5
Patient reaction		
Told Dr	26	92.85
Stopped taking drug	2	7.14
Missing	102	
Did your treatment plan change a lot recently?		
No	101	77.7
Yes	29	22.3
Do you take any acute drugs (pain killer, ATB, etc)		
No	24	18.5
Yes	106	81.5

Relationship with healthcare providers

Regarding the relationship with the care provider, 54.6% considered the physician (rather than the pharmacist) to be their primary reference regarding drugs and disease conditions. 91.5% reported that the physician explained the pathology and its severity and 92.3% that the physician asked about previous medications before prescribing new ones. 92.3% of the patients reported that the physician explained to them how to take their medications and 95.4% reported that the pharmacist did (Table 5).

Table 5: Relationship with healthcare providers

	Frequency	Valid Percentage (%)
Whom do you consult more regarding your drugs and disease?		
Dr	71	54.6
Pharmacist	59	45.4

Pharmacist		
Did the Dr ask you about drugs you are already taking before prescribing new one?		
Yes	120	92.3
No	10	7.7
Did the Dr explain to you how to take your medications?		
Yes	120	92.3
No	10	7.7
Did you fully understand how to take them?		
Yes	121	93.1
No	9	6.9
Did the Pharmacist explain to you how to take your medications?		
Yes	124	95.4
No	6	4.6
Did you fully understand how to take them?		
Yes	126	96.9
No	4	3.1

Patient attitudes and knowledge

Although about half of the study population(40.8%) responded that nothing would make them stop treatment, they thought that they are over-medicated(40%), reported postponing physician's appointments(55.4%), and considered forgetfulness(38.5%) as the most important reason that would make them stop their medications. The majority believed that they cannot skip or double doses on their own (92.3%), would not stop their treatment even if advised to do so (90.8%), and believed that their treatment is improving their condition (92.3%) (Table 6).

Table 6: Patient attitudes and knowledge

	No	Yes
Do you think you are over-medicated?	60 %	40 %
A person can take double the dose to feel better or skip a dose if he felt worse	92.3 %	7.7 %
Do you usually postpone doctor's appointments?	44.6%	55.4 %
If someone you trust advised you to stop your treatment, would you?	90.8 %	9.2 %
Do you believe that drug therapy will improve your condition?	7.7%	92.3 %
What is the most important reason that would make you stop taking your drugs?		

	Frequency	Valid (%)	Percent
Drug Cost	23	17.7	
Forgetfulness	50	38.5	
Regimen is too complicated	4	3.1	
Nothing would make me stop	53	40.8	
Total	130	100	

HbA1c measuring

The majority of the sample patients (93.8%) reported measuring the value of HbA1c regularly with a duration of 6 months (mean 5.92 ± 3.35 , range 1-24). The value of last measurement was 8.41 ± 1.81 (range 5-18) (Table 7).

Table 7: HbA1c measuring

	Frequency	Percentage (%)
Do you measure the value of HbA1C regularly?		
No	8	6.2
Yes	122	93.8
What is the duration between two measurements(in months)	Mean:5.92	SD:3.355
		Range (1-24)
What is the value of last measurement?	Mean:8.41	SD:1.817
		Range (5-18)

Adherence patterns

A large percentage of the sample patients reported forgetting to take their medications sometimes (64.6%), and (72.3%) having difficulty remembering to take all their medications. Only (16.9%) of the patient reported cutting back their medications when they felt better or worse. Although sticking to the treatment plan was considered as inconvenient by (63.1%) of patients, only (24.6%) reported forgetting their drugs at home when leaving. (17.7%) reported not taking their drugs on the day prior to the interview, (30%) in the previous 2 weeks.

The mean 8-MMMAS of the sample was found to be 5 (± 1.6) and ranged from 0 to 8 with a median value of 5. The value of the mean was taken as a determinant for adherence status where if the summation of the score of the 8-MMMAS questions was 0-5, the patient was considered as non adherent, and if the value was 6 and beyond, the patient being adherent.

64.60% of the sample was classified as adherent, and 35.40% are known to be non-adherent (Table 8).

