Some renal function parameters in individuals with metabolic syndrome

Mohammad A. Alkataan

Department of Clinical Pharmacy, College of Pharmacy, University of Mosul, Mosul, Iraq

<u>Received 16/10/2011</u> Accepted 20/5/2012

Abstract

Metabolic syndrome (MS), a cluster of risk factors for cardiovascular diseases. This syndrome characterized by: Insulin resistance, hyperinsulinemia, abdominal obesity, elevated blood pressure, lipid abnormalities and low grade inflammatory state. There are growing data demonstrated the relation between MS and renal impairment, these data revealed that individual with MS at higher risk to develop chronic kidney disease. (a) To determined the changes in renal function parameters in MS individuals.(b) determine the effect of age, sex and BMI on the measured parameters . This study was conducted during period from January to September 2011. Fifty apparently healthy individual (30 male and 20 female) were included in this work as control with age range 25±6.3 years, BMI range 21± 3.7 Kg/m² and weight range 55 ± 3.9 Kg and another fifty individual(30 male and 20 female) were selected to have at least three of the WHO criteria of MS. Data were presented as mean \pm SD, 2-sample t-Test was used to show the significance changes between the two groups. The effect of age and BMI on measured parameters were determined using Person - correlation. This study revealed that MS individual shows a significant increase in SFG,TC,TG, B.Urea, S.Cr and U.Sp-G when compared to those of the controls, while HDL-C and e-GFR shows significant reduction when compared to those of control table 1. In both group e-GFR significantly correlated to individual weight (r = 0.02), BMI (r =.0.075).SFG significantly correlated to B.Urea, S.Cr., e-GFR and U.Sp-G in MS individual (r = 0.03) but not in control group. In conclusion: MS individual show significant changes in renal function that may related to higher susaptability of this group to developing renal diseases

الملخص

المتلازمة الأيضية، هي مجموعة من عوامل الخطر لأمراض القلب والأوعية الدموية و هذا التناذر يتصف ب : مقاومة الانسولين ، فرط انتاج الانسولين ، والبدانة في البطن ، وارتفاع ضغط الدم واضطر ابات الدهون وتفاعل التهابي ضعيفة وكشفت البيانات أن هناك نموا في العلاقة بين مرض المتلازمة الايضية و امراض الكلوي , إن الافراد المصابين بالمتلازمة الايضية أكثر عرضة للنك معن الخوارد المصابين بالمتلازمة الايضية أكثر عرضة للالصابة بامراض الكلى .: (أ) تحديد التغيرات في بعض وظائف الكلى عند الأفراد المصابين بالمتلازمة الايضية (ب) تحديد التغيرات في بعض وظائف الكلى عند الأفراد المصابين بالمتلازمة الايضية (ب) تحديد عرضة للاصابة بامراض الكلى .: (أ) تحديد التغيرات في بعض وظائف الكلى عند الأفراد المصابين بالمتلازمة الايضية (ب) تحديد تأثير عوامل الجنس والعمر وم عامل كتلة الجسم على معايير المقاسق أجريت هذه الدراسة في الفترة من يناير إلى سبتمبر 2011. وقد مشملت الخمسين شخصا من الأصحاء (30 من الذكور والإناث 20) في هذا العمل كمجموعة التحكم كما سن 25 ±6.6 سنة، معامل أشتملت الخمسين شخصا من الأصحاء (30 من الذكور والإناث 20) في هذا العمل لمجموعة التحكم كما سن 25 ±6.6 سنة، معامل المتلة الجسم 12 مجموعة التحكم كما سن 25 ±6.6 سنة، معامل ألقا الإناث) لديهم ما لايقل عن ثلاثة من صفات المعتمدة لتشخيص المتلازمة الايضية سبعايير منظمة الصحة العالمية . وقد تع عرضت البيانات ألفون إذات 20 كلغ ± 3.9 و خمسين فرد آخرين (30 من الذكور و 20 وقد تم عرضت البيانات كمعدل ± الانحراف المعياري ، كما استخدم اختبار T لإظهار تغييرات بين المتلازمة الايضية معايير مناملة المعيد . وقد تع عرضت البيانات كمعدل ± الانحراف المعياري ، كما استخدم اختبار T لإظهار تغييرات بين المتلازمة اليضية العامية . وقد عرضت البيانات كمعدل ± الانحراف المعياري ، كما استخدم اختبار T لإظهار تغييرات بين المتلازمة العام المعوم عني وزم معامل في معايير . وي مالم في معرف ألموان في مالمي ألفور .. والم الفراد المعايية . وعامل مع مع معايير منعان في الد مالالمية . وي مالم ورفي معامل وي مالمتازمة الايضية معايير منور و 20 وقد معنوية في ماتو وي مالم وي مالمي . وقدم ألفور المصابين بالمتلازمة اليوي .. وي مالم معروي يام معوون ياده معاوي وي مالي المعار .. وي مالم وي مالمون يا من مع معوم معالي يالمم .. ولمما مل معايي وي مالمان ..

