

Comparative Study of Serum Magnesium, Calcium, Potassium and Sodium Levels in Diabetics and Hypertensives with Acute Myocardial Infarction

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Abstract:

Introduction: The importance of extracellular and intracellular magnesium has become gradually recognized during the last century. At the present moment, pathologies as common as diabetes, hypertension and dyslipidaemia are associated with an altered metabolism of magnesium.

Aim: Study included diabetics and hypertensives with Acute Myocardial Infarction to compare the levels of cations in these subgroups and also to find out any association of cation disturbances with these metabolic disorders.

Materials & Methods: Study was conducted in 75 patients (n=75) admitted to the intensive coronary care unit (ICCU) with chest pain and provisional diagnosis of Acute Myocardial Infarction (AMI). All the 75 patients were studied serially on 1st day of admission, 3rd day and 7th day after AMI. Patients without serious disease who were seen in the outpatient departments and healthy volunteers were taken as controls (n=30).

Results: There is significant decrease in Serum Magnesium and Serum Potassium levels in hypertensives and Diabetics on 1st day of AMI when compared to controls ($p < 0.01$) with no significant change in $S.Ca^{+2}$ and $S.Na^{+}$ levels ($p > 0.05$). There is significant decrease in Serum Magnesium levels in cases with diabetes ($p < 0.05$) when compared to cases without diabetes. There is lowering of Serum Magnesium levels in cases with Hypertension when compared to cases with out hypertension , but there is no statistical significance. There is no statistically significant change in other electrolytes in cases with and without diabetes and in cases with and without hypertension.

Conclusion: There is significant decrease in Serum Magnesium levels in Diabetics with AMI when compared to cases without diabetes and lowering of $S.Mg^{+2}$ level in hypertensive cases with AMI, with no significant change in $S.Ca^{+2}$ and $S.Na^{+}$ levels. Low Magnesium Levels may be a cause or effect of Diabetes

Keywords: AMI (Acute Myocardial Infarction), Diabetes Mellitus, Magnesium.

I. Introduction

Study included diabetics and hypertensives with Acute Myocardial Infarction to compare the levels of cations in these subgroups and also to find out any association of cation disturbances with these metabolic disorders. The importance of extracellular and intracellular magnesium has become gradually recognized during the last century. At the present moment, pathologies as common as diabetes, hypertension and dyslipidaemia are associated with an altered metabolism of magnesium, and this divalent cation is even being considered as a potential tool in the prevention or co-adjuvant treatment of ventricular arrhythmias, coronary heart disease and cirrhosis of the liver, among others.¹

Habitually low intakes of magnesium and resulting abnormal magnesium metabolism are associated with etiologic factors in various metabolic diseases, in particular: cardiovascular; blood pressure; skeletal growth & osteoporosis; and diabetes mellitus.² Magnesium is essential for potassium transport. Evidence suggests that a deficit of magnesium is closely interrelated to potassium deficiency and refractory potassium repletion. Numerous experiments and clinical observations have credited magnesium with a positive influence on the incidence of migraine attacks.³ Reduced erythrocyte magnesium (Mg) levels have been reported in the chronic pain syndromes: fibromyalgia syndrome (FS), chronic fatigue syndrome (CFS), myofascial pain syndrome (MPS) eosinophilia myalgia syndrome (EMS) and systemic lupus erythematosus (SLE). Calcium interacts with sodium, potassium and magnesium to help regulate blood pressure. Sodium helps regulate blood pressure and water balance in cells. Helps maintain acid-base balance and aids in muscle contraction and nerve impulse transmission.⁴

II. Materials and Methods

Study was conducted in 75 patients (n=75) admitted to the intensive coronary care unit (ICCU) of department of Cardiology and Acute medical care unit department of Medicine, Gandhi Hospital, with chest

pain and provisional diagnosis of Acute Myocardial Infarction (AMI). All were electrocardiographically proved cases of AMI ranging in age from 32-76 years with mean age of 54.1 ±11.2 (Males-45 and Females-30).

Patients without serious disease who were seen in the outpatient departments and healthy volunteers were taken as controls (n=30) with mean age of 49±9.9 (Females-14, Males-16). Diabetics were excluded and no control subject has a history or electrocardiographic evidence of ischemic or rheumatic heart disease, hypertension, congestive heart failure or renal failure. None was receiving any sort of medication.

