# Association between HbA1c and Serum Lipid Profile among a sample of Iraqi Patients with Type2 Diabetes Mellitus

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### Abstract:

**Background:** Diabetes mellitus is considered a public health concern. Around 1.4 million of Iraqis have diabetes. Reported T2DM prevalence in Iraq ranges from 8.5% (IDF—age-adjusted) to 13.9%. A local study including more than 5400 people in the city of Basrah, Southern Iraq, reported a 19.7% age-adjusted prevalence of diabetes in subjects aged 19 to 94 years. Diabetes is a serious risk factor for atherosclerotic cardiovascular disease (ASCVD) and an important cause of mortality. ASCVD is the commonest cause of death in the western world. Diabetes was identified as a high risk condition for ASCVD. In adults with diabetes with ASCVD or multiple ASCVD risk factors it is important to prescribe high intensity statin to reduce LDL at least by 50%.

**Objective:** To investigate the association between dyslipidemia and HbA1c and to detect the benefit of using statins to decrease the risk of CVD.

**Cases and methods:** A prospective, single-dose, non-randomized study was carried out on 1st November 2020 until 31th July 2021 at a private clinic in Wasit governorate-Iraq; and included patients with type 2 diabetes. Clinical and biochemical laboratory assessment and re-assessment were carried out before and after 3 months of receiving Rosuvastatin 20 mg/ day. A paper questionnaire form was used, including sociodemographic and clinical features (age, gender, weight, height, waist circumference and biochemical markers [total cholesterol, HDL, LDL, TG and HbA1c]).

**Results:** A total of 256 type 2 DM patients were included in the study; receiving 20 mg of rosuvastatin as a single dose for 3 consecutive months. Of the study group 83 (32.4%) were males and 173 (67.6%) were females. The mean age of males was  $(52.1\pm10.49)$  years and that of females was  $(53.1\pm10.41)$  years. The mean values of (BMI, Waist circumference WC, HbA1c, LDL, TG, and cholesterol) for the studied sample after treatment were significantly lower than mean values before treatment, except that for HDL where it was significantly higher after treatment of rosuvastatin single dose among all patients, P<0.05; in spite of that gender and age groups shows no significant differences between each other, P>0.05. In patients with total cholesterol level 200 or more; a significant elevation in mean of HDL; P<0.01

**Conclusions:** All age groups and both gender have benefited from rosuvastatin treatment in the reduction of lipid cholesterol as well as HbA1c. Rosuvastatin can be used by type 2 diabetics regardless of age and gender. HbA1c can be used as a predictor of dyslipidemia in type 2 diabetes.

Keywords: cardiovascular diseases, diabetes mellitus, dyslipidemia, HbA1c, HDL, rosuvastatin

### Introduction

Diabetes mellitus (DM) is considered as a public health concern. Diabetes is a serious risk factor for Atherosclerotic cardiovascular disease (ASCVD) and an important cause of mortality. Risk factors for developing type 2 diabetes (T2DM) are: Older age, high plasma glucose, metabolic syndrome, high body mass index (BMI), and high glycated hemoglobin (HbA1c > 6%). [1-5,] Around 1.4 million of Iraqis have diabetes. Reported T2DM prevalence in Iraq ranges from 8.5% (IDF—age-adjusted) to 13.9%. A local study including more than 5400 people in the city of Basrah, Southern Iraq, reported a 19.7% ageadjusted prevalence of diabetes in subjects aged 19 to 94 years. [6, 7] Diabetic patients are 2 to 4 times more likely to die due to ASCVD than non-diabetics. It

\*Department of Internal Medicine, Al Zahraa Teaching Hospital, Wasit Health Directory, Ministry Of Health, Iraq. E-mail: <u>deef561@gmail.com</u> may lead to microvascular diseases (retinopathy, neuropathy, nephropathy) and/or macrovascular diseases (ASCVD) that manifest with coronary artery disease (CAD), stroke and peripheral arterial disease)[8-9]. It may also lead to the impairment of the cardiac diastolic performance and muscle contractility. [10] About two thirds of adults with diabetes over the age 65 die as a result of (CAD) and the risk is increased with the addition of other risk factors.[11, 12] HbA1c predicts the risk of developing diabetic complications in patients with DM.[13] ASCVD is the commonest cause of death in the western world. [14] Diabetics usually suffer from dyslipidemia which characterized is by hypertriglyceridemia (TG) with reduced levels of high-density lipoprotein (HDL) cholesterol. [15, 12] Diabetes had been identified as a high-risk condition for ASCVD. In adults with diabetes with ASCVD or multiple ASCVD risk factors it is important to

JFac Med Baghdad 2021; Vol.63, No. 3 Received: Aug., 2021 Accepted: Sept., 2021 Published: Oct., 2021 prescribe high intensity statin to reduce LDL at least by 50%. [16]. Dyslipidemia is a common problem in the Iraqi population. A study conducted on diabetic patients in the Kurdistan region of Iraq reported that hyperlipidemia was more dominant among diabetic patients with retinopathy [17].

