Original article

Ginseng Panax and Its Impact on Mitigating the Negative Effects of Tin Chloride on Kidney Function and Serum Proteins in Male Rabbits

Salma Hamed¹*^D, Marfoua Ali¹^D, Fayrouz Khaled²

¹Department of Zoology, Faculty of Science, Omar Al-Mokhtar University, El-Beida, Libya ²Department of Chemistry, Faculty of Science, Omar Al-Mukhtar University, El-Beida-Libya **Corresponding Email.** <u>salma.hamed@omu.edu.ly</u>

Abstract

The existing investigation aimed to locate *ginseng*'s ability to resist the harmful effects of tin chloride "SnCl₂" on the kidneys and serum proteins. Male rabbits were used in this investigation. The number of study groups was 4, each containing 5 rabbits, so the total number of rabbits used was 20. For twelve weeks, the animals underwent oral therapy per day. The experiment was designed as follows: The "Control group" (group 1) rabbits were given corn oil only. "*Ginseng* group" (group 2) rabbits were treated only with "100 mg/kg body weight" of *ginseng*. "Tin chloride (SnCl₂) group" (group 3) rabbits were treated with "20 mg/kg body weight" of SnCl₂ only. Finally, the "*Ginseng* + SnCl₂ group" (group 4) rabbits were treated with SnCl₂ and *ginseng* together. The outcomes displayed a substantial rise in the "*Ginseng* group" rabbit in the following serum parameters: "total protein (TP), albumin (A), and globulin (G)". A substantial "P<0.05" decrease of the "urea and creatinine." The exact opposite occurred in the "SnCl₂ group", where serum proteins "TP, A, and G" decreased substantially, and "urea and creatinine" levels increased markedly. *Ginseng* minimizes the harmful effects of SnCl₂ on all tested parameters, and this was demonstrated in the "*Ginseng* + SnCl₂ group". In conclusion, *Panax ginseng* has an important role in reducing the harmful effects of SnCl₂.

Keywords: Male Rabbit Kidney, Tin Chloride, Panax Ginseng, Urea, Creatinine.

Introduction

"Tin (Sn)" is a heavy metal found in the crust of the earth. Its average concentration is about 2 milligrams per kilogram [1]. Tin is widely used in our daily lives, in lining beverage and food cans, in container plating, and in alloys such as pewter and bronze. In the manufacture of rodenticides and insecticides, plastic pipes, and paints [1]. Inorganic components of tin, such as "tin oxide, tin chloride, and tin sulfide," are formed by the interaction of tin with oxygen, chlorine, and sulfur. Tin is also utilized in the manufacture of colors, perfumes, toothpaste, and soap. Organic tin compounds, "triphenyltin, tributyltin, and dibutyltin," are formed when tin interacts with carbon. Organic forms of tin are not found in the environment but are often synthetic materials [1]. Inorganic tin is present in the bones and other parts of the body, including the skin, kidneys, lymph nodes, tongue, lungs, liver, and spleen. Nephrotoxicity, neurotoxicity, genotoxicity, low calcium in bone, and immunotoxicity are among the serious effects of inorganic tin on the body [2,3]. Excess tin can cause anemia, in addition to the risks posed by tin on the kidneys [4].

Stannous ions are poisonous, and it seems that this toxicity is mediated by the formation of "reactive oxygen species (ROS)". Rabbit's kidneys are damaged by prolonged exposure to tin chloride [5]. To appreciate kidney function, serum "urea, and creatinine are used as biomarkers. Creatinine phosphate metabolism in muscles produces creatinine, whereas the liver generates "urea" from amino acids [6]. Rising "urea and creatinine" are kidney damage indicators. The kidney's filtration system deteriorates as kidney cells collapse, leading to elevated "urea and creatinine" levels [7]. "Urea and creatinine" parameters are used to estimate the results [8]. The usage of herbs and their products is widely spread due to their perceived efficacy and lack of negative effects [9]. Among these herbs, *P. ginseng* has a lengthy history and is today one of the most widely used medicinal plants in the world [10]. *Ginseng*'s pharmacological effects are mainly caused by its saponins, also known as ginsenosides, the plant's principal and bioactive components [11]. Most of the actions of *ginseng*, such as its anti-inflammatory, anti-cancer, antioxidant, and vasorelaxation qualities, seem to be attributed to its saponins [12]. According to [13], ginseng boosts the antioxidant defense system [14], stating that *ginseng* could reduce the negative effects of tin chloride (SnCl₂) on the kidney function parameters and serum protein concentrations of male rabbits, the current investigation was conducted.

