

DIYALA UNIVERSITY FACULTY OF MEDICINE

CASE STUDY REPORT DYSLIPIDEMIA AMONG TYPE 2 DIABETIC PATIENTS

A Graduation Project Report
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Prepared by :
Aisha Ahmad Sabeeh

Supervised by ;

Prof. Dr. Ali Mousa J'afar

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إهداء؛

تحية طيبة ؛

أحمد الله عزَّ وجل الذي وفقني لإتمام هذا البحث العلمي ،ومتعني بالصحة والعافية ومدنى بالعزيمة ، فله الحمد حمداً طيباً كثيرا.

إلى أعز الناس وأقربهم إلى قلبي والدتي العزيزة ؛ تاج فخر طالما حملته على رأسي، فلك كامل الشكر والعرفان وإلى والدي العزيز من كان ومازال سندي ووسام عزتي و كبريائي

اللذين كانا عونا وسندا لي ، و كان لدعائهما المبارك أعظم الأثر في تسيير سفينة البحث حتى ترسو على هذه الصورة.

إلى جدتي الغالية التي كانت معي طيلة رحلتي في هذه السنوات الست وكان لها فضل كبيرً علي "

إلى عمي أثير الذي لم يتوان يوماً في مساعدتي ودعمي

إلى اخوتي فاطمة وعبدالرحمن

إلى صديقتي العزيزتين مريم عبد السلام وإيلاف يعقوب من كانتا معي في رحلة بحثي وقدمتا إلي المساعدة والدعم خلال هذه الفترة اشكركما كثيراً.

كما وأود أن أتقدم بالشبكر إلى

أستاذي ومشرفي و موجهي الفاضل الأستاذ الدكتور علي موسى: اتقدم اليه بجزيل الشكر والعرفان لاختياره لهذا الموضوع و توجيهه وتشجيعه المستمر ومتابعته للبحث حتى ظهر بأحسن صورة نرجوها

الدكتور على ثامر الذي ساعدني في إتمام هذا البحث و كان له فضل علي الدكتور/ زيد المدفعي لجهوده المبذولة في التحليل الإحصائي

أعضاء لجنة المناقشة الكرام لتفضلهم بقبول مناقشة هذه الدراسة.

كما وأتقدم بالشكر الكثير لجميع الأساتذة و الزملاء (في مجموعتي) الذين قدموا لي المساعدة مهما كانت طبيعتها، وإلى كل من قدم لي تشجيعا مهما بلغت درجته.

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ABSTRACT:

Purpose: To find out the prevalence of dyslipidemia along with its associated risk factors and comorbidities among T2DM patient in Diyala province.

Patients and Methods: The present retrospective cross sectional study based on the available biochemical data of type-2 diabetic patients

This study employed in Divala province collecting inpatients from Baguaba teaching hospital, outpatients from clinics, college and the nearby places.

Data on socio-demographic characteristics, and clinical factors were collected using a structured questionnaire through face to face interviews. This cross-sectional questionnaire was applied to 109 adults , 54 with T2DM and the other 55 are without, gathering data during the period between 1 of March 2022 and 10 of April 2022. Inclusion Criteria allowed for Adult T2DM patients (age ≥30years), Exclusion criteria for control cases and T1DM

Results: Fifty four diabetic subjects were studied and compared to fifty five non-diabetic subjects (control). The age of onset was 50.9±10.5 years (30-87 years). The duration of diabetes was <5 years in 15 (27.8%) subjects, 5-10 years were 16 (29.6%) and >10 years in 23 (42.6%). our present study indicates that the most common recognized abnormality was hypertriglyceridaemia, positive correlation between HbA1c% and total cholesterol and LDL, There was a significant increase in the presence of proteinurea in diabetic group in those with diabetic nephropathy.

Conclusion: A high significant of dyslipidemia was found among T2DM patients in the study area.

The findings of this study should be taken into account to conduct appropriate intervention measures on identified risk factor reduction and implement routine screening, treatments, and prevention of dyslipidemia.

Introduction

Diabetes mellitus is a syndrome with disordered metabolism and inappropriate hyperglycemia due to either a deficiency of insulin secretion or to a combination of insulin resistance and inadequate insulin secretion to compensate for the resistance.¹

It has many causes , most commonly type 1 or type 2 diabetes. Type 1 diabetes is generally considered to result from autoimmune destruction of insulin-producing cells (β cells) in the pancreas, leading to marked insulin deficiency, whereas type 2 diabetes is characterised by reduced sensitivity to the action of insulin and an inability to produce sufficient insulin to overcome this 'insulin resistance'. ².