### Cases & Methods:

A prospective non randomized single dose study was carried out at a private clinic in Wasit governorate, Iraq; and included patients with type 2 diabetes. The study started on 1<sup>st</sup> November 2020 until 31<sup>th</sup> July 2021.

**Inclusion criteria:** All type 2 diabetic patients who were managed in the private clinic during the study period.

**Exclusion criteria:** Patients with Type 1 DM, thyroid dysfunction, any lipid-lowering or anti-obesity agents and history of Rosuvastatin intake.

A convenience sampling was carried out at a private clinic, 4 hrs./ day, 6 days/week for 3 consecutive months and biochemical lab assessment and reassessment was carried out before and after three months of receiving Rosuvastatin 20 mg/ day.

A paper questionnaire form was used and included the following data: Demographic and clinical features (Age, gender, measuring weight, height, waist circumference and biochemical markers including [total cholesterol (TC), HDL, LDL, TG and HbA1c levels] were analyzed twice (before and after three months of taking Rosuvastatin 20 mg/ day).

**Statistical analysis:** Descriptive analyses of the variables were expressed as frequencies and percentages for categorical data. The mean and standard deviation was used for quantitative data. A paired two sample t test was used to assess the differences between variables before and after management. We used  $P \le 0.05$  to determine statistical significance.

**Ethical Approval:** This study was approved by MOH, Wasit Health Directorate. The researcher paid for the biochemical lab tests for patients before and after the treatment, after taken verbal consent from them.

### **Results:**

A total of 256 patients with type 2 DM were included in this study; receiving 20 mg of rosuvastatin as a single daily dose for 3 consecutive months. Of the study group. 83 (32.4%) were males and 173 (67.6%) were females, with no significant difference between the mean ages of males ( $52.1\pm10.49$ ) and females ( $53.1\pm10.41$ ), table 1.

### Table 1: Mean age of males and femalesparticipating in the study

Gender	Ν	%	Mean	$\pm$ SD	P-
			age		Value
Male	83	32.4	52.1	10.49	0.47
Female	173	67.6	53.1	10.41	_
Total	256	100	52.8	10.43	

Paired samples t-test in table 2 displays that the mean values of (BMI, WC, HbA1c, LDL, TG, and cholesterol) among studied samples after treatment were significantly lower than mean difference before treatment, except that for HDL; where it was significantly higher after treatment, P < 0.001. The mean values of VLDL shows no significant difference according to the treatment, (P=0.27), table 2.

### Table 2: Differences in the means of variablesbefore and after treatment

Variables		Mean	SD	P value	
DML tra/m <sup>2</sup>	Before treatment	30.8	5.57	- <0.001	
BMI kg/m <sup>2</sup>	After treatment	28.4	4.16	<0.001	
WC cm	Before treatment	107.2	9.35	-0.001	
	After treatment	101.7	11.48	< 0.001	
HbA1c mg/dl	Before treatment	9.4	2.09	-0.001	
	After treatment	7.9	1.32	- <0.001	
HDL mg/dl	Before treatment	42.4	15.96	- <0.001	
	After treatment	53.1	14.59	- <0.001	
LDL mg/dl	Before treatment	114.9	38.10	-0.001	
	After treatment	68.6	38.68	- <0.001	
TC	Before treatment	187.4	118.57	- <0.001	
TG mg/dl	After treatment	110.1	78.26	- <0.001	
Total Cholesterol mg/dl	Before treatment	185.9	44.68	- <0.001	
	After treatment	164.8	29.98	- <0.001	
VLDL mg/dl	Before treatment	46.0	42.12	0.27	
	After treatment	42.4	23.05	- 0.27	

According to the age group and gender, mean values of HbA1c were reduced significantly after treatment among age groups (<40 years and  $\geq$  40 years) P=0.046, P<0.001 respectively and among gender, P<0.05. Figure 1.