و الكثافة النوعية للبول عند المصابين بالمتلازمة الايضية (ص = 0.03) الاستنتاج: يظهر المصابين بالمتلازمة الايضية تغيرات كبيرة في وظائف الكلى التي قد تتعلق بارتفاع احتمالية إصابتهم بأمراض الكلى .

Introduction

Metabolic syndrome (MS), a cluster of risk factors for cardiovascular diseases . This syndrome characterized by: Insulin resistance, hyperinsulinemia, abdominal obesity, elevated blood pressure, lipid abnormalities and low grade inflammatory state (12,3). There are growing data demonstrated the relation between MS and renal impairment, these data revealed that individual with MS at higher risk to develop chronic kidney disease ⁽⁴⁾. Chen and his collogues described that MS individuals had 80-130% higher risk to develop renal impairment than non-MS subject ⁽³⁾. Also they described the relationship among increase serum fasting glucose (SFG)more than 6.1 mmol/L, hypertension, low e-GFR and Microalbuminuria⁽²⁾. Muntner *et al*, described the strong association between renal manifestation and lipid abnormalities that include low high density lipoprotein (HDL-C) and high serum triglycerides (TG) in MS individuals ⁽⁵⁾. Kambham et al, described focal segmental glumerolosclorsis association with central obesity that now referred as (obesity related glumerolopathy)⁽⁶⁾. Moreover the low grade inflammatory that seen in MS individuals may related to release of adipocytic-cytokines Leptin,IL-6 ,TNF-α that include: and adipoaectin that involve at least partially in promoting renal impairment as described by Wiss⁽⁷⁾. Urinary specific gravity (U.Sp-G) that reflect the glumerolar filtration and dilution/ concentration ability of kidney, was related to MS by the fact that MS individual have elevated SFG and for each 1% increase in SFG there are 0.004 unit increase in U.Sp-G as described by Schumann and Schweitzer⁽⁸⁾. This study was design to describe the changes some renal function parameters in in individuals with MS. Also define the changes in e-GFR and urinary specific gravity in MS

individuals. The effect of age, BMI and individuals weight on these parameters were also studied.