Hyperglycaemia causes both acute and long-term problems. Acutely, high glucose and lack of insulin can result in marked symptoms, metabolic decompensation and hospitalisation. Chronic hyperglycaemia is responsible for diabetes-specific 'microvascular' complications affecting the eyes (retinopathy), kidneys (nephropathy) and feet (neuropathy).²

The worldwide prevalece of DM has risen dramatically over the past two

decades, from an estimated 30 million cases in 1985 to 382 million in 2013 (Fig. 1). Based on current trends, the International Diabetes Federation projects that 592 million individuals will have diabetes by



the year 2035 (see http://www.idf.org/). Although the prevalence of both type 1 and type 2 DM is increasing worldwide, the prevalence of type 2 DM is rising much more rapidly, presumably because of increasing obesity, reduced activity levels as countries become more industrialized, and the aging of the population. In 2013, the prevalence of diabetes in individuals from age 20-79 ranged from 23 to 37% in the

10 countries with the highest prevalence (Tuvalu, Federated States of Micronesia, Marshall Islands, Kiribati, Vanuatu, Cook Islands, Saudi Arabia, Nauru, Kuwait, and Qatar, in descending order of prevalence).³

Dyslipidemia:

Disorders of lipoprotein metabolism are collectively referred to as "dyslipidemias." Dyslipidemias are generally characterized clinically by increased plasma levels of cholesterol, triglycerides, or both, variably accompanied by reduced levels of HDL cholesterol. Because plasma lipids are commonly screened, dyslipidemia is frequently seen in clinical practice.

Obesity and insulin resistance are frequently accompanied by dyslipidemia characterized by elevated plasma levels of TG, low HDL-C, variable levels of LDL-C, and increased levels of small dense LDL. The increase in adipocyte mass and accompanying decreased insulin sensitivity associated with obesity have multiple effects on lipid metabolism.

In insulin-resistant patients who progress to type 2 diabetes mellitus, dyslipidemia remains common, even when the patient is under relatively good glycemic control.³

Literature Review;

Dyslipidemia is one of the risk factors for increased morbidity and mortality in diabetes. [17]

It's one of the major modifiable risk factors for cardiovascular diseases (CVD) in a type-2 diabetic (T2DM) patient. Dyslipidemia in T2DM patients is attributed due to increased free fatty acid flux secondary to insulin resistance, this will result in in an overproduction of triglyceride-rich lipoproteins from the liver, decreased clearance of triglyceride-rich lipoproteins, and, in some

cases, an altered postprandial lipoprotein metabolism. Accumulating clinical evidence has suggested serum triglyceride (TG) is a leading predictor of atherosclerotic cardiovascular disease, comparable to low-density lipoprotein (LDL)-cholesterol (C) in populations with type 2 diabetes, so early detection and treatment of dyslipidemia in type-2 diabetes mellitus can prevent risk for atherogenic cardiovascular disorder.[13]

Diabetic kidney disease (DKD) substantially worsens plasma lipid profile thereby potentiated atherogenic risk.[18]

Objective:

To determine the prevalence of dyslipidemia and its associated risk factors and complications among T2DM patients in Diyala province;

To clarify the association between glycated hemoglobin (HbA1c) and the lipid profile in patients with type 2 diabetes mellitus.

To clarify protein urea relation to lipid profile in diabetic patients

Methods and Materials: Study Design Study Area, and Ber

Study Design, Study Area, and Period

It is a retrospective cross- sectional study based on the available biochemical data of type-2 diabetic patients

This study employed in Diyala province which is located in Eastern Iraq, 57 km from the capital city, Baghdad.

The study was conducted during the period from 1 of March to the 10th of April of 2022.

collecting data from inpatients in Baquaba teaching hospital, outpatients from clinics, and the nearby places.

Data on socio-demographic characteristics, and clinical factors were collected using a structured questionnaire through face to face interviews.

This data regarding presence of DM, duration of diabetes, age, gender, residency, level of education, glycemic control, presence of complications, presence of any chronic diseases, previous measures of lipid profile and if there is evidence of dyslipidemia or not and whether it's controlled

This cross-sectional questionnaire was applied to 109 adults, 54 with T2DM and the other 55 are without, Data on HbA1c, Random blood glucose (RBG), total cholesterol (TC), triglycerides (TG), high-density lipoprotein cholesterol (HDL-C), and low-density lipoprotein cholesterol (LDL-C) levels were collected from participants along with RFT(urea & creatinine), also general urine examination for the presence of protein-urea.