Figure 1: Differences in the means of HbA1c according to the age groups and gender of study population before and after treatment

Similarly, according to the age group and gender, mean values of total cholesterol were reduced significantly after treatment among age groups (<40 years and  $\geq$  40 years, P=0.049 and P<0.001 respectively and among both gender, P<0.01, Figure 2.



## Figure 2: Differences in the means of cholesterol according to the age groups and gender of study population before and after treatment

When the patients were grouped according to their HbA1c level (<7 and  $\geq$ 7), significant reductions were observed after treatment in mean of (BMI, WC, LDL, and TG) with significant elevation in mean of HDL (P<0.05) for the first group and significant reductions were observed after treatment in mean of (BMI, WC, LDL, TG, and cholesterol) with significant elevation in mean of HDL (P<0.05) for the second group. There was no significant difference in mean VDLD after treatment in both groups (P>0.05), table 4.

Table 3: Differences in the means of variablesbefore and after treatment according to theHbA1c of studied sample

Variables	HbA1	Before		After	After		
	с	treatm	treatment		treatment		
		Mea	SD	Mea	SD		
		n		n			
BMI	<7	31	4.57	29	4.1	0.014	
kg/m <sup>2</sup>	≥7	30.8	5.55	28.4	4.19	<0.00 1	
WC cm	<7	106. 4	7.48	98.2	11.1 5	0.004	
	≥7	107.	9.45	101.	11.4	< 0.00	
		3		9	8	1	
HDL mg/dl	<7	38.6	15.69	55	14.1	<0.00 1	
	≥7	42.7	38.08	52.9	14.6 6	<0.00 1	
LDL	<7	116.	37.49	63.5	36.3	< 0.00	
mg/dl		3			8	1	
-	≥7	114. 8	38.28	69	38.9	<0.00 1	
TG mg/dl	<7	186. 8	72.03	103	67.6	<0.00 1	
	≥7	187.	121.4	110.	79.1	< 0.00	
		4	5	6		1	
Cholester ol mg/dl	<7	182. 8	43.05	165. 2	26.1	0.13	
-	≥7	186. 2	44.11	164. 7	30.2 6	<0.00	
VLDL mg/dl	<7	50	45.1	34.7	0 22.1 1	0.25	
0	≥7	46	42	43	23.1	0.416	

In patients with total cholesterol level less than 200, a significant reduction was observed after treatment in means of (BMI, WC, LDL, TG, and HbA1c) with a significant elevation in mean of HDL, P<0.01. In patients with total cholesterol level 200 or more, a significant reduction was observed after treatment in means of (BMI, WC, LDL, TG, and HbA1c) with a significant elevation in mean of HDL; P<0.01. No significant difference was found in the mean of VDLD after treatment whatever the level of total cholesterol, P>0.05, table 4.

Table 4: Differences in the means of variablesbefore and after treatment according to the totalcholesterol of studied sample

cholester of of studied sample								
Variabl	Choleste	Before		After	After			
es	rol	treatment		treatm	treatment			
		Mea	SD	Mea	SD			
		n		n				
BMI	<200	30.8	5.36	28.5	4.00	< 0.0		
kg/m <sup>2</sup>						01		
	≥200	30.8	5.88	28.3	4.40	< 0.0		
						01		
WC cm	<200	107.	9.16	101.	11.5	< 0.0		
		6		8	0	01		
	≥200	106.	9.66	101.	11.5	< 0.0		
		4		4	0	01		
HDL	<200	41.4	16.6	52.2	16.4	< 0.0		
mg/dl			9		0	01		
	≥200	44.3	14.1	54.7	13.8	< 0.0		
			6		0	01		
LDL	<200	97.4	26.5	55	16.4	< 0.0		
mg/dl			5		0	01		
	≥200	148.	34.2	94.7	53.1	< 0.0		
		4	7		0	01		
TG	<200	160.	82.0	95.3	56.4	< 0.0		
mg/dl		1	1		0	01		

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	21.				
≥200	239.	155.	138.	103.	< 0.0
	4	19	3	10	01
200	9.3	2.11	7.8	1.30	< 0.0
					01
≥200	9.5	2.02	7.9	1.40	< 0.0
					01
200	43	42.6	42.4	23.1	0.88
				0	
≥200	51.7	40.7	42.3	23.0	0.08
		6		0	
	200 ≥200 200	4 2200 9.3 ≥200 9.5 2200 43	$\begin{array}{cccccccccccccccccccccccccccccccccccc$	$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	$\begin{array}{c ccccccccccccccccccccccccccccccccccc$

