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Protective and Curative Effect of Costus Roots on Hepatotoxicity in Male Albino Rats

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Abstract:

The purpose of this study was to ascertain the impact of costus (CS roots powder concentrations 2.5% and 5% on the hepatotoxicity of male albino rats. Forty-two male albino rats were employed in the experiment. The rats were separated into seven main groups, including omit it curative and protective groups, after being fed the optimal diet for one week (six rats omit it each). Therapeutic teams: Group 1 was served a basal diet as the negative control group. Group 2 was a positive control group that received a basal diet along with three weekly injections of carbon tetrachloride (CCl₄) (1 mg/kg body weight) to cause hepatotoxicity. The group (2) plus 2.5% (CS) powdered roots was fed to the third group. Group (4) was fed the same as group (2) plus 5% (CS) powdered roots. Defense Organizations Group (1)'s curative group and negative control group were similar. After eight weeks, the group (5) positive add the control group was given a basal diet plus a CCl₄ injection (1 mg/kg body weight). The group (5) plus 2.5% (CS) powdered roots was fed to the group (6). The group (7) received the group (5) plus 5% (CS) powdered roots. Phytochemical analysis for costus roots was determined. The investigation documented that providing hepatotoxicity CS root powder at 2.5% and 5% in the curative add and protective groups increased body weight gain, feed intake and feed efficiency ratio. Our results could be add as follows: diets enriched with 2.5% and 5% CS roots powder improved blood lipid levels and reduced kidney and liver function risks.

Keywords: Hepatotoxicity - Costus roots - Liver function - Kidney function - antioxidant

Introduction

The treatment of liver illness is still difficult for hepatologists and liver diseases continue to be major health problems. [1,2]. Particularly in Egypt, liver fibrosis is a significant public health concern. Fibrosis of the liver is a long-term wound-healing response to recurrent injury that is related to the etiology of the lesion. However, liver fibrosis is a dynamic condition that may reverse after the injury's underlying cause is dealt with. However,

cirrhosis develops when an injury is prolonged, which reduces the likelihood that it will heal [3,4].

The ccl4 model is a special model that mimics all significant features of human liver fibrosis, especially in those with hepatitis C virus infection. These features include inflammation, portal hypertension, ascites, fibrous formation, and possibly fibrosis regression. It is also very easy to compare, has excellent reproducibility, and has been used in several previous investigations. [5]. CCl₃, a dangerous product of CCl₄, binds to lipoprotein, causing lipids to oxidize, increasing liver malonaldehyde (MDA), releasing reactive oxygen species, and lowering liver glutathione (GSH). Following this, the properties of hepatocyte the bloodstream, mitochondrial, and lysosomal membranes alter, resulting in hepatocyte enlargement and lysis, which boosts the blood level of the enzyme alanine transaminase (ALT) and the enzyme aspartate transaminase (AST)[6]. Acute damage, the start of fiber synthesis, and severe fibrosis are the three stages that make up the development of liver fibrosis. Kupffer cell activation & the beginning of an inflammatory response, which promotes the production of cytokines and chemokines, as well as the attraction and activation of neutrophils and lymphocytes, which contribute to liver necrosis, are characteristics of the phase of acute ccl4-mediated liver fibrosis [7].

Herbal medicines are frequently utilized for the treatment and prevention of a variety of illnesses and frequently contain highly active pharmacological substances that are typically thought to be harmless and nontoxic in comparison to synthetic drugs [8]. Dry seeds of costus (SC) have been utilized by folk medicine since ancient times for the treatment of many diseases and conditions, like allergies & asthma, vomiting, certain bronchitis, ulcer, throat infections, arthritis, coughing, tuberculosis, gastrointestinal difficulties & many others [9,10]. The costus of *Saussurea* is also known as the Indian costus or costus [11]. For many thousands of years, *S. costus* has been applied as a medicine, plus its root of this species is the major part utilized for medicinal purposes, and it was suggested by Prophet Muhammad to "treat with Indian burning incense, for the purpose of treatment for seven disorders" [12]. *S. costus* roots, besides other types, were used for cancer treatment by ancient physicians & more recently, by Japanese scientists & *S. costus* compounds and extractions have been listed in numerous pharmacopoeias [13]. Whenever consumed as a nutritional product, the Food and Drug Administration of America concerns *S. costus* to be commonly accepted safe [14].