Venous blood was collected under aseptic precautions after consent is taken and RBS, Blood Urea and Serum Creatinine were estimated to rule out diabetes and renal pathology among controls. Serum Magnesium, Calcium, Potassium and Sodium were estimated in both control and study groups. S. Magnesium was estimated by calmagite kit method.⁵ S.calcium was estimated by O-cresolphalein kit method⁶ and S. electrolytes were estimated by flame photometry method⁷

The following data were recorded in study group- age, sex, medication at the time of admission, treatment for hypertension and heart failure was registered. All patients were monitored with continuous electrocardiogram (ECG) registration. All the 75 patients were studied serially on 1st day of admission, 3rd day and 7th day after AMI. Written consent was taken from each patient for participating in the study. Five ml of venous blood was collected on each occasion in plain sterile bottle and allowed to clot. Serum was separated immediately for estimation of serum creatinine, serum magnesium, calcium and electrolytes. Two ml of blood was taken in oxalate fluoride bottles on 1st day for estimation of Blood sugar and Blood urea. Random blood sugar was measured by King and Asatoor method. Blood urea was measured by Di acetyl monoxime (DAM) colorimetric method and serum creatinine by Jaffes alkaline picrate method.

Inclusion criteria

- 1) Patients with fresh, first attack of AMI proved by history, ECG findings or cardiac enzyme analysis.
- 2) Patients with or without diabetes and hypertension.

Exclusion criteria

- 1) Patients with renal failure.
- 2) Patients with other cardiovascular problems like cardiomyopathy, rheumatic or valvular heart diseases based on history and echocardiography reports.
- 3) Estimated time of infarction > 24 hrs prior to admission i.e. window period > 24hrs.

III. Results

Seventy five patients (n=75) with established ischemic heart disease were studied and their mean age being 54.1±11.2 ranging from 32-70 years. They were studied serially on 1st, 3rd and 7th day after Myocardial Infarction attack. Hence 75 patients were included under the study group (n=75). Study group (Group-I) was classified into three sub groups i.e. sub group Ia, Ib, Ic, i.e. 1st day, 3rd day and 7th day follow up study respectively. Out of 75 cases 25 cases were hypertensive (n=25) and 20 cases (n=20) were diabetic.

Thirty subjects without any heart disease and renal pathology, who were in control group the mean serum Magnesium was 1.08 ±0.26 mmol/litre and mean Ca⁺² was 9.45±1.134 mg%. Mean serum Na⁺ and K⁺ were 137.38 ±3.67 meq/l and 4.21 ± 0.394 meq/L respectively. (Table-1) Statistical analysis was made between control and subgroups Ia and Ib and Ic. Analysis is performed by student t-test. There is significant decrease in all the parameters on 1st day of myocardial infarction (p<0.01) with more significance (p<0.001) in case of serum Magnesium and Serum Potassium levels, but there was no significant difference between control and groups Ib and Ic (p>0.05) in all the parameters. Intra group comparisons are showing a progressive increase in serum Magnesium and serum Potassium, S.Ca⁺² and S.Na⁺ levels by 7th day of follow up reaching near control values. (Table -1)

Table-1 Mean + SDs of Serum electrolytes in control and study groups on three days of follow up

Parameter	Control Group (n=30)	Study Group (n=75) Group I		
		Ia (1 st day) (n=75)	Ib (3 rd day) (n=75)	Ic (7 th day) (n=75)
Serum Magnesium (mmol/L)	1.08 ±0.26	0.74 ±0.21***	0.956 ± 0.176	1.11 ± 0.21
Serum Calcium (mg/dl)	9.45 ± 1.13	7.04 ±0.84**	8.6 ± 0.84	9.34 ± 0.97
Serum Sodium (meq/L)	137.38 ± 3.67	129.1 ±10.12**	136 ± 6.55	139 ± 4.05
Serum Potassium (meq/L)	4.21 ± 0.39	2.92 ±0.76***	3.41 ± 0.59	3.69 ± 0.601

