

## Histopathological Effect of Potassium Peroxymonosulfate in Albino Mice Treated with Vitamin C

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### ABSTRACT

**Background** Disinfectants and surfactants are non-biodegradable contaminants that build up in bodies of water. Vegetation, planktons, fish, and zooplanktons absorb these disinfectants, which then enter the food web. Disinfectants introduced into aquatic systems accumulate in the food chain, causing a variety of harmful consequences and even mortality in aquatic creatures.

**Objectives** The present study aimed to evaluate the toxic effect of potassium peroxymonosulfate on the internal organs in albino mice.

**Methodology** :Eighty adult Swiss albino mice at the age of 7-8 weeks were divided into four groups (20 mice per group). 1st group was administered potassium peroxymonosulfate orally with dose (1449.7 mg/kg, b.wt) daily for eight weeks, 2nd group was administered vitamin C (250 mg/kg, b.wt) orally at the same time for eight weeks, 3rd group was administered potassium peroxymonosulfate (as same 1<sup>st</sup> group) and at the same time vitamin C as 2<sup>nd</sup> group orally for eight weeks, 4th group considered as negative control group. All of the animals were put to sleep, and portions of liver, kidney, intestine, heart, spleen, and brain were removed for histological analysis.

**Results:** our data reveal that the toxic effect of potassium peroxymonosulfate in the liver characterized by necrosis of hepatocytes with mitotic division nuclei, dilated central vein with mononuclear cells aggregation in portal area around portal vein and bile duct. Congestion of blood vessels with marked vacuolar degeneration of hepatocytes. The results reveal that vitamin C has an effect on the liver that is characterized by mononuclear cells aggregation around central vein and around portal area with proliferation of Kupfer cells. The microscopic section of the kidney showed mononuclear cells infiltration in the glomeruli and between renal tubules. The microscopic section of the lung showed mononuclear cells infiltration in the interalveolar septa. The microscopic section of the intestine hyperplasia of goblet cells and mild mononuclear cell infiltration in the lamina propria, and mucosal gland hyperplasia of lymphoid tissue. Results showed that the effect of potassium peroxymonosulfate and vitamin C in the kidney showed necrosis with acute cellular swelling of renal tubules and inflammatory cells in the lumen of renal tubules.

**Keywords:** peroxymonosulfate, vitamin C, liver, kidney, Brain

### INTRODUCTION

Disinfectants and cleaning agents prepared with potassium peroxymonosulfate have been shown to be more effective than chlorine-based disinfectants and cleaning agents against a variety of infectious microorganisms, including viruses and bacteria (Eleraky and Gasparini, 2002). and they have been used in the hygiene management of healthcare environments (Ogura and Doidge, 2015). This drug is particularly effective against a wide range of infectious pathogens. Furthermore, the US Environmental Protection Agency (EPA) claims that potassium peroxymonosulfate-based disinfectants (PPD) are efficient against norovirus, methicillin-resistant *Staphylococcus aureus* (MRSA), hepatitis B virus, and hepatitis C virus. (Environmental Protection Agency, 2014). Disinfectants and surfactants are non-biodegradable contaminants that build up in bodies of water. Vegetation, planktons, fish, and

zooplanktons absorb these disinfectants, which then enter the food web. Disinfectants introduced into aquatic systems accumulate in the food chain, causing a variety of harmful consequences and even mortality in aquatic creatures. ( Summarwar ,2013). Disinfectant use has resulted in a number of issues, including fungicide resistance and potentially detrimental consequences on human health (Phillips *et al.*, 2008).

As a result, innovative solutions that are successful in reducing mycotic diseases while still being safe for fish and the environment are still urgently needed. (Khosravi *et al.*,2012). Disinfection has a number of drawbacks, including the generation of potentially harmful disinfection by-products (DBPs) such trihalomethanes (THMs) and haloacetic acids (HAAs) 7. (2017, Bianco). The interaction of chlorine with natural organic materials produces this generation. Many THMs have been found as genotoxic mutagens that can be harmful to aquatic life and even people, with some of them being carcinogenic. (Ruales-Lonfat *et al.*, 2016). For the formation of sulfate, free radicals, peroxymonosulfate is a unique chemical oxidant. Potassium peroxymonosulfate, often known as oxone, is the most common salt form of peroxymonosulfate.