Methods

Chemicals

Tin chloride $"SnCl_2"$ (1 mg/L) was obtained from the Chemistry Department of the Faculty of Science. *Panax ginseng* powder was obtained from a local pharmacy, where 7 grams were dissolved in 42 ml of normal saline.

Animal Management

20 male rabbits were used, weighing 1.641 ± 27.2 kg and 6 months old. Water and food were available ad libitum throughout the three-month experiment. Every animal was housed in a different cage and weighed once a week.

Experiment Design

The experiment included 4 equal-sized groups of rabbits, each containing 5 rabbits; the "first group" served as a "control group," where rabbits were given only one milliliter of corn oil for three months, twice a week, via oral gavage. In the "second group," the rabbits were given *Panax ginseng* alone at an oral dose of 0.01 g/kg [15]. The "third group" also received orally 20 mg/kg/day of SnCl₂ alone. The "fourth group" received a mixture of *ginseng* + SnCl₂ orally.

Biochemical tests and enzyme activity

After being drawn, after "20 min" at "25°C", the blood clotted in a plain tube. All of the following were classified as either normal or negative. A refractometer was used to measure specific gravity. On the day of sample collection, all results were noted as either positive or negative and compared to the medical record. By centrifugation at "860×g" for "20 min," extract the serum, which in turn was kept until needed at "-20°C." Kidney function parameters were estimated using an automated analyzer and a kit approach.

Statistical analysis

Conducted a data analysis using "Graph Prism Pad and Minitab version 17" and used "ANOVA analysis and Tukey multiple comparison tests" to locate statistically significant differences. Which was estimated to be "P < 0.05" [16].

Results

The mean values of serum "total protein (TP), albumin (A), and globulin (G)" are observed in "Table 1 and Figures 1-3." When administered *P. ginseng* alone, "TP, A, and G" levels increased substantially. However, in the SnCl₂-administered group, the opposite occurred; "TP, A, and G" levels were lower than in the control group. *Ginseng* mitigated the effects of SnCl₂ in the *ginseng* + SnCl₂ group, as the measured parameters were close to those in the control group.

In "Table 2 and Figures 4 and 5", it was observed that the mean values of "urea and creatinine" in male rabbits were lower for "the *ginseng* group" when contrasted to "the control group," and the opposite was observed in "the SnCl₂ group," where levels of "urea and creatinine" were substantially higher contrasted to "the control group. In "the *ginseng* + SnCl₂ group," levels of both "urea and creatinine" were close to those "the control group," due to the role played by *ginseng* in reducing the harmful effects of SnCl₂.

remedied groups							
Experimental groups							
Parameter	Control Mean± SEM	Ginseng Mean± SEM	SnCl ₂ Mean± SEM	Ginseng +SnCl ₂ Mean± SEM			
Globulin (U/L)	2.82±0.126ª	2.48 ± 0.162^{a}	2.81 ± 0.154^{a}	2.80 ± 0.099^{a}			
Albumin (U/L)	3.92 ± 0.060^{b}	4.43±0.088ª	3.56 ± 0.109^{b}	3.99 ± 0.076^{b}			
T. Protein (U/L)	6.74 ± 0.104^{ab}	6.91 ± 0.121 a	$6.36\pm0.128^{\rm b}$	$6.79\pm0.091a^{\rm b}$			

Table 1. States values of serum	"total protein, albumin,	and globulin" of rabbit males in all					
remedied groups							

Each treatment group has n = 5; values are presented as means \pm SEM. Different "superscript letters a, b, and c" within columns differed substantially ("p<0.05").



