Adverse Effects of Monosodium Glutamate on Liver and Kidney Functions in Adult Rats and Potential Protective Effect of Vitamins C and E

Manal Said Tawfik1*, Nawal Al-Badr2

1Department of Food Science and Technology, Faculty of Agriculture-El-Shatby, Alexandria University, Alexandria, Egypt; 2Department of Food Nutrition, King Saud University, Riyadh, Saudi Arabia.
Email: *mana92ti@yahoo.com

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ABSTRACT
Monosodium glutamate (MSG) was administrated to rats at doses of 0.6 and 1.6 mg/g body weight for 14 days. Body weight and relative liver and kidney weight of rats were significantly increased. On regard to liver functions, the activities of alanine aminotransferase (ALT) and γ-glutamyle transferase (GGT) significantly increased in the serum, on MSG administration, meanwhile, serum total protein, albumin and serum total bilirubin significantly decreased. MSG had adverse effects on kidney functions as serum urea and serum creatinine were significantly increased. Vitamin C (0.3 mg/g body weight) and vitamin E (0.2 mg/g body weight) co-administrated with MSG, significantly restored the body weight and relative liver and kidney weights to control levels. In the presence of vitamin C and vitamin E the activities of ALT and GGT in the serum were also significantly reduced to become comparable with control trial. Consequently, serum total protein, albumin and serum total bilirubin were significantly increased in the serum, while serum urea and serum creatinine were significantly decreased after administration of each vitamin. The results showed that MSG at doses of 0.6 and 1.6 mg/g of body weight may cause an adverse effect on the hepatic and renal functions which might be due to oxidative stress induced by MSG on the liver and renal tissue. Supplementation of vitamin C and vitamin E was capable of ameliorating MSG-induced oxidative stress on hepatic and renal functions.

Keywords: Monosodium Glutamate; Hepatic Stress; Renal Stress; Alanine Aminotransferase; γ-Glutamyle Transferase; Creatinine; Total Bilirubin

1. Introduction
Monosodium glutamate (MSG) known as AJI-NOMOTO is the sodium salt of glutamic acid [1]. MSG contains 78% of glutamic acid, 22% of sodium and water [2]. Glutamate is one of the most common amino acids found in nature and is the main component of many proteins and peptides of most tissues. Glutamate is also produced in the body and plays an essential role in human metabolism. It is a major component of many protein-rich food products either in free or bound state of animal such as meat, fish, milk and cheese or vegetable origins such as mushroom and tomato [3]. MSG is commonly used as a flavor enhancer especially in Chinese, Thai-ese and Japanese foods [4,5].

When MSG is added to food, it provides a flavoring function similar to the naturally occurring free glutamate which differ from the four classic tastes of sweet, sour, salty and bitter [6]. As food additive, MSG is described and listed on food labels as a “Flavoring” or “hydrolysed vegetable protein”. Through its stimulation of the or-sensory receptors and by improving the palatability of meals, MSG influences the appetite positively, and induces weight gain. Despite its taste stimulation and improved appetite enhancement, reports indicate that MSG is toxic to human and experimental animals [7]. MSG could produce symptoms such as numbness, weakness, flushing, sweating, dizziness and headaches. In addition to these MSG symptom complex, ingestion of MSG has been alleged to cause or exacerbate numerous conditions, including asthma, urticaria, atopic dermatitis, ventricular arrhythmia, neuropathy and abdominal discomfort [8].

MSG has a toxic effect on the testis by causing a significant oligozoospermia and increase abnormal sperm morphology in a dose-dependent fashion in male Wistar rats [9]. It has been implicated in male infertility by causing testicular hemorrhage, degeneration and alteration of sperm cell population and morphology [10]. It has been reported that MSG has neurotoxic effects resulting
in brain cell damage, retinal degeneration, endocrine disorder and some pathological conditions such as addiction, stroke, epilepsy, brain trauma, neuropathic pain, schizophrenia, anxiety, depression, Parkinson’s disease, Alzheimer’s disease, Huntington’s disease, and amyotrophic lateral sclerosis [2]. It cannot be stated that MSG is the cause of such varied conditions as epilepsy and Alzheimer’s disease, although there may be concerns of its involvement in its etiology [2].

The Food and Drug Administration (FDA) of the United States reports that MSG is safe and that it should be maintained on the “Generally Recognized as Safe” (GRAS)-list of foods. MSG is thus reportedly permitted as a safe food additive that needs no specified average, daily intake or an upper limit intake requirement [11].