Saussurea costus is used separately or in mixture with other medications in Indian health systems. Because its roots include antispasmodic, antifungal, and antibacterial effects, *S. costus* should be used in more human-related healthcare and pharmaceuticals [15,16]. Considering the worldwide increase in resistant to drugs diseases and the expensive nature of therapy, plants used for medicinal purposes with antifungal abilities is a potential factor in innovative antimicrobial drugs in the type of pure elements or natural extracts [17]. SO, this study was conducted to investigate the effect of feeding rats with hepatotoxicity on costus roots on serum liver functions, kidney function, lipid profile and antioxidant enzymes.

Materials and methods

Materials:

1-Plants:

Costus roots was obtained from Agriculture Research Center, Egypt.

2-Basal diet:

Starch and corn oil were bought at the local market. The Cairo Company for Chemical Trading, in Cairo, Egypt, provided the casein, cellulose, vitamins, minerals, dextrin, L-cysteine, choline chloride.

3- Carbon tetra chloride (CCl₄):

Carbon tetrachloride (CCl₄) was obtained from El-Gomhoryia Company for Chemical Industries, Cairo, Egypt as 10% liquid solution.

4-Rats:

Adults male albino rats (n = 42) of Sprague- Dawely strain weighing (150-180g) were purchased from Helwan Farm of Experimental Animals, Helwan, Egypt.

5-Chemical Kits

Chemical kits manufactured by Egyptian American Corporation and given by Alkan Organization for laboratory service are employed for monitoring triglycerides, cholesterol, LDL-C, urinary acid, ammonia nitrogen, creatinine level, and transaminases.

Methods:

1-Animal ethics statement:

The Institutional Animal Care and Use Committee (ARC-IACUC) at Agriculture Research Center is organized and operated according to the world Organization for Animal Health (OIE) and the Eighth Edition of the Guide for the Care and Use of Laboratory Animal (2011). Ethical committee approved this study via authorized veterinarian with applying minimum constrain to the animals and using approved sample collection methods.

2-Dry costus roots preparation

Dry costus roots were prepared by being ground in an electric blender, then root powders were added to the daily basic diet.

3-Determination of phytochemical qualitatively analysis of costus roots by HPLC:

The colorimetric measurements were performed in the manner described by [18, 19].

4-Induction of hepatotoxicity in experimental rats:

Protective group:

Intraperitoneal infusion of CCl₄ (50%) in olive oil as the solvent (1ml/kg) was administered at three intervals per week to both groups to elicit hepatotoxicity throughout the last week of the research. [20], proposed a method for sacrificing rats twenty-four hours after the previous CCl₄ treatment.

Curative group:

In both groups, CCl₄ (50%) in olive oil (1 ml/kg) was administered through the abdomen 3 times a week for one week to elicit toxicity to the liver. The trial ended for 8 weeks throughout the testing period.

5-Concept of the study:

Animals were habituated for a week beforehand the trial began. Water was available all day. The rats were separated into seven groups, including therapeutic and protection groups (each with six rats). Curative groups: Group (1): Negative control group was fed on basal diet. Group (2): Positive control group was fed on basal diet + CCl₄ injection (1mg/kg body wt.) from the first day Groups (3 & 4) were fed as the group (2) + 2.5% & 5% (CS) roots powder. Protective groups: Group (1): Negative control group was the same as Group (1) in curative group. Group (5) Positive control group was fed on basal diet + CCl₄ injection (1 mg/kg body wt.) in the last week. Groups (6 & 7) were fed as the group (5) + 2.5% & 5% (CS) roots powder.

6-Blood sampling:

Following a night of fasting after the experiment, the rats were sedated, slaughter & blood from them was extracted from the aorta. The serum was separated by centrifuging the blood samples for fifteen minutes at 3000 rpm per minute. The blood sample was properly split into dry, sterile Wassermann tubes with a pipette from Pasteur and placed at -20°C until testing.

