ORIGINAL ARTICLE

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Estimation of magnesium level in type 2 diabetes mellitus and its correlation with HbA1c level

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Summarv

Background: Type 2 diabetes mellitus (T2DM) is a common disease; the total number of diabetes patients is expected to reach 366 million by 2030. Magnesium has received considerable attention for its potential role in improving insulin sensitivity and preventing diabetes and its cardiovascular complications. Hypomagnesaemia is linked to poor control of type 2 diabetes mellitus, and depletion of serum magnesium occurs exponentially with duration of disease. The aim of this study was to measure serum magnesium level and the correlation of magnesium level with HbA1c in type 2 diabetes mellitus.

Methods: In this cross-sectional study, we included 100 diabetic patients; blood is taken for measurement of both magnesium and HbA1c levels. Patients with risk factors of magnesium deficiency were not included.

Results: Of the total 100 patients recruited in our study, majority had diabetes for 8.0 years duration. The mean age of the patients was 53.97 ± 10.65 years, most of them were on oral hypoglycaemic agents (84.9%) followed by combination of oral hypoglycaemic agents and insulin (10.5%), and a small percentage were on insulin alone (4.7%). The mean of serum magnesium and HbA1c levels was 1.88 ± 0.25 mg/ dL and 8.38% ± 1.53%, respectively. The majority of the patients had a normal level of magnesium (95.0%); however, most of them had uncontrolled blood glucose (82.0%). The study showed that the serum magnesium level and HbA1c are not significantly correlated (P = 0.462).

Conclusion: Normal magnesium levels were observed in majority of patients, there is no significant correlation between serum magnesium and HbA1c levels in patients with T2DM, but larger-scale clinical trials are needed in future.

KEYWORDS HbA1c, magnesium, type 2 DM

1 | INTRODUCTION

Type 2 diabetes mellitus (T2DM) is a chronic disease resulting from a complex inheritance-environmental interaction along with other risk factors such as obesity and a sedentary lifestyle.¹ The total number of diabetes is expected to reach 366 million by 2030.² A high

prevalence rate has been observed in developing countries and in populations undergoing "westernization" or modernization.¹ In Iraq, the prevalence increased from 5% in 1978 to 19.7% in 2012.³

In spite of innovative methods in the management of diabetes mellitus, the morbidity and mortality continue to be high.⁴ The deleterious effect of diabetes in these patients is related to complications,

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including both macrovascular and microvascular complications.^{5,6} Diet is widely believed to play an important role in the development of type 2 diabetes (T2D) and the associated complications.^{5,6}

The glycated haemoglobin (HbA1c) is widely used for evaluation of diabetes control.⁷ The HbA1c reveals the overall blood glucose levels over a period of 2-3 months, and the common use of the HbA1c assay is to assess changes in metabolic control that follow an alteration in treatment.⁸

Magnesium (Mg) is one of the important components of many foods such as grains, nuts and green leafy vegetables, and it plays a key role in many fundamental biological processes, including energy metabolism. Mg has received considerable attention for its potential role in improving insulin sensitivity and preventing diabetes and its cardiovascular complications.^{9,10} It is claimed that Mg deficiency is common in diabetic patients and there is an inverse relationship between Mg intake and incidence of T2DM.¹³

It was founded that poor glycaemic control, insulin resistance and low Mg level were strongly associated with increased the prevalence of microalbuminuria.¹⁴ Hypomagnesaemia is linked to poor control of type 2 diabetes mellitus, and depletion of serum magnesium occurs exponentially with duration of disease.¹⁵ It is now established that diabetes can by itself induce hypomagnesaemia which in turn induces or worsens diabetes mellitus.¹⁶

Although the assessment of serum magnesium level has been reported in many studies from various countries, the comparative studies within the diabetic population based on glycaemic control are limited. For this reason, we decided to study the level of serum magnesium in T2DM.

1.1 | Aim of the study

The aim of this study was to find the correlation between magnesium and HbA1c levels in type 2 diabetes mellitus.

2 | METHODS

2.1 | Study design and sampling

This is a cross-sectional study, which included 100 patients with type 2 DM, each week approximately 5-7 patients were selected randomly, aged between 30 and 70 during the period of 6 months (January 2018 to June 2018).

The study conducted at Azadi Teaching Hospital at the department of internal medicine in cooperation with Duhok General Directorate of Health. All patients' participating in this study signed informed consent to have their data collected and used for research. This study was approved by the Duhok General Directorate of Health Ethics Committee.

2.1.1 | Inclusion criteria

All cases of type 2 diabetes mellitus without comorbidities and the age of patients was between 30 and 70 attending Azadi Teaching Hospital in Duhok.

2.1.2 | Exclusion criteria

Patients with renal failure, patients who suffered from acute myocardial infarction in last six months, patients on diuretics, history of alcohol abuse, magnesium supplements/magnesium-containing antacids, malabsorption and patients with chronic diarrhoea.

2.2 | Blood sampling

After an overnight fasting, 5 mL of venous blood was collected by standard procedure without tourniquet from each patient under complete aseptic conditions; 2.5 mL was placed in lithium heparin test tube. Serum was obtained by low-speed centrifugation, the separation was done without undue delay, and after separation, the serum samples were stored at 2-8°C and then used directly for magnesium analysis. Magnesium has been measured by Cobas 6000 (Roche, Mannheim, Germany) using colorimetric end-point method. The other 2.5 mL was placed in EDTA test tube and used for HbA1c analysis. HbA1c was measured by enzyme-linked immunosorbent assay (ELISA) test. The Statistical Package for Social Sciences (SPSS) was used for data analysis, and data were presented as mean ± standard deviation. Comparisons in variables were analysed using independent *t* test and Pearson correlation. *P*-values were regarded significant when ≤0.05.