Subjects and Methods

This study was conducted during period from January to September 2011. This study received approval from Ethics and scientific committee in department of clinical pharmacy - university of Mosul. Fifty apparently healthy individual (30 male and 20 female) were included in this work as control with age range 25 ± 6.3 years, BMI range 21 ± 3.7 Kg/m² and weight range 55 ± 3.9 Kg and another fifty individual(30 male and 20 female) were selected to have at least three of the WHO criteria of MS that include: BMI ≥ 25 Kg/m², BP \geq 140-110 mmHg, serum fasting glucose \geq mmol/L, hypertriglyceridemia 7.1 with diminish of HDL level ⁽⁹⁾. The age range 25 ± 5.76 years, BMI range 27 ± 4.8 Kg/m² and weight range $69.8\pm$ 65Kg. The effect of sex was eliminated by using symmetrical number of each sex. Serum fasting glucose assayed by oxidase /peroxidase colorimetric glucose method⁽¹⁰⁾,</sup> Total serum cholesterol bv Richmond-enzymatic methods⁽¹¹⁾, HDL- c measured by Lopez-Virella method⁽¹²⁾ and serum triglycerides were measured using Fossati-enzymatic method ⁽¹³⁾ serum creatinine measured by Jaffa reaction method⁽¹⁴⁾Blood urea was measured using Mc Neely method⁽¹⁵⁾ and urinary specific gravity was measured by estimated while. Uriometer GFR was calculated using the following equation⁽¹⁶⁾: e-GFR= (140-age)× Wt/ 72× S.Cr

note: in female this equation multiplied by 0.85

Data were presented as mean \pm SD, 2-sample t-Test was used to show the significance changes between the two groups. The effect of age and BMI on measured parameters were determined using Person - correlation.

Results

This study revealed that MS individual shows a significant increase in SFG,TC,TG, B.Urea, S.Cr and U.Sp-G when compared to those of the controls, while HDL-C and e-GFR shows

significant reduction when compared to those of control table 1. In both group e-GFR significantly correlated to individual weight (r = 0.02), BMI (r = .0.075).

SFG significantly correlated to B.Urea, S.Cr., e-GFR and U.Sp-G in MS individual (r = 0.03) but not in control group.

Table (1):- Biochemical parameters in individuals with MS and control groups.

	81
Control	MS individual
5.17 ± 0.36	7.34± 0.29***
5.13±0.46	6.61±.71***
1.67 ± 0.25	1.75±0.5**
0.99 ± 0.15	$0.90 \pm 0.13 *$
6.12 ± 0.83	8.76±1.16**
0.94 ± 0.19	1.4±0.05***
97.9 ± 30.7	66.33± 9.08**
1.0213 ± 0.0038	$1.0355 \pm 0.0046 **$
	5.17 ± 0.36 5.13 ± 0.46 1.67 ± 0.25 0.99 ± 0.15 6.12 ± 0.83 0.94 ± 0.19 97.9 ± 30.7

Note: P<0.05=*,P<001=**, P<0.001=***

Discussion

This study demonstrated that a significant changes in all renal function parameters measured in this work and this related to many factors. The significant increase in S.Cr, B.Urea and U.Sp.-G with significant reduction in GFR may related to sustain hyperglycemia that associated with MS that cause irreversible damage to renal structures and this result agree with results obtained by Segure *et al*⁽¹⁷⁾.</sup>The significant reduction in GFR may related to the significant elevation in S.TG that seen in MS individuals and this agree with result obtained by Samulsson et al. how described significant deterioration in GFR with significant elevation in S.TG that reflect as

increase in S.Cr⁽¹⁸⁾. The significant increase in S.Cr may related to significant reduction in HDL-C in MS individuals and this agree with results obtain by Muntner et al. how described a negative relation between S.Cr and HDL-C in MS individuals⁽⁵⁾. The significant reduction in e-GFR that seen MS individuals agree with results obtained by Chen et al. how demonstrated a significant reduction in GFR less than 60 ml/min/1.73 m^2 in MS individuals and related it to the sustain increase in BP that occur due to increase in adrenergic activity that caused by sustain hyperglycemia and dyslipidemia in this $group^{(3,19)}$. The sustain high BP lead to peripheral arteries damage that ended with left ventricular heart failure leading to significant reduction in renal perfusion that in turn leading to significant reduction in GFR and increase $S.Cr^{(20-22)}$. In conclusion : MS individual show significant changes in renal function that may related to higher susptability of this group to developing renal diseases .