Inclusion Criteria: Adult T2DM patients (age ≥30years)

Exclusion Criteria: control cases who do not suffer from any type of diabetes mellitus and T1DM patients as were five patients who had type 1 diabetes

Statistical Analysis:

Statistical analyses were performed using SPSS statistical package for Social Sciences (version 20.0 for windows, SPSS, Chicago, IL, USA). Quantitative data are represented as mean, SD, minimum and maximum. Qualitative data are expressed as number and percentage. Student's t-test was used to study the difference between diabetics and control groups.

Pearson's correlation was used to test the relation between HbA1c% with lipid profile,

Chi-square test was used to test the relation between diabetes with other factors.

P value of <0.05 was considered statistically significant.

Results:

Fifty four diabetic subjects were studied and compared to fifty five non-diabetic subjects (control). The age of onset was 50.9±10.5 years (30-87 years). The duration of diabetes was <5 years in 15 (27.8%) subjects, 5-10 years were 16 (29.6%) and >10 years in 23 (42.6%). Fifty three diabetics were controlled (98.1%) and only 1 diabetic was not controlled (1.9%).

Demographic characteristics are presented in the following tables.

Table 1: Demographic characteristics of the studied groups

| | | Group | | | | |
|-----------|---------|------------|-------|-------|-------|--|
| | | DM Control | | | | |
| | | Count | % | Count | % | |
| Age group | <=39 | 2 | 3.7% | 25 | 45.5% | |
| | 40-49 | 7 | 13.0% | 19 | 34.5% | |
| | 50-59 | 17 | 31.5% | 9 | 16.4% | |
| | 60-69 | 16 | 29.6% | 1 | 1.8% | |
| | 70-79 | 9 | 16.7% | 1 | 1.8% | |
| | >=80 | 3 | 5.6% | 0 | 0.0% | |
| Gender | Male | 27 | 50.0% | 13 | 23.6% | |
| | Female | 27 | 50.0% | 42 | 76.4% | |
| Residence | Rural | 14 | 25.9% | 6 | 10.9% | |
| | Urban | 40 | 74.1% | 49 | 89.1% | |
| Education | Unknown | 7 | 13.0% | 0 | 0.0% | |
| | High | 21 | 38.9% | 17 | 30.9% | |
| | Low | 26 | 48.1% | 38 | 69.1% | |

Table 2: Presence of diabetic complications and other chronic diseases in the studied groups

| | | | Gr | oup | |
|-----------------|-----|-------|---------|-------|--------|
| | D | М | Control | | |
| | | Count | % | Count | % |
| CVD | Yes | 29 | 53.7% | 3 | 5.5% |
| | No | 25 | 46.3% | 52 | 94.5% |
| Renal | Yes | 21 | 38.9% | 0 | 0.0% |
| | No | 33 | 61.1% | 55 | 100.0% |
| Neurological | Yes | 32 | 59.3% | 2 | 3.6% |
| | No | 22 | 40.7% | 53 | 96.4% |
| Chronic disease | Yes | 40 | 74.1% | 7 | 12.7% |
| | No | 14 | 25.9% | 48 | 87.3% |

Table 3: Comparing glucose, HBA1c, urea and creatinine between diabetic subjects and control

| | | Group | | | | | | | | |
|--------------------------------|--------|--------|--------|--------|---------|-------|-------|--------|--|--|
| | | DI | М | | Control | | | | | |
| | Mean | SD | Min | Max | Mean | SD | Min | Max | | |
| HBA1c% P=0.005 | 9.49 | 2.09 | 5.40 | 14.00 | 5.39 | .67 | 3.99 | 7.50 | | |
| RBS mg/dl P=0.005 | 278.43 | 107.89 | 106.00 | 540.00 | 99.70 | 17.08 | 71.00 | 166.00 | | |
| Urea mg/dl P=0.005 | 55.21 | 39.01 | 10.67 | 160.00 | 24.07 | 7.27 | 11.00 | 38.03 | | |
| Creatinine mg/dl P=0.005 | 1.50 | 1.20 | 0.38 | 5.00 | 0.68 | 0.21 | 0.40 | 1.30 | | |

There was a significant difference between diabetic and control groups in HBA1c, RBS, urea and creatinine (P<0.05). Student's t-test HBA1c%, RBS, urea and creatinine were higher in the diabetic group.

