

# Histological Studies Of The Effects Of Monosodium Glutamate On The Testis Of Adult Wistar Rats

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## Abstract

Histological effects of Monosodium glutamate (MSG) commonly used as food additive on the Testis of adult Wistar rats were carefully studied. The rats of both sexes (n=24), average weight of 185g were randomly assigned into two treatments (n=16) and control (n=8) groups. The rats in the treatment groups received 3g and 6g of MSG thoroughly mixed with their feeds for fourteen days, while the control rats received equal amounts of feeds without MSG added. The rats were fed with growers' mash purchased from Edo Feeds and Flour Mill Ltd, Ewu, Edo State and were given water liberally. The rats were sacrificed on day fifteen of the experiment. The testis was carefully dissected out and quickly fixed in 10% buffered formaldehyde for routine histological study after H&E method.

The histological findings after H&E methods indicated that the treated sections of the testis showed some cystic degenerative changes, with some vacuolations appearing in the stroma when compared to the control sections. There were atrophic and degenerative changes with the group that received 6g of MSG more severe. These findings indicate that MSG consumption may have a deleterious effect on the sertoli cells and leydig cells of the testis and may probably have adverse effects on spermatogenesis, spermiogenesis and testosterone production in adult male Wistar rats. This is in consonance with previous work done and reported by Onakewoh et al.

## INTRODUCTION

Monosodium glutamate (MSG) otherwise known as AJI-NO-MOTO is the sodium salt of glutamic acid. 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. Monosodium glutamate contains 78% of glutamic acid, 22% of sodium and water<sup>1</sup>. 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 such as meat, fish, milk and some vegetables<sup>2</sup>.

MSG is produced in many countries around the world through a fermentation process of molasses from sugar cane or sugar beets, as well as starch and corn sugar. In this fermentation process, bacteria, which may be genetically modified, are grown aerobically in a liquid nutrient medium. These bacteria have the ability to synthesize glutamic acid outside of their cell membranes and excrete it into the medium to accumulate there<sup>2,3</sup>. 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.

In Nigeria, most communities and individuals often use Monosodium glutamate as a bleaching agent for the removal of stains from clothes. There is a growing apprehension that its bleaching properties could be harmful or injurious to the body, or worse still inducing terminal diseases in consumers when ingested as a flavor enhancer in food. Despite evidence of negative consumer response to MSG, reputable international organizations and nutritionist have continued to endorse Monosodium glutamate, and reiterate that it has no adverse reactions in humans. The Directorate and Regulatory Affairs of Food and Drug Administration and Control (FDA & C) in Nigeria, now NAFDAC has also expressed the view that MSG is not injurious to health<sup>4</sup>. The Food and Drug Administration (FDA) of the United States reports that Monosodium glutamate is safe and that it should be maintained on the "Generally Recognized as Safe" (GRAS)-list of foods; being listed on food labels as a "Flavoring" or "hydrolyzed vegetable protein". MSG is thus reportedly permitted as a safe food additive that needs no specified average, daily intake or an upper limit intake requirement<sup>1</sup>.

Through its stimulation of the orosensory receptors and by improving the palatability of meals, MSG influences the

appetite positively, and induces weight gain<sup>5</sup>. Despite its taste stimulation and improved appetite enhancement, reports indicate that MSG is toxic to human and experimental animals<sup>6</sup>.

In 1968, the first published report of an adverse reaction to Monosodium glutamate appeared in the *New England Journal of Medicine* where it was reported that Monosodium glutamate was neurotoxic; killing brain cells, causing retinal degeneration, endocrine disorder and also associated with a number of pathological conditions such as addiction, stroke, epilepsy, brain trauma, neuropathic pain, schizophrenia, anxiety, depression, degenerative disorders such as Parkinson's disease, Alzheimer's disease, Huntington's disease, and amyotrophic lateral sclerosis<sup>1</sup>. The toxic effect of MSG was reported by the work done on the testis, causing significant oligozoospermia and increase abnormal sperm morphology in a dose-dependent fashion in male Wistar rats<sup>7</sup>. It has also been established that MSG may be implicated in cases of male infertility as it causes testicular hemorrhage, degeneration and alteration of sperm cell population and morphology<sup>8</sup>.