***(p<0.001)**(p<0.01)

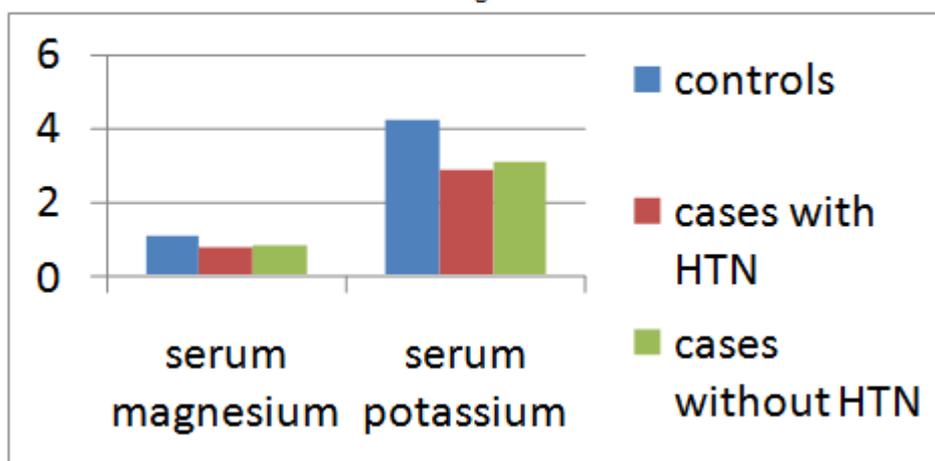
Mean levels of S.Mg⁺², Ca⁺², Na⁺ and K⁺ were compared in cases with and without Diabetes and Hypertension. [Table-2] There is significant decrease in Serum Magnesium and Serum Potassium levels in hypertensives and Diabetics when compared to controls (p<0.01) with no significant change in S.Ca⁺² and S. Na⁺ levels (p>0.05). Serum Magnesium level in cases with Diabetes is significantly lower than in cases without diabetes (p<0.05). There is lowering of S.Mg⁺² in cases with hypertension in AMI cases but there is no statistical significance. There is no significant change in other electrolyte levels in the different groups (Table-2) Fig-1 & Fig-2.

Table-2 Mean±SD of electrolyte levels in cases with and without hypertension and diabetes respectively.

	Control (n=30)	Cases with Hypertension (n=25)	Cases with Diabetes (n=20)	Cases without Diabetes and Hypertension (n=30)
Serum Magnesium (mmol/L)	1.08 ± 0.26	0.74 ± 0.14	0.68 ± 0.24*	0.789 ± 0.24
Serum Calcium (mg/dl)	9.45 ± 1.13	8.23 ± 1.07	8.35 ± 1.00	8.4 ± 0.25
Serum Sodium (meq/L)	137.38 ± 3.67	127.8 ± 11.47	132.3 ± 10.5	133.8 ± 0.88
Serum Potassium (meq/L)	4.21 ± 0.39	2.88 ± 0.77	3.3 ± 0.94	3.1 ± 0.94

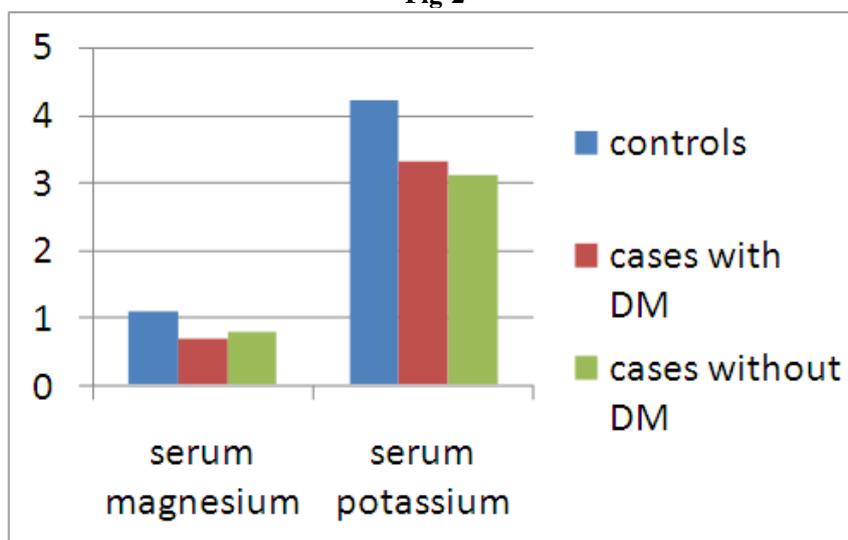
*(p<0.05)

