

## HISTOLOGICAL AND HISTOPATHOLOGICAL CHANGES INDUCED BY DIMETHOATE (PESTICIDE) ON TESTES, KIDNEY AND LIVER OF MALE RABBITS IN YEMEN.

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### ABSTRACT

The present study deals with the changes induced by dimethoate pesticides, which commonly used in Yemen especially in khat plant and other vegetables and fruits. Twenty of adult male rabbits were divided into three groups, the first one was served as control group. The rest administered 1/20 (1.34 mg. Kg. b.wt.) and 1/10 (2.68 mg. kg. b.wt.) of LD<sub>50</sub> of pesticides respectively. All Animals were housed in separate cages under standard condition and stayed for two weeks before starting the experiments. The results revealed structural changes which more severe in rabbits given high doses of pesticide especially in liver and testes. The liver showed degenerative changes of hepatic cells and focal necrotic hepatocytes with lymphocytic infiltration and destructive of bile ducts lining epithelium in addition to congestion and widening of sinusoids and hepatic venules. The seminiferous tubules of the testes showed a considerable atrophy associated with marked reduction of spermatogonial cell number, partial failure of spermatogenesis was clearly detected. The spermatogonial cells revealed pyknotic nuclei, sertoli cells were detached from the basal lamina. The spermatids were undergoing several degenerative changes involved shrinkage or collapse of chromatin. The kidney tissue showed renal oedema and focal areas of tubular necrosis and high vacuolated cytoplasm.

### INTRODUCTION

The widespread use of pesticides all over the world for many years in plant production and public health programs has the main causes of serious environmental problems. These pollutants reach the human body and from animals in the daily diet and in the long term exposure those pollutants exert serious health hazards on the cells and tissues<sup>(1, 2)</sup>. The most important widespread and insecticides are four classes, organochlorines, carbamates, organophosphates and pyrethroids<sup>(3)</sup>.

These pesticides are usually mixed with water and sprayed on plants. They are usually introduced into the body of various biological systems by impregnation through the cuticle or skin, through inhalation or through ingestion. The acute toxicity of these pesticides in human beings is a serious health problem in numerous regions of the world, particularly in the developing countries, where more toxic organophosphate and carbamate insecticides are increasingly replacing the less toxic and more persistent organochlorine pesticides<sup>(4)</sup>. Dimethoate is one of the class of insecticides referred to as organophosphates, these chemicals act by interfering with the activities of cholinesterase an enzyme that is essential for the proper working of the nervous system of both humans and insects.

The present research designed to evaluate the possible toxicity of the dimethoate in some organs of rabbits like liver, testes and kidneys.

### MATERIALS AND METHODS

Twenty mature male rabbits were divided into four equal groups. All rabbits were housed and marked, then stayed for one week before starting the experiments. They fed on standard rodent pellets and vegetables as a source of vitamins.

Applied pesticide. Dimethoate is blue liquid organophosphorus pesticide (40% pure). The first group is control and received saline solution. The second, third, and fourth groups treated with 1/20 LD<sub>50</sub> of dimethoate (75 mg/kg. b.w.) 1/15 LD<sub>50</sub> (10.0 mg/kg. b.w.) and 1/10 LD<sub>50</sub> (15.0 mg/kg. b.w.).

The administration of all doses was received orally using stomach tube day after day (Ten doses were treated). Control and pesticide treated rabbits were sacrificed and rapidly dissected. The morphological features of the organs were recorded and tissue samples of liver, testes and kidneys were cut into small pieces, then fixed rapidly in neutral buffered formalin 10%. Then processed for paraffin embedding, histological sections of 7 micron thick were cut and stained with haematoxylin and eosin. They were then microscopically examined.

### RESULTS

**Liver:** The livers of the investigated control rabbit consisted of the classical hepatic lobules. The hepatocytes were arranged in cords radiating from the central veins with intervening blood sinusoids (figures 1 and 2). The hepatic cells or hepatocytes are polyhedral in shape, large in size and have well defined cell boundaries, one or two prominent nuclei are distinct between these cells certain phagocytic stellate cells known as Kupffer cells.