Liver is the largest gland in the mammalian body. The hepatocytes have metabolic functions that deal with very essential processes such as detoxification, deamination, transamination, removal of ammonia in the form of urea, biosynthesis and release of the non-essential amino acids and plasma proteins with the exception of immuno gamma globulins, gluconeogenesis, storage of glycogen, conversion of carbohydrates and proteins into lipids, synthesis of lipoproteins, phospholipids and cholesterol, oxidation of fatty acids, storage of iron in the form of ferritin as well as storage of vitamins A, D and B12. Several functional tests have been formulated to explore hepatic status [12-16]. Several enzymes have been determined to explore hepatic status such as alanine amino transferase (ALT) and aspartate amino transferase (AST). In addition some other tests include measurement of serum lactic dehydrogenase (LDH), gamma glutamyl transpeptidase (GGT), alkaline phosphatases and 5-nucleotidase activities are employed [17-19].

The Kidney is a paired organ located in the posterior abdominal wall, whose major functions include the removal of toxic metabolites and waste products from the blood and regulation of the amount of fluid and electrolytes balance in the body. To test functions of the kidneys routine urinalysis is used to measure serum urea, creatinine, sodium and potassium and serum bicarbonate [13,18-20].

Vitamins have indispensable role in almost all biochemical reactions and they are ideal antioxidants able to increase tissue protection from oxidative stress due to their easy, effective and safe dietary administration in a large range of concentrations [21,22]. Vitamin E (α-Tocopherol [α-Toc]) is the primary membrane bound, lipid-soluble, chain-breaking antioxidant that protects cell membranes against lipid peroxidation [23-26].

Vitamin E pre-treatment has been reported to be beneficial in preventing formaldehyde-induced tissue damage in rats [24,27]. The preventive effect of vitamin E on cypermethrin or endotoxin-induced oxidative stress in rat tissues is suggestive of its antioxidant activity [28-33].

L-ascorbic acid is the first to become depleted on the exposure to oxidative stress [34]. Normal L-ascorbate level has a therapeutic benefit due to its ability to reduce the oxidative stress by reacting with superoxide and hydroxide radicals as well as alkyl, peroxy and alkoxyl radicals, thereby it can neutralize these radicals and stop the initiation and propagation of chain reaction [35,36].

Although some information is available on the MSG-induced oxidative stress and toxicity, the studies on the effect antioxidants, especially those consumed in food on MSG-induced toxicity and oxidative stress are lacking. Moreover, most of studies, so far on MSG, have been carried out on very high doses (4 mg/g and above) except of the study of Onyema et al., 2006 [37]. Thus, There is need to substantiate whether MSG induce oxidative stress at an appreciably lower doses and study the effect of nutritional antioxidants on it.

The objective of this research was designed to study the effect of MSG at two oral low doses of 0.6 and 1.6 mg/g body wt on liver and kidney functions in albino rats. The possible protective effect of vitamin E (at oral dose of 0.2 mg/g body wt) and vitamin C (at oral dose of 0.3 mg/g body wt) against MSG induced hepatic and renal stress was also investigated.

2. Materials and Methods

2.1. Animals

Adult albino rats of both sex with average weight of 95 g were obtained from King Saud University Animal Care Center. The animals were housed in cages under standard hygienic condition and were fed with rat chow and water ad labium. In order to optimize treatment doses, all animals were fasted for 1 h prior to treatment administration.

2.2. Chemicals

MSG was purchased from BDH laboratory (Poole, UK), vitamin C and vitamin E were obtained from Sigma-Aldrich Chemicals Co. (St. Louis, MO, USA). All chemical reagents were of analytical grades and purchased from Sigma Chemical Co. (St. Louis, MO, USA).

2.3. Experimental Design

Animals were randomly assigned into seven groups of (n = 8); Group 1 and groups from 2 to 7 served as control and experimental groups respectively. Groups 2 and 3 were given 0.6 mg/g body weight and 1.6 mg/g body weight of MSG respectively on a daily basis; groups 4
and 5 were given 0.6 mg/g body weight and 1.6 mg/g body weight of MSG plus 0.3 mg/g body weight vitamin C on a daily basis; groups 6 and 7 were given 0.6 mg/g body weight and 1.6 mg/g body weight of MSG plus 0.2 mg/g body weight vitamin E on a daily basis. Group 1, received equal amount of feeds without MSG and vitamins added for fourteen days. The rats were sacrificed on the fifteenth day of the experiment. The choice of MSG doses, vitamin E and C doses was based on previous reported studies [37,38].