Figure 1. "Globulin Values (mean± SEM)" every two weeks of rabbit males in all study groups.

Ginseng SnCl₂ Sn+*Gin*

https://doi.org/10.54361/ajmas.258210



Figure 2. "Albumin values (mean± SEM)" every two weeks of rabbit males in all study groups.



Figure 3. "Total protein values (mean± SEM)" every two weeks in rabbit males in all study groups.

Table 2. States values of "urea and creatinine" of rabbit males in all remedied groups.

Experimental groups							
Parameter	Control Mean± SEM	Ginseng Mean± SEM	SnCl ₂ Mean± SEM	Ginseng +SnCl ₂ Mean± SEM			
Urea (mg/dl)	38.50 ± 0.512^{b}	35.82±1.075 ^b	46.98±1.816 ^a	39.19±1.102 ^b			
Creatinine (g/dl)	0.76 ± 0.042^{b}	0.59 ± 0.021^{b}	1.20 ± 0.084^{a}	0.76 ± 0.042^{b}			

Each treatment group has n = 5; values are presented as means \pm SEM. Different "superscript letters a, b, and c" within columns differed substantially ("p < 0.05").



Weeks

Figure 4. "Urea values (mean± SEM)" every two weeks of rabbit males in all study groups.

https://doi.org/10.54361/ajmas.258210



Figure 5. "Creatinine values (mean± SEM)" every two weeks of rabbit males in all study groups.

Discussion

In the current investigation, it was found that when male rabbits were given only ginseng, levels of the "total protein, albumin, and globulin" in serum increased, whereas levels of these parameters decreased when the rabbits were treated with $SnCl_2$ compared to "the control group". This concurs with what [17] found. Levels of "total protein, albumin, and globulin" were adjacent to values of "the control group" in the group administered with $ginseng + SnCl_2$.

The decrease in plasma "total protein" from SnCl₂ treatment was primarily due to lower "albumin and globulin" levels. For animals exposed to a polluted environment, low plasma proteins are due to changes in the liver's generation of free amino acids and proteins, or changes in how these substances are metabolized [17,18]. [17,19] found that when male rabbits were given only *ginseng*, the major cause of decreased blood protein was also been attributed to the loss of too much due to nephrosis. It might also arise from "reduced protein synthesis, increased proteolytic activity, or degradation". Liver cells may be damaged by SnCl₂, which partially leads to a significant decrease in blood proteins.

Regarding kidney function, "urea and creatinine" levels decreased in male rabbits administered with *ginseng* alone. Conversely, a substantial increase in "urea and creatinine" levels was observed when rabbits were given SnCl₂. *Ginseng* reduced the harmful effects of SnCl₂ in the group receiving both *ginseng* and SnCl₂, bringing them close to control values.

If the renal tubules are damaged or the pressure or volume of blood flowing through them decreases, as in nephritis, the renal tubules lose their ability to rid the blood of "urea" and other waste products, and the level of "urea" in the blood increases. Therefore, high blood "urea" levels are considered one of the most important indexes of impaired kidney function in rabbits administered with SnCl₂ [17,20]. According to [21], Bowman's capsule and the cells lining the renal convoluted tubules were changed by SnCl₂. Failure of the liver and the drop in proteins of the plasma support the idea that protein breakdown produces urea. As a result of the liver and its functions being affected by SnCl₂, increased urea concentrations occur in the plasma of animals given SnCl₂.

Conclusion

The harm to the kidneys, the swelling of the renal tubular epithelium, and the proximal tubule degeneration represent harmful effects of increasing the concentration of tin compounds in the human body and the animal, and ginseng can be used as a dietary supplement to improve them.

Conflicts of Interest

The authors declared no conflict of interest.