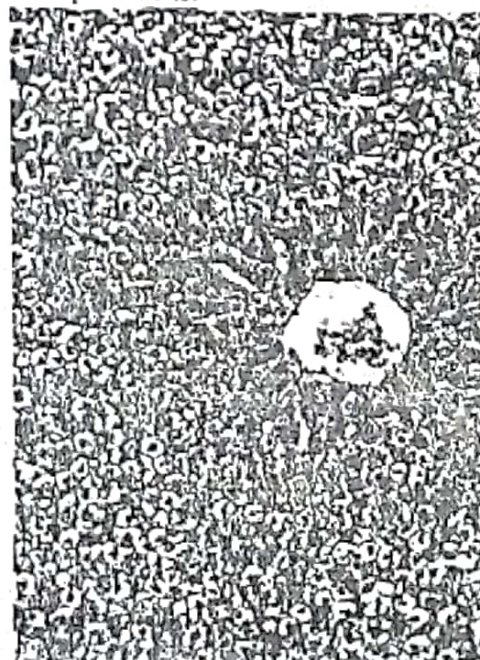


Fig. (1): Liver (Gr.1) showing hexagonal classic hepatic lobules. H&E Stain X 100

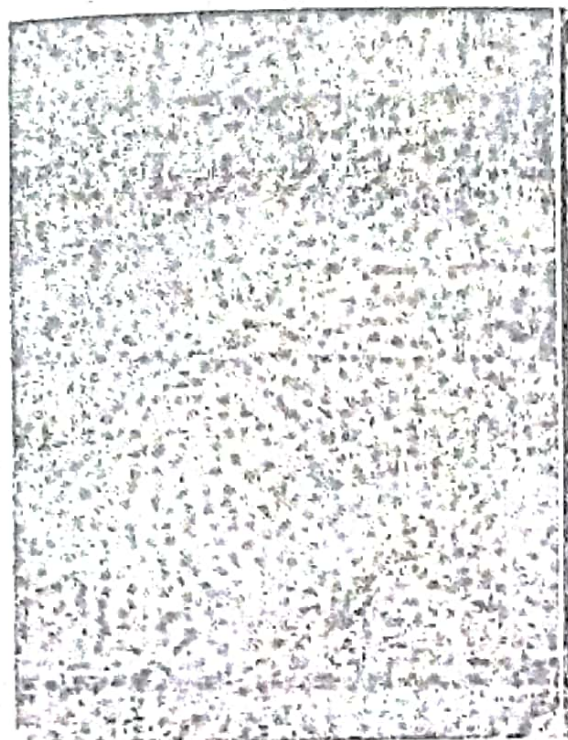


Fig. (2): Liver (Gr I) polygonal eosinophilic hepatocytes with spherical basophilic nuclei. H&E stain X 1000

#### Liver tissue of group II, III and IV:

The liver cells showed variable changes of degeneration depend on the dose, vascular degeneration the nuclei of the liver cell showed pyknosis, and karyolyses and disintegrate of the chromatin into several fragments (figure 3). Some of blood vessels appeared dilated and filled with blood cells. The wall of the blood vessels showed more thickening. The hepatic cell showed hydropic degeneration evidenced by vacuolation of their cytoplasm.

Non-Kupffer cells were prominent, there was venous congestion of hepatic parenchyma in which the sinusoids were mostly dilated and engorged with blood (figure 4).



Fig. (3) : Liver (Gr. II & III), showing congestion of portal area, prevascular oedema and hydropic degeneration H&E stain X 1000



Fig. (4): Liver. (Gr. IV). Showing pyknosis and karyorrhexis of hepatocytes H&E stain X 1200

**Kidney:** The control kidney (group I) showed normal cellular pattern and arranged of the renal glomeruli and tubules (figure 5). The Kidney showed number of uriniferous tubules which composed of nephron and collecting tubules. The malpighian corpuscles consisted of tuft of blood capillaries. The layers of glomerulus is formed of two layer, externally simple squamous epithelium rest on basement membrane. Internally consisted of flattened epithelial cell (figure 6).

Treated animals the kidney tissues showed more histological changes. According to the induced doses. The tissue showed extensive haemorrhagic areas and congested blood vessels in between the renal tubules of cortical and medullary regions of the kidney tissues. Degeneration of renal epithelium. Renal oedema and focal areas of tubular necrosis in the form of ill defined cell membranes and highly vacuolated cytoplasm (figures 7 and 8).



Fig. (5): Kidney: (Gr. I) showing renal cortex with renal corpuscles and proximal and distal convoluted tubules. H&E stain X 1000



Fig. (6): Kidney: (Gr. I): showing renal corpuscle and glomerular capillaries. H&E stain X 1000



Fig. (7): Kidney: (Gr. II, III). Showing necrosis of tubules of kidney. H&E stain X 1000



Fig. (8): Kidney: (Gr. IV) showing infiltration with lymphocytes and degeneration of epithelium lining of duct. H&E stain X 1000

Testis: In the control group the testis, the seminiferous tubules possessed well differentiated spermatogenic cells. The latter cells and spermatocytes were arranged in four layers. The sertoli cells were observed at regular intervals of testes lobules. Each seminiferous tubules showed of typical arrangement of spermatogenic cells, including spermatogonia, spermatocytes and spermatids (figures 9 and 10).