In terms of radical generation, activated peroxymonosulfate has recently been proven to be more efficient than hydrogen peroxide and persulfate (Rastogi *et al.*, 2009) .ROS have been linked to a wide range of diseases, including arthritis and connective tissue disorders, as well as carcinogenesis, aging, physical damage, infection, and acquired immunodeficiency syndrome (Kumar, 2013). oxidative stress is linked to pathological diseases that lead to cardiovascular problems, such as hypertension, hypercholesterolemia, and diabetes. (Kumar,2012). The predominant cytolytic and haematolytic activity of detergents (surfactants) is related to degradation of cell membrane integrity, mitochondrial activities, and cellular metabolism (Swenson and Ho-Yeon , 1992). The severity of the poisonous impact is determined by the agent's concentration and chemical properties, such as the length of the alkane chain and unsaturations (Swenson *et al.*, 1992).cute and chronic detergent toxicity has been studied in vitro (Menard *et al.*, 2011 ), ex vivo on tissues of experimental animals, and in vivo utilizing several animal models ( Johnson, 2004). Advanced oxidation techniques based on the formation of sulfate radicals have been shown to be successful in degrading a variety of resistant micro pollutants in aqueous solutions, including hormones.(Brienza,2014), Pharmaceuticals (Zhang, 2014), pesticides, and pool water decontamination (Anipsitakis, 2008) are only a few examples.

Vitamin C can be found in abundance in a variety of natural foods, including fresh fruits and vegetables. Vitamin C is a potent antioxidant with the potential to donate a hydrogen atom and generate the ascorbylfree radical, which is generally stable. Antioxidant vitamins such as vitamin E, vitamin C, and -carotene have been shown to reduce oxidative damage and the risk of certain chronic diseases (Rouhier *et al.*, 2008; Verma *et al.*, 2007). Vitamin C is a water-soluble antioxidant that has been shown to neutralize reactive oxygen species (ROS) and lessen oxidative stress (Rouhier *et al.*,2008; Verma *et al.*, 2007). Vitamin C is a powerful reducing agent and free radical scavenger in biological systems (Duarte *et al.*, 2005, Jasim *et al.*,2019 a).Vitamin C is a water-soluble antioxidant that has been shown to neutralize reactive oxygen species (ROS) and lessen oxidative stress (Rouhier *et al.*,2008; Verma *et al.*, 2007). Vitamin C is a powerful reducing agent and free radical scavenger in biological systems (Duarte *et al.*, 2005). It works as the initial line of defense against oxidative stress, protecting lipid membranes and proteins. Vitamin C, as a water soluble molecule, can neutralize free radicals and prevent free radical damage both within and outside the cells. Vitamin C is a great supply of electrons for free radicals looking for a way to re-establish their stability. Vitamin C can provide free radicals electrons and hence reduce their reactivity. (Rouhier *et al.*, 2008; Bindhumol *et al.*, 2003) .

## MATERIALS AND METHODS

Eigty mice aged 7-8 weeks and weighted ranged from 30-35gm,In this study, animals from the College of Vet. Med./University of Baghdad's animal house were used. They were housed and cared for in a traditional animal facility under strict temperature control (20±5°C). Pellets were

provided to the animals, as well as water. Each group of mice was kept in a plastic cage with hard-wood chip bedding for the duration of the study. To maintain a clean environment, the bedding was changed on a regular basis.

### **Preparation of potassium peroxy monosulfate**

Dissolve 1.4497 mg of potassium peroxy monosulfate in 10 ml distilled water to prepare concentration 14.497 mg/ml and to be given at dose volume of 0.1 ml/10 g B.W.