Table 8: 8-MMMAS Answers

Question	Yes	No
1- Do you sometimes forget to take your pills?	64.6 %	35.4 %
2- People sometimes miss taking their medications for reasons other than	30%	70 %

forgetting. Thinking over the past two weeks, were there any days when you did not take your medicine?		
3- Have you ever cut back or stopped taking your medication without telling your doctor, because you felt worse when you took it?	10 %	90 %
4- When you travel or leave home, do you sometimes forget to bring along your medications?	24.6 %	75.4 %
5- Did you forget to take your medications yesterday?	17.7 %	82.3 %
6- When you feel like your disease is under control, do you sometimes stop taking your medicine?	16.9%	83.1 %
7- Taking medication everyday is a real inconvenience for some people. Do you ever feel hassled about sticking to your treatment plan?	63.8%	36.2 %
	Sometimes/Often/Always	Never/Rarely
8- How often do you have difficulty remembering to take all your medications?	72.3%	27.7%
	Adherent	Non-Adherent
MMMAS percentage	64.6%	35.4 %

Comparison between adherent and non adherent patients

Sociodemographic characteristics did not seem to predict the variation of adherence, as none of which showed any significance in Chi-square test (Table 9)

Table 9: Comparison between adherent and non adherent patients concerning sociodemographic predictors

Predictor	Non-adherent (%)	Adherent (%)	P-value
Sex			
Males	47.7	60.5	0.166
Females	52.3	39.5	

Educational Level:			
Illiterate	31.8	23.3	
Elementary	47.7	41.9	
Intermediate	18.2	15.1	
Secondary	0	11.6	
University	2.3	8.1	0.071
Marital Status:			
Single	0	9.3	
Married	90.9	80.2	
Divorced	6.8	3.5	
Widowed	2.3	7	0.078
Employment status:			
Unemployed	56.8	39.5	
Employee	27.3	37.2	
Self-employed	15.9	23.3	0.172
Daily break:			
No	63.6	50	
Yes	29.5	44.2	
Missing	6.8	5.8	0.280
Monthly income:			
[500000-1000000]	77.3	67.4	
[1000000-2000000]	18.2	20.9	
>2000000	4.5	11.6	0.377
Care provider at home:			
No	56.8	64	
Yes	43.2	36	0.429

Concerning healthy lifestyle factors, smoking status ($p=0.012$) and the level of stress and anxiety ($p=0.009$) were found to be significantly different between adherent and non adherent patients while other factors as doing regular sports, measuring important health indicators , and adherence to a healthy diet were all not found to be significantly important predictors of high adherence (table 10).

Table 10: Comparison between adherent and non adherent patients concerning healthy lifestyle

Predictor	Non-adherent (%)	Adherent (%)	P-value
Smoking status:			
Non smoker	68.2	41.9	
Ex-smoker	22.7	32.6	
Smoker	9.1	25.6	0.012
Regular sports:			
No	68.2	57	
Yes	31.8	43	0.216

Level of stress and anxiety:			
None	0	15.1	0.009
Some times	68.2	62.8	
All the time	31.8	22.1	
Memory (forgetfulness)			
Never	22.7	20.9	0.973
Sometimes	65.9	67.5	
All the time	11.4	11.6	
Do you regularly measure your BP, BS, Cholesterol, etc?			
No	15.9	16.3	0.957
Yes	84.1	83.7	
Did the Dr recommend a certain diet for you to follow?			
No	6.8	5.8	1
Yes	93.2	94.2	

Regarding drug related factors, neither the drug number nor the duration since administration of treatment were suggested to negatively affect the adherence level.

Medical insurance, difficulty paying for drugs and experiencing a drug related side effects were not found to be significantly different between adherent and non adherent patients.

However, statistically significant predictors of low adherence were found in monthly cost in LL ($p=0.049$), changing treatment plan a lot recently (0.010) and taking acute drugs (pain killer, ATB, etc) ($p=0.014$) (Table11).

Table 11: Comparison between adherent and non adherent patients concerning drug related factors

Predictor	Non-adherent (%)	Adherent (%)	P-value
Number of chronic drugs			
1-4	47.7	62.8	0.107
4-8	50	31.4	
>8	2.3	5.4	
Duration since administration of first chronic drug			
6-54	58.1	69.8	0.265
54-108	23.3	20.9	
>108	18.6	9.3	
Monthly cost in LL			
8000-200000	70.5	87.2	

200000-400000	25	11.6	
>400000	4.5	1.2	0.049
Medical insurance			
No	38.6	54.7	
Yes	61.4	45.3	0.084
Difficulty paying for drugs			
No	70.5	66.3	
Yes	29.5	33.7	0.630
Ever experienced a drug related side-effect?			
No	72.7	81.4	
Yes	27.3	18.6	0.255
Did your treatment plan change a lot recently?			
No	90.9	70.9	
Yes	9.1	29.1	0.010
Do you take any acute Drugs (pain killer, ATB, etc)			
No			
Yes	6.8	24.4	
	93.2	75.6	0.014

With respect to patient beliefs, patients who believed they are over-medicated, who thought they could skip and double doses, who usually postpone physician's appointments, who would stop the treatment based on others' advice and who don't think that drug therapy will improve their condition were not found to have significantly lower adherence rates (Table 12).