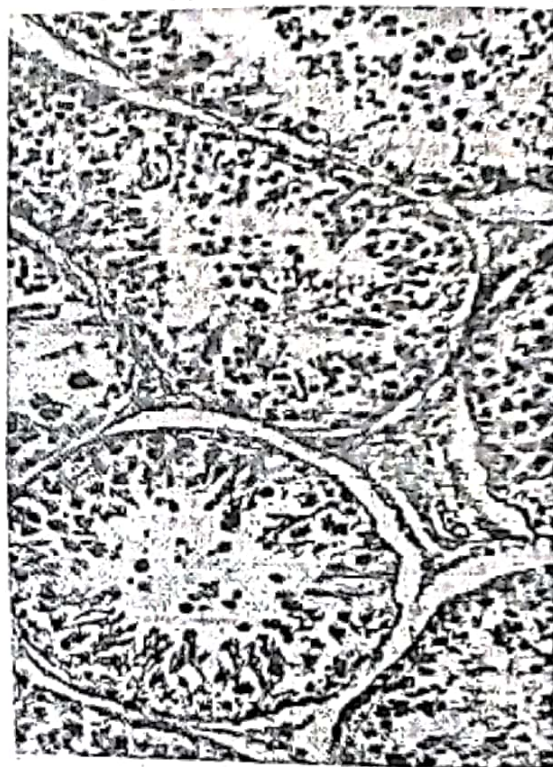


Fig. (9): Testis: (Gr. I). showing seminiferous tubules with spermatogenic cells in a normal state. H&E stain X 1000

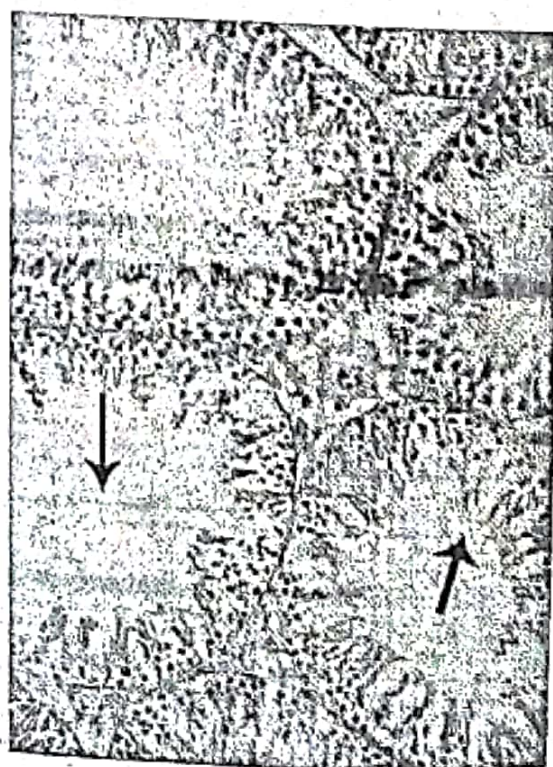


Fig. (10): Testis (Gr. I) showing testis lobules with typical arrangement of spermatogenic cells with sperm. H&E stain X 1000

**Treated testes:** They showed marked changes of the seminiferous tubules according to the dose there was marked reduction in the number of spermatogonia and spermatocytes. Many of these cells appeared lysed or degenerated. The basement membrane showed more thickening and stromal fibrosis (figure 11). The inter tubular connective tissues appeared loose and degenerated. The spermatogonia appeared markedly atrophied and degenerated and necrosis of the Sertoli cells and spermatocytes series (figure 12) detached from the basal lamina.



Fig. (11): Testis (Gr III) Showing pervascular and interstitial oedema H&E stain X 1200



Fig. (12): Testis (Gr IV) Showing congestion in interstitial blood vessel and haemorrhage & necrosis of tissue. H&E stain X 1200

## DISCUSSION

*Testis* is considered to be the most important organ in the reproduction system of males. It serves two roles the first is hormonal and the second is reproductive. The hormonal roles in values in the synthesis and secretion of the testosterone by the interstitial cells which found in groups between the seminiferous tubules. The second function involves spermatogenesis.