References

- Hegel R., Pollex R. Genetic and physiological insight into metabolic syndrome. Am J Physiol Regul Integr Comp Physiol 2005;289:663-7
- Chen J., Muntner P., Hamm L., *et al.* Insulin resistance and risk of chronic kidney disease in nondiabetic U.S.J Am Soc Nephrol. 2003;14(2):469-77.
- 3. Chen J., Hamm L., Jones D., et al. the metabolic syndrome and chronic kidney disease in U.S. adult Ann Inter Med 2004;140:167-174.
- 4. Hoehner C., Greenlund K., CaspernM., et al. Association of insulin resistance syndrome and Microalbuminuria among non-diabetic native American : the inter-tribal heart project. J Am Soc Nephrol 2002;13:1626-1634.
- Muntner P., Coresh J., Smith J., et al. Plasma lipid and risk of development impaired renal function:the atherosclerosis risk in community study. Kidney Int 2000;58:293-301.
- 6. Kambham N., Markowitz G., Valeri A., et al .obesity related glumerolopathy :an emerging epidemic. KidneyInt 2001;59:1498-1509.
- Wiss B. The inflammatory syndrome: the role of adipose tissue cytokines in metabolic disorder linked to obesity. J Am Soc Nephrol 2004;15: 2792- 1800.
- Schmann G, Schweitzer S. Examination of urine In Kalplan L.,Pesce A.,(eds). Clinical chemistry: Theory, Analysis and Correlation. Mosby 2ed Ed. St.Louis USA.1989 pp:820.
- 9. Vega G. Obesity : the metabolic syndrome and cardiovascular disease. Am Heart J 2000;142:1108-1116.

- ventricular heart failure leading to significant reduction in renal perfusion that in turn leading to significant reduction in GFR and increase S.Cr⁽²⁰⁻²²⁾. In conclusion : MS
 - Richmond W. Preparation and properties of a cholesterol oxidase from nocardia Sp. and its application to the enzymatic assay of total serum cholesterol in serum. Clin. Chem.1973; 19: 1350-1356
 - 12. Lopez Virella M., Stone P., Colwell J. Cholesterol determination in high density lipoproteins separated by three different methods. Clin. Chem.1977; 23 : 882-884.
 - 13. Fossti P., Prencipe L. Serum Triglycerides determination colorimtrically with an enzyme that produces hydrogen peroxides. Clin. Chem.1982; 28 : 2077-2085
 - Cockcroft D., Gault M. Predication of creatinine clearance from serum creatinine Nephron 1976;16:31-41
 - 15. McNeely M. Renal function In Gradwholes: Clinical laboratory methods and diagnosis. In Sonnen Wirth A. and Jarett L.8TH Ed. Mosby St. Louis USA.1980 pp:504-515.
 - 16. Bishop M.,Fody E.,Schoeff L. Clinical chemistry :Procedure, Principle and correlation .Lippincott Willams New York USA 2005.pp:302-303.
 - 17. Segura J., Compo C., Rollaan C., et al. Hypertension damage in metabolic syndrome is associated with glucose metabolism distribances. J Am Soc Nephrol 2004;15:429-446.
 - 18. Samuelsson O., Attaman P., Knight-Gibson C., et al. Complex apo-lipoprotein B containing lipoprotein particles are associated with higher rate of progression of human chronic renal insufficiency. J Am Soc Nephrol 1998;9:1482-1488.
 - 19. Resnick L.Cellular Ca⁺⁺ and Mg⁺⁺ metabolism in pathophysiology and treatment of hypertension and related metabolic disorders. Am J Med 1992;93:115-205.
 - 20. Boden G., Jadali F., White J. Effects of fat on insulin –stimulated carbohydrate metabolism in normal men. J Clin Inves 1991;88:960-966.

- 21. Boullin D.The action of extracellular Cat-ions on the release of the sympathetic transmitter from peripheral nerve. J Physol 1967;189:85-99.
- 22. Seelng M. Consequences of Mg⁺⁺deficiency, enhancement of stress action: preventive and therapeutic implication. J Am Coll Nutr 1994;13:429-446.