The testes are paired organs in males lying in the scrotal sac, and functions in the production of spermatozoa and secretion of male sex hormones, principally testosterone<sup>9</sup>. Since MSG causes significant oligozoospermia and increase abnormal sperm morphology in a dose-dependent fashion in male Wistar rats, it is relevant to investigate some of its histological effects on the testis. It is probable that the adverse effects of MSG may affect the normal physiological action of the testis and hence this investigation.

## **MATERIALS AND METHODS**

**ANIMALS:** Twenty four (24) adult male Wistar rats with average weight of 185g were randomly assigned into three groups A, B and C of (n=8) in each group. Groups A and B of (n=16) serves as treatments groups while Group C (n=8) is the control. The rats were obtained and maintained in the Animal Holdings of the Department of Anatomy, School of Basic Medical Sciences, University of Benin, Benin city, Nigeria. They were fed with growers' mash obtained from Edo feed and flour mill limited, Ewu, Edo state) and given water liberally. The rats gained maximum acclimatization before actual commencement of the experiment. The Monosodium glutamate (3g/ sachet containing 99+% of MSG) was obtained from Kersmond grocery stores, Uselu, Benin City.

**MONOSODIUM GLUTAMATE ADMINISTRATION:** The

rats in the treatment groups (A and B) were given 3g and 6g of MSG, thoroughly mixed with the growers' mash, respectively. The control © group received equal amount of feeds (growers' mash) without MSG added for fourteen days. The rats were sacrificed on the fifteenth day of the experiment. The testis was quickly dissected out and fixed in 10% buffered formaldehyde for routine histological techniques. The 3g and 6g MSG doses were chosen and extrapolated in this experiment based on the indiscriminate use here in Nigeria due to its palatability. The two doses were thoroughly mixed with fixed amount of feeds (550g) in each group, daily.

**HISTOLOGICAL STUDY:** The tissue were dehydrated in an ascending grade of alcohol (ethanol), cleared in xylene and embedded in paraffin wax. Serial sections of 6 microns thick were obtained using a rotatory microtome. The deparaffinized sections were stained routinely with haematoxyline and eosin. Photomicrographs of the desired sections were made for further observations.

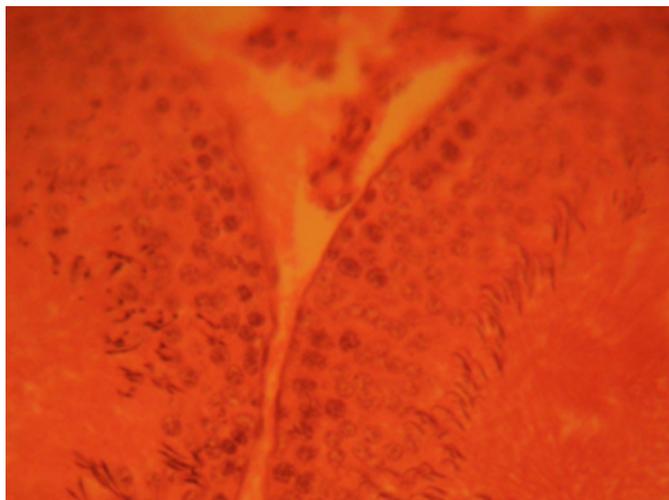
## **RESULTS**

The control sections of the testis showed normal histological features with the cross section of the convoluted seminiferous tubules showing stratified epithelium which consists of two distinct populations of cells; the spermatogenic cells and the sertoli cells. The leydig cells within the supporting tissues in the interstitial spaces between the tubules were all visible. (Figure 1).

The treatment sections of the testis showed some histological changes that were at variance with those obtained in the control. There was evidence of cystic degenerative changes, some intercellular vacuolations appearing in the stroma when compared to the control sections. There were atrophic and degenerative changes with the group that received 6g of MSG more severe (Figure 2 & 3).