Our result revealed that there were degeneration and necroses of spermatocytes, there results agreed with that mentioned<sup>(3 and 4)</sup> by the toxicity uses of mercuric chloride in rats.

Also the some observation also recorded by the toxicity of midazolam<sup>(5)</sup> causes sever histological abnormalities of spermatogenesis. The same results were elucidated<sup>(6 and 7)</sup> recorded that nicotine has the same toxic effect on fine structure of the testis. Degeneration of human male germ cells was recorded<sup>(8)</sup> the same result recorded in treated rabbit<sup>(9)</sup> by light.

*Liver* is an organ of vital importance and it is considered to be the first organ in detoxification of any toxicants. Degenerative changes in the liver of animals administered excessive amount of Dime thoate have been reported<sup>(10)</sup>. Also, there were mononuclear cell and lymphatic cell infiltration in the parenchyma and portal area, these finding are in agreement with those reported<sup>(10, 11)</sup> in mercury intoxicated rabbit. The hepatocytes in animals under experiment suffered from fatty changes, vascular degeneration in addition to other areas of Coagulative necroses. The same result recorded<sup>(8)</sup>.

The results obtained in the present study showed that dimethoate induced hepatic focal areas cell hyperplasia accompanied by areas of liver cell necrosis, congested and dilated central veins and sinusoids were also observed<sup>(9)</sup> by barium intoxication and fluoride<sup>(10)</sup> intoxication.

*Kidney:* The present observation about the extensive haemorrhag and congested blood vessels between the renal tubules in cortical and medullary regions. The samillar results obtained<sup>(9)</sup> after injection with carbamate in kidney of adult male rats. Also reported on mice<sup>(9)</sup> after feeding with daily dose of lannate (25 g b. wt.) for 5010 days.

The same results also recorded<sup>(10)</sup> of female mice after induced of methomyl insecticide.

The damaging effects of organophosphorus compounds on malpighian corpuscles were noticed<sup>(9, 10)</sup> in mice, rabbits.

Observed tubular degeneration in kidney tissue of rat exposed to azadirachtin pesticides observed the same result of dimethaht in rat<sup>(11, 12)</sup>.

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## التغيرات النسيجية والمرضية بواسطة الدايمثويت (مبيد حشري) في الخصية، الكلي والكبد لذكور الأرناب في ذمار

باليمن

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تتناول هذه الدراسة تأثير أحد المبيدات العضوية الكاربامينية الذى يستخدم حاليا فى اليمن فى مقاومة الآفات الزراعية مثل زراعات القات والفاكهة والخضروات المنتشرة فى اليمن وهو الدايمثويت على الأرناب الذكور.

تم تحديد الإصابة على بعض الذكور من الأرناب فى مرحلة البلوغ وقد استخدم فى هذه الدراسة عدد ٢٠ أرناب وقسمت على أربعة مجموعات الأولى ضابطة بينما تمت معالجة الثانية والثالثة والرابعة بالجرعات المختلفة من المبيد (٢٠/١، ١٠/١، ٥/١) من الجرعة المميتة على الترتيب. تم اعطاء الجرعة لمدة ٤ جرعات على مدى ٨ أيام (يوم بعد يوم) عن طريق الفم باستخدام لى معوى وذبحت هذه المجموعات بعد اسبوع من نهاية الجرعات.

وقد اظهر الفحص النسيجى لكل من كبد - كلى وخصية هذه الارانب المعالجة بالجرعات المختلفة تأثير مباشر مثل احتقان فى الاوردة المركزية للكبد وقد تعرضت الخلايا الكبدية من مظاهر التحلل مصحوبة بوجود فجوات فى السيتوبلازم وتحلل فى انوية الخلايا وازدياد فى نشاط خلايا كوفر. أما الكلى ادت الى تحلل أنبوبات الكلوية داخل محفظة ملبيجى وكذلك ضمور فى محفظة ملبيجى وتحلل الأوردة والشريابين الدموية وكذلك التحلل المانى فى الخلايا الطلائية المبطنة لأنبيبات البولية للكلى.

أما الخصية التى حقنت بالجرعة المضاعفة فقد ظهرت بالنبيبات المنوية للخصية تغيرات نسيجية مرضية تمثلت فى تهدم وضمور كل من الخلايا الجرثومية وخلايا المنى الانبيبات وندرة الطلائع المنوية. وكذلك ظهور تجايف داخل الانبيبات التى ظهرت خالية من المكونات تماما.