2.4. Body, Relative Liver and Kidney Weight

Body weight of each animal was determined before treatments and before sacrifice. Liver and kidneys of each animal were dissected out and weighed.

2.5. Blood Collection and Separation

At the end of treatment period the animals were sacrificed 24 hr following the last given dose. Blood samples were withdrawn and collected in glass tubes. Serum was separated by centrifugation at 3000 rpm for 10 min and stored at −80°C bending biochemical analysis.

2.6. Biochemical Assays

2.6.1. Assessment of Liver Function

2.6.1.1. Alanine Aminotransferase (ALT)

Alanine aminotransferase catalysed the transfer of an amino group between the amino acids L-alanine and L-glutamate. The ketoacids formed in this process were α-ketoglutarate and pyruvate. The pyruvate formed reacted with dinitrophenylhydrazine to produce a corresponding dinitrophenylhydrazone, which was measured with the spectrophotometer at 505 nm [39].

2.6.1.2. Protein

The peptide bonds of plasma protein in plasma reacted with biuret reagent which contained copper in alkaline solution to form a violet colored which was measured with a spectrophotometer at 540 nm [40].

2.6.1.3. Bilirubin

Plasma was added to a solution of sodium acetate and caffeine (sodium benzoate) which was then added to diazotized sulfanilic acid to form a purple azo bilirubin. The sodium acetate buffered the pH of the diazotized sulfanilic acid. The reaction was terminated by the addition of ascorbic acid which destroyed the excess diazo reagent. A strongly alkaline tartrate solution was then added to convert the purple azobilirubin to blue azobilirubin and the intensity of the color was measured at 600 nm with a spectrophotometer [39].

2.6.1.4. Albumin

Bromocresol green, a dye at an acid pH of 3.8 preferentially bound albumin to produce a shade of green color which was measured with a spectrophotometer at 630 nm [41].

2.6.1.5. γ Glutamyltransferase (GGT)

For determination of GGT activity, the serum sample was added to a substrate solution containing glycylglycine, MgCl₂ and γ glutamyle-p-nitroanilide in 0.05 M tris (free base), pH 8.2. The mixture was incubated at 37°C for 1 min interval for 5 min. The activity of GGT was calculated from the absorbance values [37].

2.6.2.2. Serum Creatinine

Creatinine Analyzer-2 (Beckman Coulter Inc., USA) in combination with a specific kit of reagents (Hichem Creatine Pak, Elan Diagnostics, USA) were employed to calculate creatinine content of the serum samples [42].

2.7. Statistical Analysis

Results obtained from the experiment were analyzed using analysis of variance (ANOVA), while comparisons were made using the Dunnet’s test at P < 0.05 level of significance.

3. Results and Discussion

3.1. Body Weight

Comparison of the final body weight with the initial body weight in each group revealed a significant difference and hence the increase of about 34.6%, 44.5% was considerable in group 2 and group 3 respectively (MSG treated animals) compared with an increase of about 31.8% in Group 1 (control) (Table 1). MSG intake could induce an increase in energy intake [43] which could lead to obesity [44] or alter the levels of carbohydrates, lipids and proteins in rats [45].

The controlling of body weight in MSG + V.C treated animals (31.5%, 30.9%) in group 4 and group 5 and MSG + V.E treated animals (30.5%, 30.2%) in group 6 and group 7 reflected the possible role of vitamins which
Table 1. Changes in body, liver and kidneys weights of control and rats treated with mono sodium glutamate (MSG) in the absence and presence of Vitamin C (V.C) or Vitamin E (V.E) expressed in grams.