### **Preparation of vitamin C**

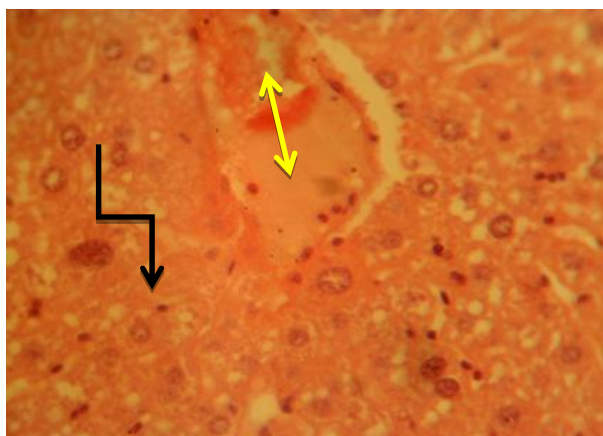
Dissolve 250 mg of vitamin C in 10 ml distilled water to prepare concentration 25 mg/ml and to be given at dose volume of 0.1 ml/10 g B.W.

### **Experimental design**

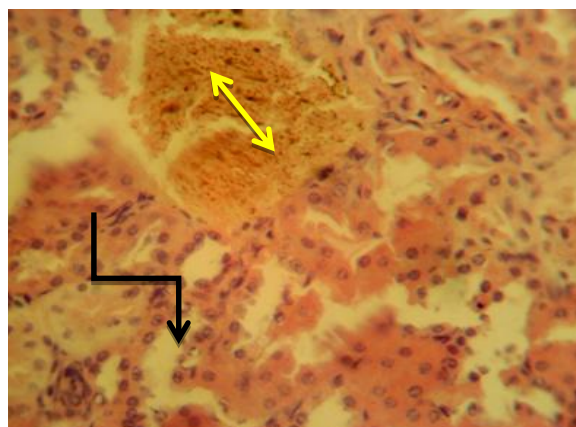
Eighty adult Swiss albino mice at the age of 7-8 weeks were divided into four groups. The 1st group (n=20) mice were administered orally with (1/10) 1449.7 mg/kg body weight of potassium peroxy monosulfate daily for 8 weeks. The 2nd group (n=20) mice were administered (250 mg/kg, b.wt) vitamin C daily for 8 weeks. The 3rd group (n=20) mice were treated in the same route with the vitamin C 250 mg/kg, b.wt./ daily for 8 weeks and the same time treated with (1/10) 1449.7 mg/kg body weight of potassium peroxy monosulfate. The 4th group (n=20) considered as control negative. Pieces of liver and kidney, intestine, spleen, heart and brain for standard histological investigation, they were fixed in 10% normal buffer formalin for 72 hours (Luna, 1968).

## **RESULT AND DISCUSSION**

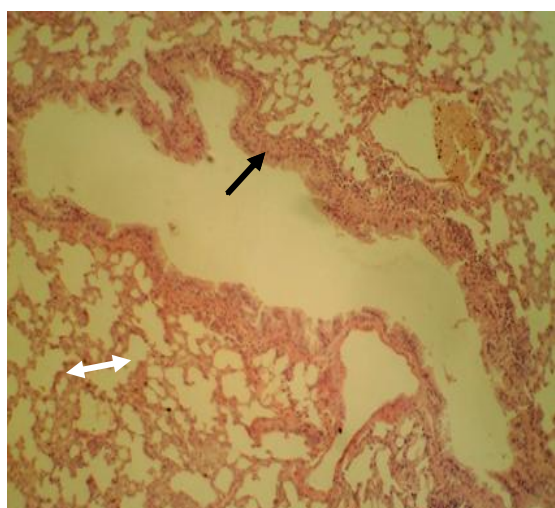
Histopathological changes of animals treated with potassium peroxy mono sulfate at 8 weeks post treatment the main lesion in liver characterized by shows edematous material in the dilated central vein with necrosis of hepatocyte (figure 1), also in kidney shows mononuclear cells infiltration and hemorrhagic area in addition to necrosis of renal tubules (figure 2), additionally in the shows focal lymphocytic aggregation with hyperplasia of epithelial cell of bronchi and increased thickness of interalveolar septa due to congested capillaries blood vessels in addition few RBCs (figure 3), also in intestine show hyperplasia epithelial cell of mucosal gland with increase mucin secretion with marked mononuclear cells infiltrations edematous material and sub mucosa and congestion of blood vessels (figure 4), also in alterations in the histopathology of animals treated with potassium peroxy mono sulfate and treated at same time with vitamin C at 8 weeks post treatment the main lesion in liver characterized by shows necrosis with vacuolar degeneration of hepatocyte and congestion central vein (in figure 5), also in kidney shows atrophy of glomeruli with necrosis and acute cellular swelling of renal tubules in addition to mononuclear cells between renal tubules and around glomerula with Bowman space, in addition to inflammatory cells in the lumen of renal tubules (in figure 6), also in lung shows hyperplasia of the epithelial cells of bronchiole with increased thickness of interalveolar connective tissue in addition to infiltration of the inflammatory cells (in figure 7), also in intestine show hyperplasia of lymphoid tissue and congested of blood vessels (in figure 8). Also in the animals after administered vitamin C, the main lesion in liver shows mononuclear cells aggregation around portal area with proliferation of Kupfer cells (in figure 9), also in kidney shows mononuclear cells infiltration in the glomeruli and between renal tubules (in figure 10). Histopathological changes of animals 4th group that considered control (negative group) No important microscopic findings have been found.