Table 4: Comparing lipid profile in the studied groups

| | Group | | | | | | | |
|--|--------|--------|--------|--------|---------|-------|--------|--------|
| | | D | М | | Control | | | |
| | Mean | SD | Min | Max | Mean | SD | Min | Max |
| Total cholesterol mg/dl P=0.014 | 228.24 | 80.62 | 103.50 | 420.00 | 193.46 | 63.20 | 105.00 | 475.00 |
| Triglyceride mg/dl P=0.005 | 244.38 | 115.85 | 43.20 | 525.59 | 148.48 | 99.42 | 38.00 | 496.00 |
| HDL mg/dl P=0.051 | 43.97 | 11.29 | 20.80 | 80.00 | 48.68 | 13.47 | 29.00 | 92.00 |
| LDL mg/dl P=0.075 | 133.00 | 44.73 | 41.00 | 250.00 | 117.09 | 47.62 | 49.00 | 329.00 |

There was a significant increase in diabetic than control groups for total cholesterol and TG (P<0.05); while HDL and LDL were not significantly different (P>0.05). Student's t-test HDL and LDL were not significantly different (P>0.05).

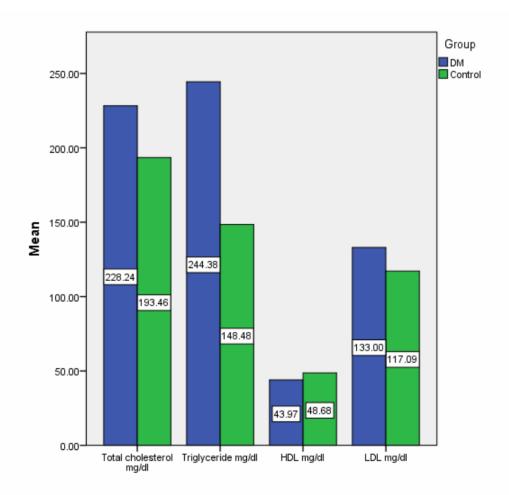


Figure 1: Lipid profile in the studied groups

Studying proteinurea, the following table show the distribution among the studied groups

Table 5: Presence of proteinurea in the studied groups

| | | | Gro | oup | Total |
|-------------|-----|-------|--------|---------|--------|
| | | | DM | Control | |
| Proteinurea | Yes | Count | 21 | 1 | 22 |
| | | % | 38.9% | 1.8% | 20.2% |
| | No | Count | 33 | 54 | 87 |
| | | % | 61.1% | 98.2% | 79.8% |
| Total | | Count | 54 | 55 | 109 |
| | | % | 100.0% | 100.0% | 100.0% |

There was a significant relation between the two groups with proteinurea being more in the diabetic group (P=0.005).

Studying the diabetic group only and the relation of proteinurea to lipid profile is presented in the following table

Table 6: Lipid profile according to the presence of proteinurea

| | | Proteinurea | | | | | | | |
|--|--------|-------------|--------|--------|--------|--------|--------|--------|--|
| | | Υ | es | | | N | 0 | | |
| | Mean | SD | Min | Max | Mean | SD | Min | Max | |
| Total cholesterol mg/dl P=0.005 | 284.55 | 88.11 | 138.00 | 420.00 | 192.41 | 50.10 | 103.50 | 290.00 | |
| Triglyceride mg/dl P=0.134 | 274.08 | 92.29 | 80.81 | 420.00 | 225.49 | 126.32 | 43.20 | 525.59 | |
| HDL mg/dl P=0.043 | 47.64 | 9.25 | 28.80 | 80.00 | 41.63 | 11.96 | 20.80 | 70.90 | |
| LDL mg/dl P=0.005 | 161.85 | 46.14 | 74.00 | 250.00 | 114.65 | 33.00 | 41.00 | 184.50 | |

There was a significant increase in the presence of proteinurea group for total cholestero, HDL and LDL (P<0.05); while TG was not significantly different (P>0.05). Student's t-test