References

- 1. Amir R, Alam MT, Bari B, Sehar N, Siddiqi ZH, Khatoon A, et al. Qalaee (Stannum) metallic origin element, its chemical compounds, action, therapeutic uses, adverse effect, and pharmacological activity studies: A Review. J Drug Deliv Ther. 2024;14(7):136-41.
- 2. Antero A, Mirja K, Tiina S, Monica N. Gold and Gold Mining. In: Nordberg GF, Fowler BA, Nordberg M, editors. Handbook on the Toxicology of Metals. 4th ed. Amsterdam: Academic Press; 2015. p. 817-43.
- 3. Paul H, Peter W. Tin and inorganic tin compounds. Geneva: World Health Organization; [date unknown].

- 4. Carolyn H. Toxicological profile for tin and tin compounds. Atlanta, GA: Agency for Toxic Substances and Disease Registry; 2005 Aug.
- 5. Fatma M. EL. Antioxidant effect of vitamin E and selenium on lipid peroxidation, enzyme activities and biochemical parameters in rats exposed to aluminum. J Trace Elem Med Biol. 2004;18(2-3):113-21.
- 6. Putu OA, I Nyoman S, I Gusti AAS. The influence of peel extract Musa paradisiaca formatypica against kidney histology, urea levels, and creatinine Rattus novergicus after intensive exercise. Bali Vet J. 2022;14(5):578-85.
- 7. Sarah W, Maggie L. Handbook of laboratory animal Management and welfare. 4th ed. West Sussex: Wiley-Blackwell; 2013.
- Andi E, Irma S, Sasmita CDB, Amaliah A, Dewi Y. Nephrotoxicity effect of Ginseng Bugis (Talinum paniculatum (Jacq) Gaertn) leaves ethanolic extract on creatinine, urea, and kidney histopathological features. Int Food Res J. 2024;31(2):417-22.
- 9. Peter CK, Liew S. Traditional Chinese herbal medicine and anaesthesia. Anaesthesia. 2002;57(11):1083-9.
- 10. Michael KAL, Jonathan M, Chun-su Y. Herbal medicines and perioperative care. JAMA. 2001;286(2):208-16.
- Nguyen HT, Takuhiro U, Osamu M, Young HK, Yukihiro S. Pharmacological Effects of Ginseng on Liver Functions and Diseases: A Minireview. Evid Based Complement Alternat Med. 2012; 2012:7.
- 12. Lorne JH, Michael JW. Inflammation, cancer, and targets of ginseng. J Nutr. 2007;137(1 Suppl):183S-5S.
- 13. Hyeong GK, Sa RY, Hye JP, Nam HL, Jang WS, Rekha S, et al. Antioxidant effects of Panax ginseng C.A. Meyer in healthy subjects: a randomized, placebo-controlled clinical trial. Food Chem Toxicol. 2011;49(9):2229-35.
- 14. David DK, Arosha NW, Chun H. Antioxidant properties of a North American ginseng extract. Mol Cell Biochem. 2000;203(1-2):1-10.
- 15. Inoue M, Wu C, Dou D, Chen Y, Ogihara Y. Lipoprotein lipase activation by red ginseng saponins in hyperlipidemia model animals. Phytomedicine. 1999;6(4):257-65.
- 16. Daryl SP. Biostatistics and microbiology: a survival manual. 1st ed. New York: Springer; 2008. 216 p.
- 17. Yousef MI, Awad TI, Elhag FA, Khaled FA. Study of the protective effect of ascorbic acid against the toxicity of stannous chloride on oxidative damage, antioxidant enzymes and biochemical parameters in rabbits. Toxicology. 2007;235(3):194-202.
- 18. Viviana R, Hector B. Effects of 2,4-dichlorophenoxyacetic acid on polyamine synthesis in Chinese hamster ovary cells. Toxicol Lett. 1991;56(1-2):151-7.
- 19. Abdul Rauf S, Aziz F, Alam J, Syed SA. Toxic effects of talstar, a new synthetic pyrethroid, on blood and liver of rabbit. Pak J Zool. 1990;23(4):289-300.
- 20. Monica C, John NMC. A Laboratory Manual for Rural Tropical Hospitals. A Basis for Training Courses. Edinburgh: Churchill Livingstone; 1985. 209 p.
- 21. Fatma EL, Mokhtar IY, Malak AZ. Stannous chloride induces alterations in enzyme activities, lipid peroxidation and histopathology in male rabbit: Antioxidant role of vitamin C. Food Chem Toxicol. 2005;43(11):1743-52.