Fig-1



Serum Mg & Sr. K levels in Controls, Cases with and without Hypertension

Fig-2



Serum Mg & Sr. K levels in Controls and in Cases with and without Diabetes mellitus

IV. Discussion

There is significant decrease in Serum Magnesium levels in Diabetics with AMI when compared to cases without diabetes ($p < 0.05$) and lowering of $S.Mg^{+2}$ level in hypertensive cases with AMI, with no significant change in $S.Ca^{+2}$ and $S.Na^{+}$ levels ($p > 0.05$). Low magnesium levels in diabetic and hypertensive cases is due to the effect of treatment with diuretics or due to glucosuria which increases urinary Mg^{+2} losses.^{8,9,10,11,12,13} Poor dietary intake, autonomic dysfunction altered insulin metabolism, glomerular hyperfiltration, osmotic diuresis recurrent metabolic acidosis, hypophosphatemia & hypokalemia may be contributory^{14,15,16,17} So contributory mechanisms for hypomagnesemia are multifactorial.

Hypomagnesemia has been reported to occur in 13.5 to 47.7% of non hospitalized patients with type 2 Diabetes Mellitus compared with 2.3 to 15% among their counter parts without Diabetes Mellitus.^{18,19,20,21,22,23}

Hypomagnesemia may induce altered cellular glucose transport, reduced pancreatic insulin excretion, defective postreceptor insulin signaling and altered insulin receptor interactions.²⁴ So Hypomagnesemia may be both cause and effect of impaired glucose metabolism. Our finding is in accordance with other studies.^{19,20,23}

In adults, low serum and intracellular magnesium concentrations are associated with Insulin resistance, impaired glucose tolerance and decreased insulin secretion.

Low level of magnesium predisposes to increase in arterial pressure as Mg^{+2} acts peripherally to produce peripheral vasodilation and hence fall in blood pressure.

Low magnesium levels may promote endothelial cell dysfunction & aggregation and thrombogenesis via increased platelet aggregation and vascular calcifications²⁵ and may also lead to the induction of proinflammatory and profibrinogenic response.^{26,27}

V. Conclusions

There is significant decrease in Serum Magnesium levels in Diabetics with AMI when compared to cases without diabetes and lowering of $S.Mg^{+2}$ level in hypertensive cases with AMI, with no significant change in $S.Ca^{+2}$ and $S.Na^{+}$ levels. Low Magnesium Levels may be a cause or effect of Diabetes. Maintaining the normal serum levels of Magnesium can prevent the development of Diabetes mellitus.