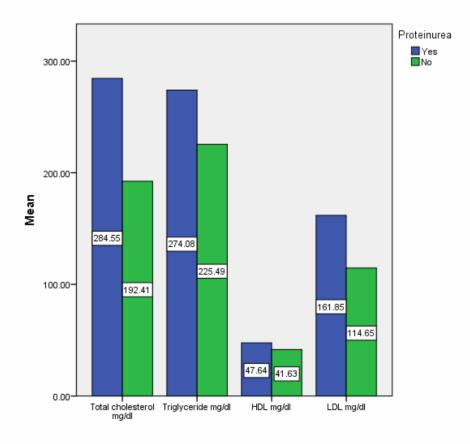


Figure 2: Lipid profile according to proteinurea

Table 7; relation between HbA1c and lipid profile In studied groups

| | | Diabetic | Control | |
|----------------|---------------------|----------|---------|--|
| | | HBA1c | | |
| T. cholesterol | Pearson Correlation | 0.484** | 0.132 | |
| | Sig. (2-tailed) | 0.005 | 0.337 | |
| | N | 54 | 55 | |
| Triglyceride | Pearson Correlation | 0.250 | 0.243 | |
| | Sig. (2-tailed) | 0.068 | 0.074 | |
| | N | 54 | 55 | |
| HDL | Pearson Correlation | -0.104 | -0.036 | |
| | Sig. (2-tailed) | 0.455 | 0.793 | |
| | N | 54 | 55 | |
| LDL | Pearson Correlation | 0.469** | 0.118 | |
| | Sig. (2-tailed) | 0.005 | 0.391 | |
| | N | 54 | 55 | |

Pearson's correlation test

In the diabetic group there was a positive correlation between HbA1c% and total cholesterol and LDL (P<0.05). No significant correlation was found with TG and HDL (P>0.05).

No significant correlation was found in the control group.

Discussion:

The relation between diabetes mellitus and serum lipid profile had been discussed a lot during the past decades [5,6,7,8]. Both lipid profile and diabetes have been shown to be the important predictors for metabolic disturbances including dyslipidaemia, hypertension, cardiovascular diseases, hyperinsulinemia, etc. [9]. Dyslipidemia as a metabolic abnormality is frequently associated with diabetes mellitus. Its prevalence is variable, depending on the type and severity of diabetes, glycaemic control, nutritional status, age and other factors. Earlier studies also indicated a strong clustering risk factor for coronary artery disease in diabetic subjects [5,10]. Over 70% of patients with type 2 diabetes mellitus had one or more types of dyslipidemia. Similarly, our study results reveal high prevalence of hypercholesterolemia and hypertriglyceridemia, which are well known risk factors for cardiovascular diseases among patients.

In diabetes many factors may affect blood lipid levels, because of interrelationship between carbohydrates and lipid metabolism. Therefore, any disorder in carbohydrate metabolism leads to disorder in lipid metabolism and vice versa [11]. Insulin resistance is a primary defects in the majority of with type-2 diabetes.

There are several studies showed that the insulin affects the liver apolipoprotein production and regulates the enzymatic activity of lipoprotein lipase and cholesterol ester transport protein, which causes dyslipidemia in diabetes mellitus.

The current study attempted to assess the prevalence of dyslipidemia and associated risk factors among T2DM patients in Diyala.

Socio-demographic factors can play role in determining dyslipidemia in diabetic patients.In the current study, dyslipidemia was significantly associated with increasing age (age≥ 50), also increased in urban area According to the findings of our study indicates that the most common recognized abnormality was hypertriglyceridaemia with mean 244.28± 115.85 followed by hypercholesterolemia with mean 228.24±80.62 [Table4]in second place and these articles support our finding respectively [4,12].

While in control group the mean of triglyceride was 148.48±99.42, total cholesterol 193.64±63.2 with P value <0.05 which make it significantly different.

We found that there is no significant difference of HDL and LDL between studied group with mean 43.97±11.29, 133±44.73 in diabetes group and 48.13±13.47, 117.09±47.62 in control group respectively, with P value>0.05 and this study disagrees with ours [14].

In the other hand other studies revealed Hypertriglyceridaemia and low HDL level was the most frequent lipid abnormality found in this study.^[13]

In the present study we found a significant difference of presence of proteinurea between the studied group, diabetic patient showed higher percentage of proteinurea than the control subjects 38.9%, 1.8% respectively [Table 5]

There was a significant increase in the presence of proteinurea group for total cholestero, HDL and LDL (P<0.05); while TG was not significantly different (P>0.05). [Table 6], Figure 2. Our results show that dyslipidemia was highly prevalent among subjects with nephropathy. We couldn't found a previous study consistent with our findings.