### Discussion:

Diabetes mellitus is more prevalence among Middle East and North Africa region with 10.9%. Moreover: it was considered to affect the largest number of adults diagnosed with DM among Western Pacific region. [18] Approximately, million and half million Iraqi have diabetes, its prevalence in Iraq ranges from 8.5% to 13.9%. According to the age-adjusted prevalence of diabetes among patients aged 19 to 94 years, a local study in Basrah, Southern Iraq, reported a prevalence up to 19.7%.[6, 7]The prevalence of dyslipidemia is high among patients with T2DM which may lead to an increased risk of CAD. When patients use statin therapy, it reduces LDL, TG, decreases the rate of VLDL synthesis, and increases HDL levels. [19] In the this study, the dose of rosuvastatin was seen to be significantly more effective in decreasing obesity and HbA1c level and improving all lipid parameters (except VLDL) after a three-month course of daily rosuvastatin 20 mg. This result is consistent with a case control study in Chinese Patients with Type 2 Diabetes Mellitus, 2019 which reported rosuvastatin to have the best effect in decreasing LDL level than other statin trials. The expected risk reduction was seen in JUPITER study.[5, 20] It was observed from the present study that the levels of HbA1c and total cholesterol means were reduced after receiving rosuvastatin to the control levels among both genders and among all age groups. This is consistent with Morieri ML, et al 2021study in North-Italy which concluded that statin treatment reduces LDL in patients with and without T2DM in different age groups. [21] In a study comparing between different statins; the reduction in TG was more marked with rosuvastatin than others and that rosuvastatin had the best effect on HDL. This effect was noticed by the VOYAGER study; 2010, a meta-analysis of 37 Randomized Controlled Trials (RCTs) which used different statin therapies.[22] Baker et al; 2010 found that rosuvastatin also but decreased insulin sensitivity statistical significance was not reached [23] while other studies reported that rosuvastatin was not associated with any change in insulin sensitivity [24, 25]. Studies done on Asian and Western populations by Nakamura H et al, 2006 reported that patients using low dose statin can decrease their risk of CVD in the same magnitude as those using high doses [26]. In addition, several RCTs reported that CVD risk was reduced more among T2DM patients who use statins than in nondiabetic patients.[27-29] Patients with type 2 diabetes can have many lipid abnormalities, elevated levels of low-density lipoprotein cholesterol and

triglycerides; and low levels of high-density lipoprotein cholesterol. Lipid abnormalities may be the result of the unbalanced metabolic state of diabetes (i.e. hyperglycaemia and insulin resistance) and improved control of hyperglycaemia does moderate diabetes-associated dyslipidaemia.[30] As a result, HbA1c may be used to predict dyslipidemia in diabetics which is similar to a study done by Hammed (2012) in Iraq which reported that early diagnosis can be reached through this inexpensive blood test. [31]

### **Conclusion:**

The study concluded that HbA1c value is associated with the level of lipid profile in diabetic patients except that for VLDL. All age groups and both genders have had benefit of rosuvastatin treatment in the reduction of lipid cholesterol as well as HbA1c, with no significant differences between groups. These results suggest that rosuvastatin can be used by type 2 diabetic patients regardless of age and gender. HbA1c can be used as a predictor of dyslipidemia in type 2 diabetes.

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# الارتباط بين الهيمو غلوبين السكري و مستويات الدهون بمصل الدم بين عينة من المرضى الارتباط بين الهيمو غلوبين المصابين بداء السكري النوع الثاني

علي مجيد حامي\* طبيب اختصاص باطنية في مستشفى الزهراء التعليمي/ دائرة صحة واسط

### الخلاصة

الخلفية: يعتبر مرض السكري من المشاكل الصحية العامة. حوالي 1.4 مليون عراقي يعانون من مرض السكري. معدل انتشار مرض السكري النوع الثاني المبلغ عنه في العراق يتراوح من 8.5٪ (معدل حسب العمر) إلى 13.9٪. كما أفادت در اسة محلية شملت أكثر من 5400 شخص في مدينة البصرة ، جنوب العراق ، عن انتشار مرض السكري بنسبة 19.7٪ حسب العمر في الأشخاص الذين تتراوح أعمار هم بين 19 و 94 عامًا. يُعد مرض السكري أحد عوامل الخطورة لأمر اض القلب والأوعية تصلب الشر ايين القلبي الوعائي وسببًا مهمًا للوفاة. كما يعد مرض السكري النوع الأكثر شيوعًا للوفاة في العالم الغربي, و قد تم اعتباره حالة عالية الخطورة في المرضى البالغين المصابين بداء السكري مع تصلب الشر ايين ، لهذا من المهم وصف العقاقير المخفضة للكوليسترول عالية الكثافة لهم لتقلبل البروتين الدهني منخض الكثافة بنسبة 50٪ على الأقل.

