A Study of Serum Magnesium Level in Type 2 Diabetes Mellitus Patients

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Abstract

Background: Type 2 Diabetes Mellitus (DM) is characterized by insulin resistance in peripheral tissues together with impaired secretion of insulin. DM is the most common metabolic disorder with magnesium deficiency, having 25% to 39% prevalence. Osmotic diuresis accounts for a portion of the magnesium loss. Plasma magnesium and intracellular magnesium concentrations are tightly regulated by several factors. Insulin is the most important factor. Insulin may modulate the shift of magnesium from extracellular to intracellular space. In the present study we try to estimate serum magnesium in type 2 DM and also to find the association with duration of T2DM. Materials and Methods: The study will be conducted in Thanjavur Medical College, Thanjavur. 25 diabetes patients and 25 healthy volunteers were included in the present study. Age group of the study subjects were between 35 and 67 years. Approximately 3 ml of fasting serum samples and 1 ml of whole blood samples will be collected in Clinical Chemistry laboratory for the estimation of fasting glucose (GOD-POD method), 2-hour post glucose levels and HbA1c (HPLC method), Lipids profile was estimated by IFCC approved clinical chemistry analyzer (fully automated). Results: There was statistically significant difference in BMI, fasting blood sugar (FBS), Post Prandial Blood Sugar (PPBS), lipid profile and Mg (in DM groups). HbA1c ($P < 0.001$) levels were statistically significance with DM patients. Serum Mg levels were correlated showed negative correlation except High Density Lipoprotein (HDL) had positive correlation. The Serum Mg levels are significantly decreased along with the duration of the diabetes. Conclusion: Hypomagnesemia is seen in type 2 DM and inverse correlation with duration of the diabetes also. Supplementation of Magnesium can prevent chronic complications related to diabetes mellitus.
1. Introduction

Type 2 diabetes mellitus (type 2 DM) is a non-autoimmune, complex, heterogeneous and polygenic metabolic disease in which body fails to produce enough insulin, characterized by abnormal glucose homeostasis [1]. It is the predominant form of Diabetes, accounting for 90% cases globally, is a worldwide health crisis, WHO predicts an incidence of 300 million by 2025 [2]. As per International Diabetes Federation (IDF) the number of diabetes is said to raise from 40.9 million to 69.9 million by the year 2025 in India. Type 2 DM is characterized by insulin resistance in peripheral tissues together with impaired secretion of insulin. DM is the most common metabolic disorder with magnesium deficiency, having 25% to 39% prevalence. Osmotic diuresis accounts for a portion of the magnesium loss [3].

Pathophysiology of Type 2 DM is due to 1) resistance to the action of insulin in peripheral tissues particularly muscle and fat but also liver 2) defective insulin secretion, particularly in response to a glucose stimulus 3) increased glucose production by the liver. Glucose itself is a crucial part of cellular ion homeostasis, increasing intracellular calcium and decreasing intracellular magnesium [4]. Prevalence of hypomagnesemia in DM is about 65% [5].

Magnesium is the fourth abundant mineral in our body and most abundant intracellular cation [6]. Magnesium is involved in many carbohydrate oxidation, enzymatic reactions, glucose transport mechanism, insulin secretion and in binding activity [7] [8]. Approximately 50% of total body magnesium is present in bones. Other 50% is found predominantly inside the cells of tissues and organs. Only 1% of magnesium is found in blood. The homeostasis of magnesium depends upon the amount of ingestion, efficiency of absorption and excretion from intestine and kidney.

Magnesium acts as a cofactor in more than 320 enzymatic reactions involving energy metabolism and nucleic acid synthesis. It is involved in several processes including hormone receptor binding and gating of calcium channels, transmembrane ion flux, regulation of adenylate cyclase, muscle contraction and neuronal activity, control of vascular tone, cardiac excitability and neurotransmitter release. Magnesium increases the body’s ability to utilize sodium, potassium, calcium, phosphorus, vitamin B complex, vitamin C, E. Intracellular magnesium is located within the mitochondria apparently because magnesium binds strongly with ATP. More metabolically active cell is the higher in its magnesium content. Plasma magnesium and intracellular magnesium concentrations are tightly regulated by several factors. Insulin is the most important factor. Insulin may modulate the shift of magnesium from extracellular to intracellular space.
In the present study we try to estimate serum magnesium in type 2 DM and also to find the association with duration of T2DM.