This finding is inconsistent with previous studies from^[19]as its shows there's a relation of protein urea with elevated triglyceride level.

In diabetic group there was a positive correlation between HbA1c% and total cholesterol and LDL (P<0.05). This previous study in Iraq in 2021 support our findings^[15].

No significant correlation was found in the control group.

Conclusion; A high prevalence of dyslipidemia was found among T2DM patients in the study area, HbA1c value associated with level of lipid profile in diabetic patients.

Also there was a significant protein urea relation to lipid profile in diabetic patients.

Recommendation:

The American Diabetes Association 2019 guidelines recommend that all diabetic patients with ASCVD or patients with a 10-year atherosclerotic cardiovascular risk > 20% should be treated with high-intensity statins (goal of 50% reduction in LDL-cholesterol) in addition to lifestyle modification^[16].

In our study we We faced difficulties in performing the investigation of lipid profile, general urine examination from inpatient in hospital as they are not routine investigations for T2DM patient, so we recommended that they should be routinely investigated in order to decrease its associated complications and comborbidities and to facilitate the collection of data in the future.

The findings of this study should be taken into account to conduct appropriate intervention measures on identified risk factor reduction and implement routine screening, treatments, and prevention of dyslipidemia.

ASCVD= Atherosclerotic cardiovascular disease

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الخلاصة:

الهدف من الدراسة: التعرف على نسبة الإصابة بعسر او اضطراب دهون الدم وما يرتبط به من عوامل الخطر والأمراض المصاحبة بين مرضى السكر النوع الثاني في محافظة ديالي.

المرضى والطرق: الدراسة المقطعية الحالية بأثر رجعي بناءً على البيانات البيوكيميائية المتاحة لمرضى السكري من النوع الثاني.

تمت هذه الدراسة في محافظة ديالى وقد تم جمع البيانات من المرضى المنومين من مستشفى بعقوبة التعليمي والمرضى من العيادات والأماكن القريبة.

تم جمع البيانات حول الخصائص الاجتماعية والديموغرافية والعوامل السريرية باستخدام استبيان منظم من خلال المقابلات وجهاً لوجه. وقد تم تطبيق هذا الاستبيان المقطعي على 109 من البالغين ، 54 منهم مصابين بالسكري النوع الثاني، بينما الخمس والخمسون الاخرون لا يعانون من السكري ، وتم جمع هذه البيانات خلال الفترة بين 1 مارس 2022 و 10 أبريل 2022.

وقد كانت معايير التضمين المسموح بها لمرضى السكري النوع الثاني البالغين من (العمر ثلاثين سنةً فما فوق) ، أمًا معايير الاستبعاد فقد كانت للحالات التي لا تعاني من السكري في الاصل ومن يعاني من السكري النوع الاول.

النتائج: تمت دراسة أربعة وخمسين شخصًا مصابًا بمرض السكري ومقارنتهم بخمسة وخمسين شخصًا غير مصابين بمرض السكر (مجموعة التحكم). كان عمر البدء 50.9 ± 10.5 سنة (30-87 سنة). كانت مدة مرض السكري أقل من 5 سنوات في 15 (27.8٪) ، 5-10 سنوات كانت 16 (29.6٪) وأكثر من 10 سنوات في 23 (42.6٪). تشير دراستنا الحالية إلى أن الاضطراب الأكثر شيوعًا هو ارتفاع دهون الدم الثلاثية، وأن هناك علاقة إيجابية بين اختبار الهيمو غلوبين الغليكوزيلاتي (HbA1c ٪) والكوليسترول الكلي والبروتين الدهني منخفض الكثافة LDL ، وكانت هناك زيادة كبيرة في وجود بروتينية في مجموعة مرضى السكري الذين يعانون من اعتلال الكلية السكري.

كما وكانت هذه الزيادة في وجود مجموعة البروتينية للكوليسترول الكلي ، HDL و LDL.

الخاتمة: تم العثور على نسبة عالية من اضطراب أو عسر دهون الدم بين مرضى السكري الثاني في منطقة الدراسة ويجب أن تؤخذ نتائج هذه الدراسة في عين الاعتبار لإجراء تدابير التدخل المناسبة بشأن الحد من عوامل الخطر المحددة وتنفيذ الفحص الروتيني والعلاجات والوقاية من اضطراب دهون الدم قدر الامكان