**(Figure1):**Histopathological section in the liver of animal at 8 weeks post-treatment with potassium peroxymonosulfate (1.4497mg/kg b.w ) shows edematous material in the dilated central vein(yellow arrow) with necrosis of hepatocyte(black arrow) (H&E stain 40X)



**(Figure 2):**Histopathological section in the kidney of animal at 8 weeks post-treatment with potassium peroxymonosulfate (1.4497mg/kg b.w ) shows mononuclear cells infiltration and hemorrhagic area(yellow arrow) in addition to necrosis (black arrow) of renal tubules (H&E stain 40X).

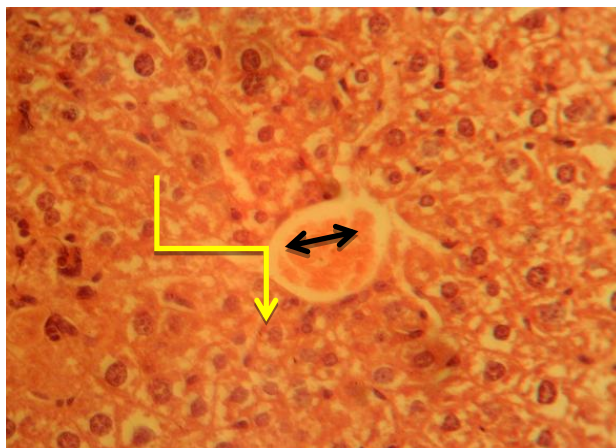


**(Figure 3):**Histopathological section in the lung of animal at 8 weeks post-treatment with potassium peroxymonosulfate (1.4497mg/kg b.w ) shows focal lymphocytic aggregation with hyperplasia of epithelial cell of bronchi(black arrow) and increased thickness of interalveolar septa(white arrow) due to congested capillaries blood vessels in addition few RBCs (H&E stain 10X).

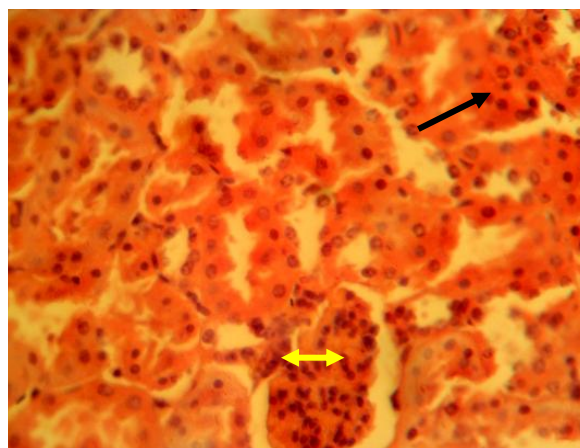


**(Figure 4):**Histopathological section in the intestine of animal at 8 weeks post-treatment with potassium peroxymonosulfate(1.4497mg/kg b.w ) show hyperplasia epithelial cell of mucosal gland with increase mucin secretion(white arrow) with marked mononuclear cells infiltrations edematous material and sub (black arrow)mucosa and congestion of blood vessels (yellow arrow) (H&E stain 40X).