3 | RESULTS

Of the total 100 patients recruited in our study, more than half of them were females with the male: female ratio of 0.54:1.00 and most of them had diabetes for 8.0 years (75%), while 35% of study population has diabetes of 3-5 years. The mean age of the patients was 53.97 ± 10.65 years and the majority of them were on oral hypoglycaemic agents (84.9%), followed by the combination of oral hypoglycaemic agents and insulin (10.5%), and a small percentage were on insulin alone (4.7%). The mean of serum magnesium level and HbA1c was 1.88 ± 0.25 mg/dL and $8.38\% \pm 1.53\%$, respectively. The majority of the patients had a normal level of magnesium (95.0%); however, most of them had uncontrolled

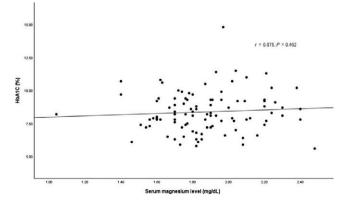


FIGURE 1 Correlation of serum magnesium level with HbA1c in type 2 diabetes patients (the correlation was not significant, P = 0.462)

TA	ΒL	E	1	General	characteristics	of stud	y patients
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Patients' characteristic	Frequency distribution			
(n = 96)	Mean	Sta. deviation		
Age (y)	53.97	10.65		
Serum magnesium level (mg/ dL)	1.88	0.25		
HbA1c (%)	8.38	1.53		
Disease duration (y) ^a	8.0	7.25		
	Frequency	Percentage		
Gender				
Male	35	35.0		
Female	65	65.0		
Treatment type				
Insulin	4	4.7		
Oral hypoglycaemic agents (OHA)	73	84.9		
Combined (insulin plus OHA)	9	10.5		
History of myocardial infarction	1	1.0		
HbA1c categories				
Controlled (HbA1c <7)	18	18.0		
Uncontrolled (HbA1c ≥7)	82	82.0		
Serum magnesium ranges				
Normal (1.5-2.5 mg/dL)	95	95.0		
Low Mg Level	5	5.0		

^aMedian and interquartile range were calculated for disease duration.

blood glucose (82.0%), as shown in Table 1. The study showed that the serum magnesium level and HbA1c are not significantly correlated with each other (P = 0.462) as shown in Figure 1.

The study did not show a significant difference in serum magnesium level between the patients with optimal and uncontrolled glucose level (P = 0.866). The serum magnesium level of both groups was in the normal range, as shown in Table 2.

4 | DISCUSSION

Many studies have been done to estimate the serum magnesium level in T2DM patients and its correlation with HbA1c, but in Duhok, very few data are available regarding the serum magnesium level in diabetic patients, and to the best of our knowledge, this study is the first to evaluate the correlation of magnesium level with HbA1c in type 2 diabetes mellitus in Duhok. Sex distribution in Table 1 represents that the ratio of females is greater than males which are agreed with many studies reported in different countries.^{15,16}

The magnesium is an essential cofactor of more than 300 enzymes including those important in glycolysis, neuromuscular transmission, synthesis of carbohydrates, proteins, lipid and nucleic acids, and it has a role in insulin's secretion, its binding and its activity.^{11,13,18}

TABLE 2 Difference of serum magnesium level in patients with optimal and uncontrolled glucose level

	Serum ma	agnesium level	P-value	
HbA1c	Mean	Sta. deviation	(two-sided)	
Optimal HbA1c	1.88	0.23	0.866ª	
Uncontrolled HbA1c	1.87	0.26		

^aIndependent *t* test was performed for statistical analysis.

The serum magnesium level in the present study was normal in 95% of cases, and our results were comparable with other studies, such as the results of Masood et al¹⁹ and Walter et al²⁰ whose reported the normal Mg level in both T2DM patients and healthy controls. But Wälti et al²¹ showed that serum Mg concentrations of 37.6% of the diabetics were below the reference range and the findings of many studies which were reported low magnesium serum levels in T2DM.²²

The HbA1c levels were uncontrolled in about 82% of patients enrolled in this study, and our results were in agreement with many studies which reported high HbA1c levels in T2DM.^{23,24}

Our results showed no correlation between serum Mg level and HbA1c level in contrast to many studies reported that there is an inverse correlation between serum Mg level and HbA1c,²⁵ some studies showed normal Mg levels in T2DM but lack data correlation to HbA1c.

The nonsignificant correlation between serum Mg level and HbA1c of this study may be due to our exclusion of patients on diuretic and patients with renal failure or may be due to dietary intake of our population. Our results were in contrast to that reported by Siddique et al²⁶ who founded that hypomagnesaemia is associated with higher level of HbA1c. Also, Tilal et al showed that there is a strong negative correlation between the serum level of magnesium and HbA1c level. Meanwhile, Navarrete-Cortes (2016) revealed that the magnesium supplementation does not improve HbA1c or insulin sensitivity in diabetic subjects with normomagnesaemia.²⁷

4.1 | Limitation of study

Being cross-sectional study and small sample size are main limitations. Larger-scale clinical trials are needed in future to determine whether there is a correlation between serum Mg level and HbA1c level in T2DM.

5 | CONCLUSION

Normal magnesium levels were observed in about 95% of patients with T2DM, and HbA1c levels were uncontrolled in 82% of patients which enrolled in this study. There is no significant correlation between serum Mg level and HbA1c level in patients with T2DM.

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ETHICAL APPROVAL

All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional and/or national research committee and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards.

CONFLICT OF INTEREST

Authors declare that they have no conflict of interest.

AUTHOR CONTRIBUTIONS

All authors have equally contributed in this study, and they all agreed on the final manuscript.

DATA ACCESSIBILITY

None.

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