<table>
<thead>
<tr>
<th>Groups</th>
<th>Initial Body Weight (g)</th>
<th>Final Body Weight (g)</th>
<th>Relative Liver Weight (g/100g)</th>
<th>Relative Kidneys Weight (g/100g)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Group 1 (Control)</td>
<td>88 ± 5.2</td>
<td>116 ± 6.4</td>
<td>3.65 ± 0.12</td>
<td>0.75 ± 0.02</td>
</tr>
<tr>
<td>Group 2 (0.6 mg/g b.w. MSG)</td>
<td>101 ± 2.1</td>
<td>136 ± 5.3</td>
<td>3.85 ± 0.17</td>
<td>0.78 ± 0.01</td>
</tr>
<tr>
<td>Group 3 (1.6 mg/g b.w. MSG)</td>
<td>101 ± 1.3</td>
<td>146 ± 5.1</td>
<td>4.16 ± 0.21</td>
<td>0.80 ± 0.01</td>
</tr>
<tr>
<td>Group 4 (0.6 mg/g b.w. MSG + 0.3 mg/g b.w. V.C)</td>
<td>92 ± 4.7</td>
<td>121 ± 4.7</td>
<td>3.68 ± 0.13</td>
<td>0.76 ± 0.01</td>
</tr>
<tr>
<td>Group 5 (1.6 mg/g b.w. MSG + 0.3 mg/g b.w. V.C)</td>
<td>94 ± 3.6</td>
<td>123 ± 3.9</td>
<td>3.77 ± 0.09</td>
<td>0.70 ± 0.02</td>
</tr>
<tr>
<td>Group 6 (0.6 mg/g b.w. MSG + 0.2 mg/g b.w. V.E)</td>
<td>95 ± 2.7</td>
<td>124 ± 3.4</td>
<td>3.60 ± 0.15</td>
<td>0.72 ± 0.03</td>
</tr>
<tr>
<td>Group 7 (1.6 mg/g b.w. MSG + 0.2 mg/g b.w. V.E)</td>
<td>96 ± 5.1</td>
<td>125 ± 3.1</td>
<td>3.67 ± 0.14</td>
<td>0.78 ± 0.04</td>
</tr>
</tbody>
</table>

Mean ± SD. N = 8 for the control; N = 8 for treated group; Mean of the final body weight is compared with that of the initial; Mean of the relative organ weight in treated group is compared with the corresponding mean of the control. 
P value of less than 0.05 indicates a significant difference between the compared means.

could lead to changes in functions against MSG oxidant effect of the organs that are regulated by the nervous system and metabolic activities [46].

### 3.2. Relative Liver and Kidneys Weight

A significant increase in the liver and kidney weight of the rats was observed after administration of MSG at two employed doses (Table 1). Thus, could be attributed to an increase in activity of inflammatory agents that could have resulted to inflammation of liver and kidney tissues [47].

Vitamin C and Vitamin E reduced the MSG-induced increase in the liver and kidney weight (Table 1), and vitamin E was more effective especially with higher intake of MSG, possibly via its action as a radical scavenger. By scavenging the radicals that contributed to oxidative stress induced by MSG, Vitamin E could help in reducing inflammation [37]. The changes in relative liver and kidneys weights have been accompanied with changes in functional aspects.

### 3.3. Liver Functions

#### 3.3.1. Serum Alanine Aminotransferase (ALT)

Table 2 shows activities of ALT enzyme that were measured in serum samples. Significant (p < 0.05) increases in the serum alanine aminotransferase was observed in the MSG-treated rats compared to the control rats (Table 2). Alanine aminotransferase (ALT) had activities means of 32.1, 40.3 and 46.6 U/L in serum of the control and MSG treated adult rats (0.6 mg/g and 1.6 mg/g body weight) respectively. The observed increase in activity was around 25.5%, 45.2% in the two doses. The ALT enzyme is a sensitive marker of liver damage [48]. Therefore, the increase in the serum ALT activity might perhaps be an indication of liver damage. MSG could dissociate easily to release free glutamate. The diminution of GLU produces ammonium ion (NH₄⁺) that could be toxic unless detoxified in the liver via the reactions of the urea cycle. Thus, the possible ammonium ion overload that may occur as a result of an increased level of glutamate following MSG intake could damage the liver, consequently releasing the ALT enzyme that may lead to its observed elevation. This increase could also be explained by free radical production which reacts with polyunsaturated fatty acids of cell membrane leading to impairment of mitochondrial and plasma membranes resulting in enzyme leakage [49].

The result seemingly agrees with the reports of Farombi and Onyema (2006) and Onyema et al. (2006) [37,50] that the activity of serum ALT increased in male rats that were fed MSG probably due to the finding that MSG induced oxidative stress in the liver. Thus, it could be concluded that MSG may be hepatotoxic at a low dose, hence should be avoided during the treatment of liver disorders. Furthermore, since ALT was a strong positive indicator of insulin resistance, diabetes mellitus and obesity [51] which are risk factors for coronary heart disease [52-54], the use of MSG even at low doses should not be encouraged because of the possible health implications. The serum ALT shows functional activity of liver. An increase in the activities of these enzymes indicates an effect due to the doses.