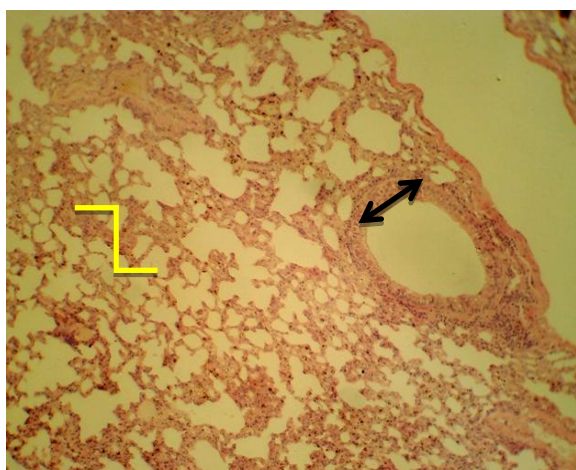




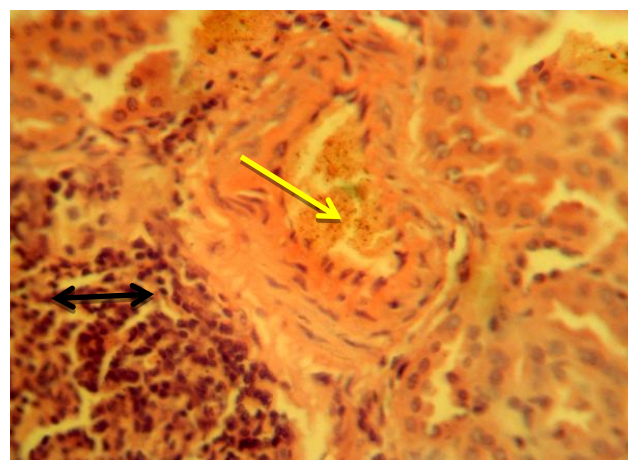
**(Figure 5):** Histopathological section in the liver of animal at 8 weeks post-treatment with potassium peroxymonosulfate and vitamin c shows necrosis with vacuolar degeneration of hepatocyte (yellow arrow) and congestion central vein (black arrow) (H&E stain 40X)



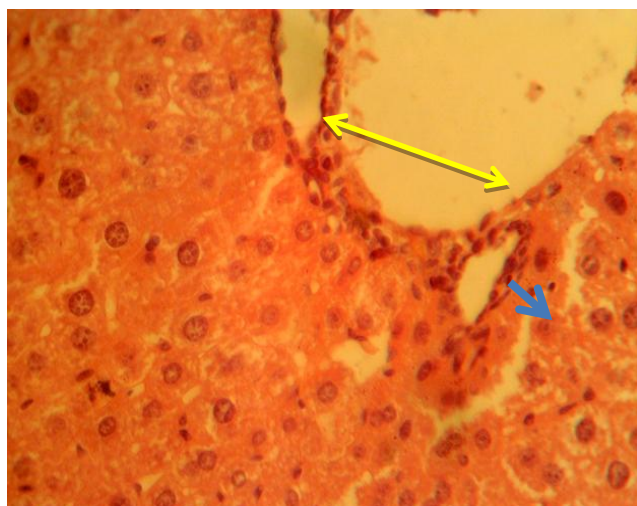
**(Figure 6):** Histopathological section in the kidney of animal at 8 weeks post-treatment with potassium peroxymonosulfate and vitamin c shows atrophy of glomeruli with necrosis and acute cellular swelling of renal tubules in addition to mononuclear cells between renal tubules and around glomerula with Bowman space (yellow arrow), in addition to inflammatory cells in the lumen of renal tubules (black arrow) (H&E stain 40X)