المستخلص

هدفت الدراسة الحالية إلى تحديد قدرة الجينسنغ على مقاومة الأثار الضارة لكلوريد القصدير "sncl2" على الكلى وبروتينات المصل. استخدمت ذكور الأرانب في هذه الدراسة. بلغ عدد مجموعات الدراسة 4 مجموعات، تضم كل منها 5 أرانب، أي ما مجموعه 20 أرنبا. خضعت الحيوانات للعلاج عن طريق الفم يومينا لمدة اثني عشر أسبوعا. صممت التجربة على النحو التالي: أعطيت أرانب المجموعة الضابطة (المجموعة 1) زيت الذرة فقط. عولجت أرانب عريق الفم يومينا لمدة اثني عشر أسبوعا. صممت التجربة على النحو التالي: أعطيت أرانب المجموعة الضابطة (المجموعة 1) زيت الذرة فقط. عولجت أرانب "مجموعة الضابطة (المجموعة 2) بجرعة • 100 ملغم/كغم من وزن الجسم" فقط من الجين سنغ. عولجت أرانب "مجموعة كلوريد القصدير "sncl2) (النب "مجموعة 2) بجرعة • 100 ملغم/كغم من وزن الج سم" فقط من الجين سنغ. عولجت أرانب "مجموعة 3) إلمجموعة 3) إلمجموعة 3) بجرعة • 200 ملغم/كغم من وزن الج سم" فقط من الجين سنغ. عولجت أرانب "مجموعة الجينسنغ + 2sncl2" "sncl2" "مجموعة 3) (المجموعة 3) (المجموعة 3) بجرعة • 200 ملغم/كغم من وزن الج سم" فقط من الجين سنغ. عولجت أرانب "مجموعة الجينسنغ + 2sncl2" (sncl2) (المحموعة 3) (المحموعة 3) بجرعة • 200 ملغم/كغم من وزن الجسم" فقط من الجين سنغ. عولجت أرانب "مجموعة الجينسنغ + 2sncl2" "(sncl2) (المحموعة 3) باجينسنغ وكلوريد الق صدير معا. أظهرت النتائج ارتفاعا ملحوظا لدى أرنب مجموعة الجين سنغ في معايير العص التالية: البروتين (TP) (المجموعة 4) بالجين سنخ و وكلوريد الق صدير معا. أظهرت النتائج ارتفاع ملحوظا لدى أرنب مجموعة الجين سنخ في معايير العل التالية ارتفي "soncl2" (TP) ، والألبومين (A) ، والغلوبيولين . (B)" كما انخفض مستوى اليوريا والكرياتينين بشكل ملحوظ "soncl2" بشكل ملحوظ الدى أرتب مجموعة الحير الحرية التين بشكل "soncl2" بالحموية (TP) ، والألبورين (B)" كما انخفض مستوى اليوريا والكرياتينين بشكل ملحوظ "soncl2" بشكل الكي (TP) ، والألبومين (A) ، والغوبيولين . (B)" ما انخفض مستوى اليوريا والكرياتينين بشكل ملحوظ يقلل الجن سنغ من الأثار النصل " SnCl2 على جميع المايير المختبرة، وقد ثبت ذلك في مجموعة الجرية والكرياتينين" بشكل ملحوظ يقلل الجين سنغ من الأثار الصل " TP م ما و G" و شكل ملحوظ بينما ارتفعت مستويات الجين عليعا ينائس من ما الأثار الضراق الحارة الحامي الحمي يلغمارما وع م