References

- [1]. Yago M, et al, *Frontiers in Bioscience* 5, July 2000
- [2]. Rude R, *Am J Cardiol* (Apr 18) 63 (14), 1989.
- [3]. Taubert K *Fortschr Med*, 112 (24), 1994.
- [4]. Romano T, *J Nutritional & Environ Med*, 7, 107-111, 1997.
- [5]. Gindler E et al (1971) *Clin Chem* 17 : 662
- [6]. Giletman H. *J 91967 Anal Biochem* 18: 52/ Bagainski E.S (1973) *5 Clin Chem Acta* 46:46
- [7]. Praful B Godkar *Textbook of Laboratory Technology*. Reprint 1999.
- [8]. Thomas Dykner Clars Helmers & Perolov wester, *Acta Med Scand* 1984 : 216, 127-132.
- [9]. Quamme GA : Renal handling of magnesium. In ; Massry & Glasscocks *Textbook of Nephrology*, 4th Edition , edited by Massry SH, Glasscock RJ , Baltimore, Lippincott Williams & WILKINS, 2001, PP 344-350.
- [10]. Dai LJ, Ritchie G, Kerstan D, Kang HS, Cole DEC, Quamme GA: Magnesium transport in the renal distal convoluted tubule. *Physiol Rev* 81: 51-81, 2001
- [11]. Dai LJ, Friedman PA, Quamme GA: Cellular mechanisms of chlorothiazide and potassium depletion on Mg^{2+} uptake in mouse distal convoluted tubule cells. *Kidney Int* 51: 1008-1017, 1997
- [12]. Dai LJ, Friedman PA, Quamme GA: Phosphate depletion diminishes Mg^{2+} uptake in mouse distal convoluted tubule cells. *Kidney Int* 51: 1710-1718, 1997
- [13]. Wong NLM, Quamme GA, O'Callaghan TJ, Sutton RAL, Dirks JH: Renal and tubular transport in phosphate depletion: A micropuncture study. *Can J Physiol Pharmacol* 58: 1063-1071, 1980
- [14]. Nijenhuis T, Renkema KY, Hoenderop JG, Bindels RJ: Acid-base status determines the renal expression of Ca^{2+} and Mg^{2+} transport proteins. *J Am Soc Nephrol* 17: 617-626, 2006
- [15]. Duarte CG: Effects of chlorothiazide and amipramizide (MK 870) on the renal excretion of calcium, phosphate and magnesium. *Metabolism* 17: 420-429, 1968
- [16]. Eknayan G, Suki WN, Martinez-Maldonado M: Effect of diuretics on urinary excretion of phosphate, calcium, and magnesium in thyroparathyroidectomized dogs. *J Lab Clin Med* 76: 257-266, 1970
- [17]. Hodler J, Roulin F, Haldimann B: Short-term effect of thiazides on magnesium and calcium metabolism and secondarily on that of phosphorus, uric acid, oxalate and cyclic AMP [in French]. *Nephrologie* 4: 60-63, 1983
- [18]. Pham PC, Pham PM, Pham PA, Pham SV, Pham HV, Miller JM, Yanagawa N, Pham PT: Lower serum magnesium levels are associated with more rapid decline of renal function in patients with diabetes mellitus type 2. *Clin Nephrol* 63: 429-436, 2005 [published erratum appears in *Clin Nephrol* 64: 248, 2005]
- [19]. McNair P, Christensen MS, Christiansen C, Madsbad S, Transbol I: Renal hypomagnesaemia in human diabetes mellitus: Its relation to glucose homeostasis. *Eur J Clin Invest* 12: 81-85, 1982
- [20]. Mather H, Nisbet JA, Burton GH, Poston GJ, Bland JM, Bailey PA, Pilkington TR: Hypomagnesemia in diabetes. *Clin Chim Acta* 95: 235-242, 1979
- [21]. De Lordes Lima M, Cruz T, Pousada JC, Rodrigues LE, Barbosa K, Cangucu V: The effect of magnesium supplementation in increasing doses on the control of type 2 diabetes. *Diabetes Care* 21: 682-686, 1998
- [22]. Walti MK, Zimmermann MB, Spinass GA, Hurrell RF: Low plasma magnesium in type 2 diabetes. *Swiss Med Wkly* 133: 289-292, 2003.
- [23]. Ma J, Folsom AR, Melnick SL, Eckfeldt JH, Sharrett AR, Nabulsi AA, Hutchinson RG, Metcalf PA: Associations of serum and dietary magnesium with cardiovascular disease, hypertension, diabetes, insulin, and carotid arterial wall thickness: The ARIC study.

- Atherosclerosis Risk in Communities Study. *J Clin Epidemiol* 48: 927–940, 1995
- [24]. Grafton G, Baxter MA : The role of Magnesium in Diabetes Mellitus, *J Diabetes Complication* 6 : 143-149, 1992.
- [25]. Ray S Signier Y : Role of Magnesium & Potassium in the pathogenesis of atherosclerosis , *Magnesium* 3 : 226-228, 1984.
- [26]. Shivakumar K: Pro-fibrogenic effects of magnesium deficiency in the cardiovascular system. *Magnes Res* 15: 307– 315, 2002
- [27]. Maier JAM, Malpuech-Brugere C, Zimowska W, Rays-siguier Y, Mazur A: Low magnesium promotes endothelial cell dysfunction: Implications for atherosclerosis, inflammation and thrombosis. *Biochim Biophys Acta* 1689: 13–21, 2004