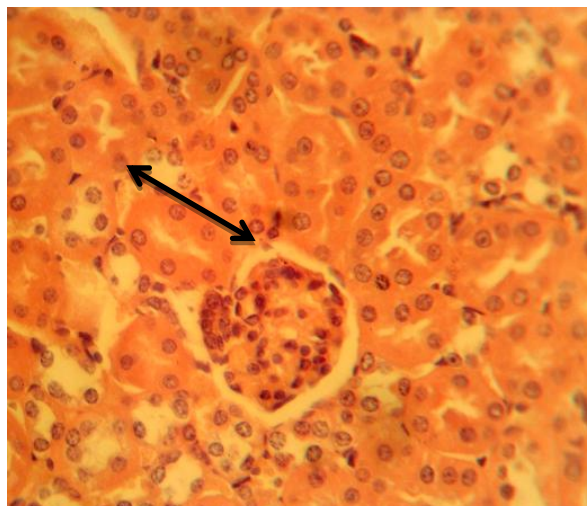
**(Figure 7):** Histopathological section in the lung of animal at 8 weeks post-treatment with potassium peroxymonosulfate and vitamin c shows hyperplasia of the epithelial cells of bronchiolus with increased thickness of interalveolar connective tissue (black arrow) in addition to infiltration of the inflammatory cells (yellow arrow) (H&E stain 10X)



**(Figure 8):** Histopathological section in the intestine of animal at 8 weeks post-treatment with potassium peroxymonosulfate and vitamin c show hyperplasia of lymphoid tissue (black arrow) and congested blood vessels (yellow arrow) (H&E stain 40X)



(Figure 9):Histopathological section in the liver of animal at 8 weeks post-treatment with vitamin c shows mononuclear cells aggregation around portal area(yellow arrow) with proliferation of Kupfer cells(blue arrow) (H&E stain 40X)



(Figure 10):Histopathological section in the kidney of animal at 8 weeks post-treatment with vitamin c shows mononuclear cells infiltration in the glomeruli and between renal tubules(black arrow) (H&E stain 40X)

Detergents, including biodegradable detergents, have been found to cause poisoning and osmoregulatory imbalances in aquatic life, particularly when present in concentrations that exceed metabolic requirement. Such xenobiotic chemicals are known to be one of the most common terrestrial and aquatic pollutants because they are persistent and mobile in soil and water.(Cox, 1998). The gills, kidney, liver, skin, heart, and brain have all been observed to be severely damaged by the detergent effluents. The goal of this study is to figure out how much oxidative and neurological damage synthetic detergent SDS causes in the brain. In order to stabilize the organism under stress, bodily processes are adjusted to resist the influence of pollutants/stressors. Due to the functional deterioration of antioxidants, oxidative stress is a critical expression of a multi-step system that culminates in an imbalance between pro-oxidant and antioxidant defense mechanisms., or an excessive buildup of superoxide radicals, or both, resulting in tissue injury ( Kavitha and Rao, 2009) . Exposure to certain pollutants, such as SDS, has been shown to produce reactive oxygen species (ROS), which can induce organ damage ( Livingstone,2001). The levels of oxidative stress were measured using MDA, a lipid peroxidation marker. Although the effect did not depend on the duration of the exposure, the considerable increase in MDA level indicated the production of super oxide radicals in fish exposed to SDS. The SOD-CAT system works by inhibiting the generation of oxy-radicals.decrease in the activity of the SOD-CAT system was most likely a reaction to toxicant stress, and failed to offset the negative effects of enhanced super oxide molecule production (Pandey *et al.*, 2003). (John *et al.*,2001). Long-term toxicant exposure hindered SOD and CAT activities in catfish tissues, which could be related to a flux of superoxide radicals resulting in an increase in cellular peroxide molecule (Zhang *et al.*,2008). GSH plays an important role in cellular antioxidant defense and metabolic pathway adjustment activities., GST decreased the amounts of xenobiotic compounds in the aquatic system, while GPx catalyzes H<sub>2</sub>O<sub>2</sub> reduction and protects against lipid peroxidation ( Li ZH *et al.*, 2010). They all work together to provide a robust line of defense against oxidative stress caused by xenobiotic contaminants ( Basha *et al.*,2012).