Administration of V.C and V.E resulted a significant reduction in the serum level of ALT enzyme at both MSG doses (Table 2). The use of V.E found to be more
Table 2. Liver function indexes of the control and treated adult rats with mono-sodium glutamate (MSG) in the absence and presence of Vitamin C (V.C) or Vitamin E (V.E)\(^1\).

<table>
<thead>
<tr>
<th>Groups</th>
<th>ALT (μ/L)</th>
<th>Serum Total Protein (g/dl)</th>
<th>Serum Total Bilirubin (mg/dl)</th>
<th>Albumin (g/L)</th>
<th>GGT (μ/L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Group 1 (Control)</td>
<td>32.1 ± 2.1(^a)</td>
<td>6.5 ± 0.1(^a)</td>
<td>0.68 ± 0.03(^a)</td>
<td>31.3 ± 2.9(^a)</td>
<td>20.5 ± 2.2(^a)</td>
</tr>
<tr>
<td>Group 2 (0.6 mg/g b.w. MSG)</td>
<td>40.3 ± 3.5(^b)</td>
<td>6.1 ± 0.2(^b)</td>
<td>0.61 ± 0.02(^b)</td>
<td>22.2 ± 3.1(^b)</td>
<td>22.1 ± 1.8(^b)</td>
</tr>
<tr>
<td>Group 3 (1.6 mg/g b.w. MSG)</td>
<td>46.6 ± 1.4(^c)</td>
<td>5.8 ± 0.2(^c)</td>
<td>0.54 ± 0.02(^c)</td>
<td>20.6 ± 4.2(^c)</td>
<td>24.3 ± 1.2(^b)</td>
</tr>
<tr>
<td>Group 4 (0.6 mg/g b.w. MSG + 0.3 mg/g b.w. V.C)</td>
<td>34.3 ± 2.8(^a)</td>
<td>6.4 ± 0.1(^a)</td>
<td>0.65 ± 0.03(^a)</td>
<td>28.5 ± 3.4(^a)</td>
<td>21.2 ± 2.3(^a)</td>
</tr>
<tr>
<td>Group 5 (1.6 mg/g b.w. MSG + 0.3 mg/g b.w. V.C)</td>
<td>35.2 ± 1.2(^d)</td>
<td>6.1 ± 0.1(^b)</td>
<td>0.58 ± 0.03(^b)</td>
<td>26.2 ± 2.2(^d)</td>
<td>22.6 ± 2.6(^d)</td>
</tr>
<tr>
<td>Group 6 (0.6 mg/g b.w. MSG + 0.2 mg/g b.w. V.E)</td>
<td>33.1 ± 1.9(^a)</td>
<td>6.4 ± 0.2(^a)</td>
<td>0.66 ± 0.02(^a)</td>
<td>30.9 ± 1.7(^b)</td>
<td>20.3 ± 1.4(^a)</td>
</tr>
<tr>
<td>Group 7 (1.6 mg/g b.w. MSG + 0.2 mg/g b.w. V.E)</td>
<td>34.2 ± 2.3(^d)</td>
<td>6.1 ± 0.1(^b)</td>
<td>0.67 ± 0.01(^a)</td>
<td>28.4 ± 3.1(^a)</td>
<td>21.2 ± 2.1(^a)</td>
</tr>
</tbody>
</table>

ALT = Alanine Amino Transferase, previously called Glutamate Pyruvate Amino Transferase (GPT). GGT = γ-glutamyltransferase. Mean ± SD. N = 8 for the control; n = 8 for treated group. Mean of treated group is compared with the corresponding mean of the control. \(P\) value of less than 0.05 indicates a significant difference between the compared means.

3.3.2. Serum Total Protein

The determined mean value of total protein in serum of control rats was 6.5 g/dl. Serum samples of treated animals contained total protein whose mean was 6.1 and 5.8 g/dl. The 6.2% and 10.8% reduction in serum total protein of MSG dosed rats (0.6, 1.6 mg/g body weight) failed to reach the statistical significance level (Table 2).