2. Materials and Methods

The present study was conducted at Thanjavur Medical College Hospital, Thanjavur in the 2013. 50 clinically diagnosed Type 2 DM patients (25 males and 25 females) were included in study group and 50 healthy individuals were included in the control group (25 males and 25 females) with Age ranged from 35 - 67 Years. Sample size of the study was selected based on the prevalence of Type 2 DM. Type 2 DM with complications, chronic kidney disease, cardiovascular diseases, drugs like (thiazide diuretics, steroids) alcoholics, and diarrhea were excluded from the study.

3 ml of fasting and 2 ml of postprandial venous blood sample taken under aseptic precaution. Fasting and postprandial blood glucose, lipid profile (Total Cholesterol, Triglyceride, HDL, LDL-calculated and VLDL-calculated) were estimated by using IFCC approved method. Serum magnesium is estimated by spectrophotometric method using xylidyl blue 1 [9]. Xylidyl blue 1 forms a red complex in alkaline solution with magnesium. The absorbance at 520 nm of there xylidyl blue 1 magnesium complex is proportional to the concentration of magnesium in the sample. Informed consents were obtained from all patients and healthy volunteers before they entered the study. Student t test done to compare BMI, fasting blood glucose and postprandial glucose levels between control and study groups. Pearson coefficient correlation between Serum Magnesium and lipid profile. One way ANOVA was performed duration T2DM and serum magnesium in study groups.

3. Results

The primary objective of the study was to compare the levels of serum Mg between T2DM and control groups, for that we had recurred 50 T2DM patients and 50 healthy volunteers after appropriate screening. Analysis of the report showed that there was a statistically significant decrease in the level of Mg in T2DM when compared with control group ("p" < 0.001) which is depicted in Table 1. Other anthropometric and biochemical parameter like BMI, FBS, PPBS and Lipid profile also presented with statistical significance.

Secondary objective was to find the association of Mg levels with biochemical parameter and duration of the T2DM. As in the literature we also found that there is negative correlation between Mg levels and all biochemical parameter, except HDL which showed a positive correlation with serum Mg levels in T2DM as in Table 2. Duration of T2DM and serum Mg levels had negative correlation, as year progress serum magnesium level come down Table 3 and Figure 1.

4. Discussion

Magnesium is an intracellular cation plays a role in cellular metabolism. Hypomagnesemia in DM has been reported in many studies [10] [11] [12]. In our
**Table 1.** Mean bmi, biochemical parameter of control and study groups.

<table>
<thead>
<tr>
<th>Variables</th>
<th>Control</th>
<th>Study</th>
<th>Statistical inference</th>
</tr>
</thead>
<tbody>
<tr>
<td>BMI</td>
<td>22.09 ± 1.48</td>
<td>25.49 ± 1.49</td>
<td>p &lt; 0.000*</td>
</tr>
<tr>
<td>S. FBG(mg/dl)</td>
<td>92.92 ± 10.30</td>
<td>132.64 ± 17.51</td>
<td>p &lt; 0.000*</td>
</tr>
<tr>
<td>S.PPBG(mg/dl)</td>
<td>125.20 ± 6.09</td>
<td>244.10 ± 69.02</td>
<td>p &lt; 0.000*</td>
</tr>
<tr>
<td>S.Mg (mg/dl)</td>
<td>2.10 ± 0.21</td>
<td>1.61 ± 0.37</td>
<td>p &lt; 0.000*</td>
</tr>
</tbody>
</table>

*P < 0.05 is statistically significant.

**Table 2.** Pearson correlation between S. magnesium, S. Fbg, S. Ppbg, In study group.

<table>
<thead>
<tr>
<th>PARAMETERS</th>
<th>&quot;r&quot; value</th>
</tr>
</thead>
<tbody>
<tr>
<td>FBG (mg/dl)</td>
<td>−0.6190*</td>
</tr>
<tr>
<td>PPBG (mg/dl)</td>
<td>−0.7379*</td>
</tr>
<tr>
<td>TC (mg/dl)</td>
<td>−0.6114*</td>
</tr>
<tr>
<td>MAGNESIUM (mg/dl) Vs TGL (mg/dl)</td>
<td>−0.5363*</td>
</tr>
<tr>
<td>MAGNESIUM (mg/dl) Vs HDL (mg/dl)</td>
<td>0.6660*</td>
</tr>
<tr>
<td>MAGNESIUM (mg/dl) Vs VLDL (mg/dl)</td>
<td>−0.5363*</td>
</tr>
<tr>
<td>MAGNESIUM (mg/dl) Vs LDL U/L</td>
<td>−0.6243*</td>
</tr>
</tbody>
</table>