In another investigation, kidney impairment caused by detergent-processed potassium peroxydisulfate therapy was examined by assessing urine indicators of renal function. Renal dysfunction is linked to changes in these biochemical markers. Increased plasma concentrations of these products would come from renal disorders that produced a decline in GFR (George *et al.*,2020). As a result, as the filtration rate decreases, the plasma concentrations of urea and creatinine rise. The treated groups had considerably greater urea levels than the positive and negative control groups in this study.This is consistent with (Shama and Wasma's ,

2011) work, which found a substantial increase in plasma urea in rats fed a methanol extract of cassava for two weeks. When compared to the control rats, the detergent treated rats had considerably higher urine concentrations of these excretory products. The detergent-treated animals' renal photomicrographs revealed a disturbed cortical structure with diffuse tubule loss, an inflammatory cell infiltrate in the interstitium, and some intact glomeruli.

This suggests that the higher plasma urea in the potassium peroxydisulfate-treated groups is due to enhanced urea secretion by renal cells and/or higher urea production by the kidneys. Because urea is one of their consequences, skeletal muscle metabolism and protein catabolism are linked. These findings are consistent with those of (Fakhouri and Fremaux-Bacchi, 2007), who stated that renal damage was most likely caused by the direct toxic action of detergent components (especially borate) on renal tubular epithelial cells and endothelium, which was exacerbated by hypovolaemia and the resulting reduction in renal toxin excretion. The severe detergent-related endothelial injury also caused thrombotic microangiopathy, with low schistocytes on peripheral smear and low platelets, which further complicated the picture, especially given the history of hamburger consumption. The present finding revealed different pathological lesions in the intestine, may be due to cytotoxic effect of detergent these result agreement with (Ogunbileje and Akinosun, 2011), who reported the detergent solution which caused multi organs injuries. Intestinal metaplasia is associated with the transformation of the stomach lining and in the earlier stages, the replaced epithelium resembles the small intestine (Madhulima, 2016). The pathological lesions that were recorded in the intestine may be due to the direct effects of potassium peroxydisulfate on intestinal mucosa which lead to stimulate goblet cells activity to produce large amount of mucin in order to remove or diluted the toxic effects of potassium peroxydisulfate. Because the function of villi is so important, and the damage to this site in this situation is so severe, the extent of gastrointestinal disorders may easily be attributed to detergents. Surf appears to have obstructed the paracellular channels generated at the connections between intestinal cells for the passage of ions and other water soluble substances to the lamina propria. (Ogochukwu and Joseph, 2009). Maciorowski (Travlos et al., 1996) also found that the effect of anionic surfactants on mollusk intestine injury was reversible. Treatment of animals with vitamin C In the current investigation, modest damages were observed in the internal organs of rats given vitamin C; these findings could imply that vitamin C's antioxidant activity protects soft tissues from the detrimental effects of free radicals created as a result of heavy salt poisoning (Jasimet al., 2019 b). The alpha tocopheroxyl radical, which is formed when exogenous radical oxidants interact with alpha tocopherol, is an example. Low-density lipoprotein cholesterol (LDL) cholesterol (LDL) cholesterol (LDL) cholesterol (LDL). The tocopheroxyl radical has the ability to be converted to alpha tocopherol by ascorbate (Neuzilet al., 1997). Reactions between iron and copper that are mediated by transition metals. For example, Fenton chemistry can lead to the creation of additional radicals when iron is reduced by ascorbate (Carr and Frei, 1999). Reduced iron, on the other hand, may be an endpoint reaction: for example, reduced iron may be the optimal state for intestinal absorption (Hallberg, 1995; Lynch, 1997).

## Acknowledgement

extend my sincere thanks, appreciation and respect to the College of Veterinary Medicine, Baghdad University for their continuous support, and I thank the physiology and toxicology department for providing full support and conducting the selected tests inside the branch.

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