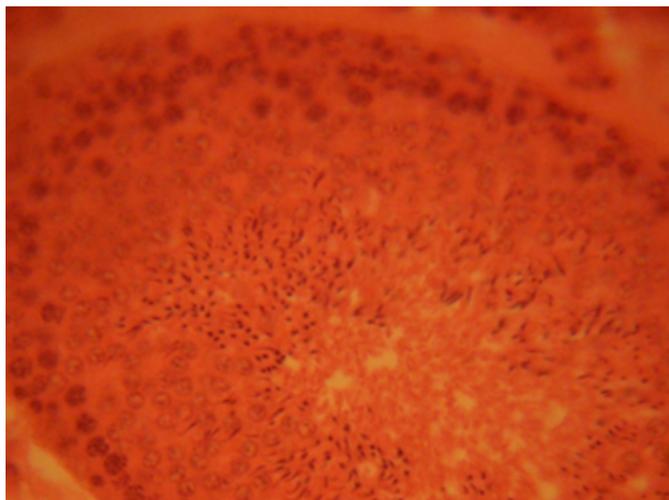
**Figure 1**

Figure 1: Control section of the testis (Seminiferous tubules) (Mag. x400).



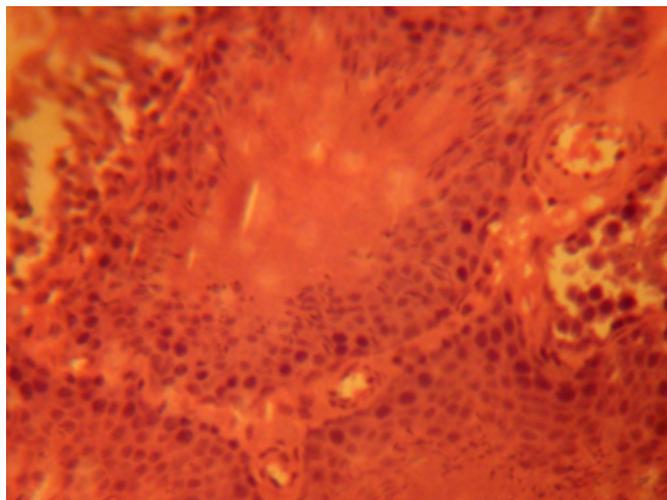
**Figure 2**

Figure 2: Treatment section of the testis (Seminiferous tubules) (3g MSG) (Mag. x400)



**Figure 3**

Figure 3: Treatment section of the testis (Seminiferous tubules) (6g MSG).(Mag. x400)



## DISCUSSION

The results (H & E) reactions showed cystic degenerative changes, with some intercellular vacuolations appearing in the stroma of the seminiferous tubules when compared to the control sections. There were atrophic and degenerative changes with the group that received 6g of MSG more severe. The vacuolations observed in the stroma of the treated testis may be due to MSG interference. The toxic effects of MSG on the testis observed in this experiment may underline the possible effects already reported <sup>7,8</sup>. The actual mechanism by which MSG induced cellular degenerative and atrophic changes observed in this experiment needs further investigation.

Degenerative changes have been reported to result in cell death, which is of two types, namely apoptotic and necrotic cell death. These two types differ morphologically and biochemically <sup>11</sup>. Pathological or accidental cell death is regarded as necrotic and could result from extrinsic insults to the cell such as osmotic, thermal, toxic and traumatic effects <sup>12</sup>. In this experiment MSG could have acted as toxins to the cells of the testis. The process of cellular necrosis involves disruption of membrane's structural and functional integrity which was also a landmark of this experiment. In cellular necrosis, the rate of progression depends on the severity of the environmental insults.

The greater the severity of insults, the more rapid the progression of neuronal injury <sup>13</sup>. The principle holds true for toxicological insults to the brain and other organs <sup>14</sup>. It may be inferred from the present results that prolonged intake and higher doses of MSG as been taken indiscriminately resulted

in increase toxic effects on the testis.

## **CONCLUSION AND RECOMMENDATION**

The results obtained in this study following the administration of 3g and 6g per day of MSG to adult male Wistar rats causes cystic degeneration, vacuolations and degenerative and atrophic changes, worse in the high dose. In inference these confirm earlier report that MSG may be a cause of male infertility. It is recommended that further studies be carried out to corroborate these findings.

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