Table 12: Comparison between adherent and non adherent patients concerning patients' beliefs

Predictor	Non-adherent (%)	Adherent (%)	P-value
Do you think you are over-medicated?			
No	61.4	59.3	
Yes	38.6	40.7	0.820
A person can take double the dose to feel better or skip a dose if he felt worse			
No	88.6	94.2	
Yes	11.4	52.3	0.327
Do you usually postpone doctor's appointments?			
No	38.6	47.7	
Yes	61.4	53.6	0.300

If someone you trust advised you to stop your treatment, would you? No Yes	88.6 11.4	91.9 8.1	0.53.9
Do you believe that drug therapy will improve your condition? No Yes	11.4 88.6	5.8 94.2	0.305

In addition good drug knowledge, positive physician-patient relationships, appropriate physician counseling, understanding how to take their medications, explaining the disease to them by physician, teaching them how to take the drugs properly, and asking them about their previous medications, as were supposed to be predictors of good adherence, were not significantly different between adherent and non-adherent patients (Table 13).

Table 13: Comparison between adherent and non adherent patients concerning relationship with healthcare providers

Predictor	Non-adherent (%)	Adherent (%)	P-value
Did the Dr discuss with you the disease and its severity? No Yes	9.1 90.9	8.1 91.9	1
Did the Dr ask you about drugs you are already taking before prescribing new one? No Yes	6.8 93.2	8.1 91.9	1
Did the Dr explain to you how to take your medications? No Yes	6.8 93.2	8.1 91.9	1
Did you fully understand how to take them? No Yes	4.5 95.5	8.1 91.9	0.717

Did the Pharmacist explain to you how to take your medications?			
No	6.8	3.5	0.406
Yes	93.2	96.5	
Did you fully understand how to take them?			
No	4.5	2.3	0.604
Yes	95.5	97.7	

On the other hand, statistically significant predictors of low adherence included most important reason that would make them stop taking their drug (mainly forgetfulness) ($p=0.009$), forgetting to take their pills ($p<0.001$), taking their medicine over the past two weeks ($p<0.001$), cutting back or stop taking their medication without telling their doctor, because they felt worse when they took it ($p<0.001$), forgetting to bring along their medications during traveling ($p<0.001$) (Table 14).

Table 14: Comparison between adherent and non adherent patients concerning remembering their medications

Predictor	Non-adherent (%)	Adherent (%)	P-value
What is the most important reason that would make you stop taking your drug			0.009
Drug cost	13.6	19.8	
Forgetfulness	56.8	29.1	
Regimen too complicated	4.5	2.3	
Nothing would make me stop	25	48.8	
Do you sometimes forget to take your pills			<0.001
No	9.1	48.8	
Yes	90.9	51.2	
Were there any days when you did not take your medicine over the past 2 weeks?			<0.001
No	45.5	82.6	
Yes	54.5	17.4	
Have you ever cut back or stopped			

taking your medication without telling your doctor, because you felt worse when you took it? No Yes	75 25	97.7 2.3	<0.001
When you travel or leave home, do you sometimes forget to bring along your medications? No Yes	45.5 54.5	90.7 9.3	<0.001
Did you forget to take your medications yesterday? No Yes	18.2 81.8	17.4 82.6	0.917

Stop taking their medicine when they feel that the disease is under control ($p < 0.001$), feeling disturbed taking medication everyday ($p < 0.001$), difficulty remembering to take all their medications ($p < 0.001$) were also shown to be significantly different (Table 15).

In addition, results showed no significant difference neither regarding HbA1c measuring regularly nor its value. Only 19.6% of the adherent patients known to be within the normal range of HbA1c value which was not significantly different with non adherent patients (Table 15).

Table 15: Comparison between adherent and non adherent patients concerning feeling disturbed and HbA1c measuring

Predictor	Non-adherent (%)	Adherent (%)	P-value
When you feel like your disease is under control, do you sometimes stop taking your medicine? No Yes	59.1 40.9	95.3 4.7	0.000
Taking medication everyday is a real inconvenience for some people. Do you ever feel hassled about sticking to your treatment plan? No Yes	6.8 93.2	51.2 48.8	0.000

How often do you have difficulty remembering to take all your medications?			
Never/rarely	4.5	39.5	0.000
Sometimes/often/always	95.5	60.5	
Do you measure the value of HbA1c regularly?			
No	6.8	5.8	1
Yes	93.2	99.2	
HbA1c value			
Within normal range	16.7	19.6	0.679
Outside normal range	83.3	80.4	

Our sample was considered to be highly adherent to treatment (64.6%) where the main cause of non-adherence in non-adherent patients was forgetfulness, where there was no other reason that makes patient stop his drug other than forgetting it.

Discussion

Our study aimed to evaluate the level of adherence in Lebanese adult outpatients suffering from diabetes mellitus, where they may present other illness and to identify the possible predictors of non-adherence in this patient group.