Additional of V.E and V.C resulted in a significant improvement in the serum total protein levels of lower MSG dosed rats (0.6 mg/g b.w.) (Table 2). Both vitamins have the ability to prevent oxidative stress which affects liver the main site for protein synthesis in the body.

3.3.3. Serum Total Bilirubin

It was found that there was a decrease (10.3%, 20.6%) in serum total bilirubin treated animals (Table 2). Control animals had a 0.68 mg total bilirubin/dl serum, whereas a mean values of 0.61 and 0.54 was recorded in serum of group 2 and 3 respectively. The difference between the compared groups was statistically significant (\(p \leq 0.05\)) (Table 2). Serum total bilirubin were found to be significantly increased in the serum after administration of vitamin C and vitamin E with MSG (Table 2).

3.3.4. Albumin

Albumin concentration decreased in the serum of MSG dosed rats. Tests performed in MSG-treated rats showed a significant decreased in serum albumin levels when compared to normal control group (Table 2). The synthetic function of liver was altered by MSG, so albumin level decreased. Albumin concentration increased in the serum of MSG administrated by additional of V.C and V.E dosed rats (Table 2).

The concentration of proteins, bilirubin and albumin in the serum can be used as indicators for the state of the liver and can be used to differentiate between different types of liver damage. The observed reduction in albumin and total bilirubin concentrations in serum (Table 2) indicated liver damage [55], arising from the uptake of the chemical compound. This may be an indication of diminished synthetic function of the liver which may consequently lead to enhanced retention of fluid in the tissues spaces [55].

3.3.5. GGT Activity

The activities of the serum enzyme markers of hepatocellular injury (GGT) increased significantly by MSG administration at concentration of 1.6 mg/g body weight. Vitamin C and Vitamin E when administrated with MSG reduced the activity of GGT (Table 2).

GGT is also elevated in most cases of liver disorders [56]. Therefore, the increased activity of GGT in the sera of MSG treated animals might have resulted from the liver injury caused by the MSG-induced oxidative stress. A similar trend was observed in the study of Onyema et al. [55].
3.4. Kidney Functions

3.4.1. Serum Urea

Blood serum samples of the rats treated with MSG contained low significant urea nitrogen as compared with urea nitrogen mean values of the control (Table 3). Mean serum urea of the control animals was 35.2 mg/dl and that of the MSG treated rats were 31.4 and 24.1 mg/dl of two MSG doses. Urea is the major nitrogen-containing metabolic product of protein catabolism. The significant reduction in serum urea concentration throughout the experimental period may be attributed to impairment of the urea cycle leading to reduced production of the metabolic product.

The serum urea nitrogen is a measure of renal function. Normally, the serum urea nitrogen level rises in heart failure, dehydration, or a high protein diet and low urea nitrogen level can be seen in liver and renal damage or in liver diseases [57].

3.4.2. Serum Creatinine

A change in serum creatinine level is also an indicator of kidney function. Therefore, in the present study serum creatinine was determined to find out the effect of MSG on renal functional markers such as serum creatinine, and serum urea nitrogen.

Concentration of creatinine in serum samples of control and treated animals were, respectively, 0.30, 0.44 and 0.54 mg/dl (Table 3). Serum creatinine of the treated rats was found to be more than that in control serum. The difference between means was significant (p ≤ 0.05).

The significant increase in creatinine content of the serum following the administration of MSG may be attributed to compromise of the renal functional capacity. MSG might have either interfered with creatinine metabolism leading to increased synthesis or the tissues might have compromised all or part of its functional capacity of tubular excretion [58-60]. Exposure to MSG may cause an adverse effect on the renal function which might be due to oxidative stress induced by MSG on the renal tissue.

Addition of V.E and V.C resulted in a significant improvement in the serum levels of biochemical parameters in kidneys (Table 3).

Farombli, 2006 [61] has shown that dietary antioxidants such as Vitamin C and Vitamin E has a modulator effects on MSG-induced serum urea oxidative damage in the liver and kidney of rats. The variation in the level of urea and creatinine are markers of renal dysfunction.

4. Conclusion

The results of the present investigation have shown that mono-sodium glutamate (MSG) at low doses is capable of producing alterations in the body weight and liver and kidney functions. These alterations appear in the liver and kidney probably because these organs are mainly responsible for detoxification of foreign compounds in the body. Vitamin C and Vitamin E have been shown to protect and restore the liver and kidney capabilities in several models injury via inhibiting oxidative damage.

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