ا**لأهداف**: التحقيق في الارتباط بين عسر شحميات الدم والهيمو غلوُبين السكري واكتشاف فائدة استخدام بعض العقاقير المخفضة للكوليسترول في تقليل خطر الإصابة بأمراض القلب والأوعية الدموية.

المنهجية. أجريت دراسة مستقبلية غير عشوائية للجرعة المفردة في عيادة خاصة في محافظة واسط – العراق بداية شهر تشرين الاول 2020 الى نهاية شهرتموز 2021. شملت مرضى السكري من النوع الثاني. تم إجراء تقييم وإعادة تقييم تحليل كيميائي حيوي سريري قبل وبعد 3 أشهر من تلقي روزوفاستاتين 20 ملغ / يوم. تم استخدام ورقة استبيان، متضمنة السمات الاجتماعية والديموغرافية والسريرية (العمر والجنس وقياس الوزن والطول ومحيط الخصر؛ الواسمات البيوكيميائية [الكوليسترول الكلي، البروتين الدهني العالي، البروتين الدهني المنخفض، الدهون الثلاثية والهيموغلوبين السكري]].

النتائج: تم تضمين ما مجموعه 256 مريض مصاب بالسكري نوع 2؛ تلقوا 20 مجم من روز وفاستاتين كجرعة واحدة يويما لمدة 3 أشهر منتالية. 83 (32.4) منهم ذكور و 173 (67.6) إناث. متوسط عمر الذكور (25.1 ± 10.4) سنة والإناث (25.1 ± 10.4) سنة. كان متوسط فرق (مقياس كتلة الجسم، محيط الخصر، البروتين الدهني المنخفض، الدهون الثلاثية الهيمو غلوبين السكري، والكوليسترول) بين جميع العينات المدروسة بعد العلاج أقل بكثير من متوسط الفرق الذي تم قياسه قبل العلاج، باستثناء البروتين الدهني العالي؛ حيث كان أعلى بشكل ملحوظ بعد العلاج، P بعد العلاج أقل بكثير من متوسط الفرق الذي تم قياسه قبل العلاج، باستثناء البروتين الدهني العالي؛ حيث كان أعلى بشكل ملحوظ بعد العلاج، P رموناس كتلة الجسم، محيط الخصر، و 300 الذي تم قياسه قبل العلاج، باستثناء البروتين الدهني العالي؛ حيث كان أعلى بشكل ملحوظ بعد العلاج، P رموناس كتلة والعمر 40 عامًا فما فوق، 20.01 والكوليسترول الكلي بشكل ملحوظ بعد العلاج بين الذكور، والإناث، والذين تقل أعمار هم عن 40 عامًا، والعمر 40 عامًا فما فوق، 20.01 P؛ دون فروق ذات دلالة إحصائية بين المجموعات، 20.5 ح.21 في الدين يعانون من مستوى الكوليسترول الكلي 200 أو أكثر؛ لوحظ انخفاض كبير بعد العلاج بوسائل (كتلة الجسم، محيط الخصر، البروتين الدهني الدين يعانون من الثلاثية، الهيمو غلوبين السكري أو مط البروتين الدهني العالي؛ قيمة الدلالة الاحصائية المقامي الدهني الدهني المنخفض، الدهون الثلاثية، الهيمو غلوبين السكري مع ارتفاع كبير في متوسط البروتين الدهني العالي؛ قيمة الدلالة الاحصائية الحصري.

**الاستنتاج:** قيمة الهيمو غلوبين السكري مرتبطة بمستوى الدهون لدى مرضى السكري. استفادت جميع الفنات العمرية وكلا الجنسين من علاج روزوفاستاتين في تقليل نسبة الكوليسترول الدهني والهيمو غلوبين السكري. يمكن لمرضى السكر من النوع 2 استخدام روزوفاستاتين بغض النظر عن العمر والجنس. كما يمكن استخدام الهيمو غلوبين السكري كمؤشر على اضطراب شحميات الدم في مرض السكري من النوع 2.

**الكلّمات المفتاحيةً:** أمراض القلب والأوعية الدموية، داء السّكري، عسر شحميات الدم، نسبة الهيمو غلوبين السكري، البروتين الدهني عالي الكثافة، روزوفاستاتين.