Yet the analysis of the possible predictors highlighted many major issues. None of the socio-demographics significantly affected the adherence level; many previous studies have found similar results [13, 14, 22].

Even though half of the patients had no medical insurance and about 67.7% of them had a difficulty in paying for drugs, this presumption did not stand upon data analysis. Neither drug cost per patient, nor the presence of a medical insurance could predict the adherence outcome in our patient group. This finding is supported by a lot of evidence from the literature where drug cost fails to be a major predictor relative to other more important criteria [23].

In addition, although a good physician-patient relationship and proper counseling by the physician had an effect in increasing the level of adherence in chronically ill patients, but the difference was not significant [8, 22, 24, 25].

Data suggest that the patient should be ‘engaged’ in the treatment plan rather than just ‘counseled’ about their drugs. It is very crucial that the physician takes into account the individual preference of patients when prescribing new medications and to involve the patient in the decision-making. This communication is especially important in older patients with multiple diseases and hence multiple drugs or ‘poly pharmacy’ [8, 22, 24, 25].

Declining memory was found to be the most major predictor of adherence in our sample. This can be associated with the idea that in older patients, declining cognitive function plays a major role in their drug intake [13]. These patients are more likely to forget to take their medications and also forget the physician’s instructions. So even if proper counseling is assumed, it will have a transient effect because none of the information will be retained by the patient. This problem can be overcome by the use of different aids. The use of clear labels and encouraging the patient to use reminders are helpful techniques to combat this problem. The physician should work with the patient on simplifying the drug regimen. Blistered medications

which divide doses over the day and week can help a great deal in remembering to take medication in appropriate doses and time-intervals. The physician should also work on simplifying the drug regimen through omitting any unnecessary medication. The number of pills per day and dose frequency can be reduced by using fixed dose combinations [8, 16].

A meta-analysis on the use of fixed-dose combination concluded that a 26% decrease in the risk of non-adherence is associated with the use of these combinations [18]. Overall it is believed that patients adhere better to once-daily medications rather than three or four-times-daily drugs, as they find it more convenient and appropriate with their daily activities [17].

Patients should be educated therapeutically about their treatments; the physician plays an important role in such an issue.

The physician with the aid of clinical pharmacist should create methods and apply ways that help in increasing the knowledge of patients about their treatment; they also should urge the patients and help them in remembering their drugs in the precise time and in the correct way.

The belief that treatment will not improve the condition was significantly related to low adherence levels. Also stopping therapy without telling the doctor because of feeling worse, feeling the disease is under control and feeling disturbed about sticking to their treatment, all were shown to be significantly different between adherent and non-adherent patients.

Such results highlight the importance of explaining for patients about the side effects and sequels that would happen without taking their medication. Patients may not be aware about the importance of treatment in preventing many hazards from occurrence.

Healthcare providers should act together in order to overcome this problem and thus by doing such a step, many problems will be avoided.

The value of HbA1C was not shown to be significantly different between adherent and non adherent patients which reflect the fact that a fluctuations in the patient's answers had occur and an overestimation of the adherence level.

Patients may not say the truth about their use of drugs and adherence to treatment regimen, and in our study we did not ask patients about details that may help us in estimating adherence rather than depending on them only as a source of information

It is important to note that this study has several limitations. The method of drug adherence was self-report which suggests a possibility of recall bias. Self-report when compared to other methods of drug adherence assessment is generally believed to overestimate adherence level [26, 27].

But yet self-report is still believed by many to be equally precise as other physical methods such as pill-count or electronic monitoring [28].

Another limitation is that the study did not compare between patients according to disease severity, disease type, or drug class. These factors were recently believed to be less significant when compared to other patient related factors such as patient attitudes or beliefs [20, 29].

Conclusion

This study focused on many issues concerning drug adherence in Lebanese diabetic outpatients. Although the studied sample was considered as highly adherent, the findings reassure the fact that patients should be frequently reminded of their treatment by several ways - like encouraging patients to use memory aids - as forgetting was shown to be the major reason of non-adherence.

As diabetes was called “the silent killer disease”, patient’s education should be regarded as a cornerstone for treatment success. Teaching them about the importance of the treatment and the severe consequences of their disease that may persist if they cut back their therapy because of feeling worse, unimproved or feeling disturbed, will motivate patients to take drugs as instructed.

So, when patients realize and understand the necessity of their treatment and the consequences of under-treatment, their beliefs will be totally changed about their therapy. Physicians and pharmacists shall put more effort in identifying patients with declining memory and try to help them take their medications sufficiently and on time, they also should make an effort in educating patients and explaining for them about their disease and the importance of taking the proper treatment on time to avoid future hazards.

More studies that assess adherence and predict factors affecting this adherence are needed to provide strong evidence on those barriers. Additional studies as well are needed to highlight the effectiveness of interventions suggested to aid and improve adherence.

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