**Table 3.** Duration wise serum mean magnesium levels in study group.

<table>
<thead>
<tr>
<th>Duration (yrs)</th>
<th>Mean</th>
<th>S.D</th>
<th>&quot;p&quot; value</th>
</tr>
</thead>
<tbody>
<tr>
<td>&lt;3 yrs (n = 10)</td>
<td>1.5360</td>
<td>0.1670</td>
<td>p = 0.0001*</td>
</tr>
<tr>
<td>4 - 6 yrs (n = 21)</td>
<td>1.2621</td>
<td>0.2846</td>
<td></td>
</tr>
<tr>
<td>7 - 9 yrs (n = 10)</td>
<td>1.044</td>
<td>0.1194</td>
<td></td>
</tr>
<tr>
<td>&gt;10 yrs (n = 9)</td>
<td>1.015</td>
<td>0.1208</td>
<td></td>
</tr>
</tbody>
</table>

*P < 0.05 is statistically significant.

**Figure 1.** Mean serum magnesium levels in relation to duration of type 2 dm.
study there is significant hypomagnesemia in Type 2 DM \( (p = 0.0001 < 0.05) \). Similar findings have been reported in studies done in serum magnesium status in Type 2 DM. \[13\] \[14\] There is negative correlation between serum magnesium levels with fasting and postprandial blood glucose level. This has been reported in a study done by Diwan, et al., Tripath, et al., Nila Masood, et al. Diabetes Mellitus is one of most common causes of magnesium deficiency although is related to poor metabolic control \[15\].

In type 2 DM patients, hypomagnesemia can be a consequence or a cause of increased insulin resistance. The reason for high prevalence may be attributed to increased urinary loss (osmotic diuresis), low dietary intake or impaired absorption of magnesium compared to healthy individuals \[16\]. Intracellular magnesium plays a key role in regulating insulin action, insulin-mediated glucose uptake and vascular tone Reduced intracellular magnesium concentrations result in defective tyrosine kinase activity, impede phosphorus bond dependent reactions of the many enzymes related to glucose metabolism, impair insulin receptor function through increased microviscosity of the plasma membrane and thus decrease insulin sensitivity and worsening of insulin resistance in diabetic patients \[17\].

Hypomagnesemia is associated with low high density lipoprotein, and increased triglyceride, very low density lipoprotein and low density lipoprotein levels. Insulin can increase free magnesium entry into the cell in insulin resistance, insulin-induced entry of magnesium is impaired \[18\]. The influence of magnesium on cell membrane ATPase activity and consequently on intracellular sodium, calcium, potassium metabolism may also play a role in diabetic complication. Hypomagnesemia when it is chronic, it increases the risk of macro and microvascular complications of DM \[19\]. Low levels of magnesium promote endothelial cell dysfunction and thrombogenesis by increasing platelet aggregation and vascular complication. Magnesium has been shown to inhibit platelet activation by inhibiting thromboxane A2 & interfering with the IIb-IIa receptor complex formation \[20\]. The influence of magnesium on cell membrane ATPase activity, consequently on intracellular Ca\(^{2+}\), Na\(^+\), K\(^+\) metabolism may also play a role in diabetic complications. Hypomagnesemia when it is chronic, increases the risk of macro and microvascular complications of DM \[15\] \[19\].

Magnesium exists only in very small amounts in blood and serum magnesium is not always precise a indicator of intracellular magnesium content, intracellular concentration measurement would have enabled better assessment of magnesium status in diabetes mellitus.

5. Conclusion

To conclude, Hypomagnesemia is seen in type 2 DM and inverse correlation between magnesium level and fasting and postprandial blood glucose levels. Periodic monitoring of Magnesium concentration and Magnesium supplementation can prevent chronic complications related to diabetes mellitus.
Limitations of the Study

Along with Magnesium, uric acid and thyroid profile estimations can be included for the study in future.

References