7-Biochemical analysis of serum:

Total cholesterol [21], triglycerides [22], HDL-C; [23], LDL-C, and VLDL-C were calculated as the equation of [24]. All provided methods for determining uric acid, urea nitrogen, and creatine [25, 26, 27], respectively. Both the AST and ALT enzymes were discovered by researchers using the methods described by [28]. Serum alkaline phosphatase (ALP) was determined according to the method described by [29]. Serum malondialdehyde (MDA) and glutathione were measured according to [30, 31].

8-Statistical evaluation:

The outcomes are shown in terms of mean standard deviation. A single-way analysis of variation, or "ANOVA," was utilized to statistically evaluate the variance in the information [32]. These computations were performed using SPSS (version 15) computer software.

Results and discussion

The phytochemical examination results revealed that costus roots have a high concentration of bioactive elements for example resins that are terpenoids, phenols, alkaloids, coumarins, flavonoids, tannins, steroids, and quinones. These phytochemical elements are essential for health care applications [33]. Our findings supported prior research, which found that *S. lappa* (synonymous *S. costus*) is a rich source of alkaloids, flavonoids, steroids, and resins. Furthermore, [34] stated that various bioactive compounds, including sesquiterpene lactones, isodehydrocostus, isozaluzanin-C, guaiainolide, santamarine, and others, were discovered and extracted from *S. costus*. The abundance of phytochemical substances in *costus* is undoubtedly responsible for its various biological operations.

Also, [35] found that, various polyphenols have been discovered to have a wide spectrum of activities that may aid in the prevention of serious illnesses. The existence of for medicinal purposes valuable bio active components such as flavonoids, tannins, alkaloids, phenolic elements, saponin, reducing sugar, and glycosides is revealed by quantitative analysis of five distinct leaf extracts of *Costus igneus*.

Table (1): Costus roots phytochemical analysis.

The components of phytochemistry	The outcomes of the tests
Resins	+
Terpenoids	+
Phenol/Polyphenols	+
Alkaloids	+
Coumarins	+
Flavonoids	+
Tannins	+
Steroids	+
Quinones	+
Volatile oil	-
Phlobatannins	-
Saponins	-
Lipids	-

+ = test positive, - = test negative

The second table shows the impact of administering liver injury rats costus roots powder on rat body weight gain, consumption of food & feed effectiveness ratio. According to the information that was brought about, there was a substantial decline in BWG, FI, and FER for the treatment positive group. (108.60 ± 5.41 ; 8.90 ± 0.52 and 12.20 ± 1.02) after the single dose of ccl4 (2mg/kg) as contrasted with the negative control group (142.20 ± 6.14 ; 11.20 ± 0.70 and 12.70 ± 0.84), respectively. Feeding acute renal failure with CS roots powder at 2.5% & 5% from the first day in curative groups showed significant increase values of BWG%; FI and FER contrasted with control positive group. Results revealed that the mean values and \pm SD for curative group fed at 2.5% CS were 117.66 ± 1.81 ; 9.92 ± 0.30 and $11.80 \pm 0.37\%$. While, the mean values and \pm SD for curative group fed at 5% CS were 122.00 ± 2.12 ; 11.02 ± 0.41 and $11.07 \pm 0.50\%$ respectively, compared to the control positive group (108.60 ± 5.41 ; 8.90 ± 0.52 and 12.20 ± 1.02), respectively.

Moreover, results revealed that protective groups which fed on basal diet containing CS roots powder at 2.5 percent & 5 percent then injected with CCL4 in the last week, improved the BWG %; FI and FER, contrasted with control positive group. The mean values for protective group fed on 2.5% CS were 138.00 ± 1.67 ; 9.80 ± 0.21 & 10.56 ± 1.67 ; and the mean values for protective group fed on 5% CS were 140.00 ± 1.87 ; 11.64 ± 0.37 & 12.00 ± 0.46 compared with control positive group 130.00 ± 5.24 ; 10.82 ± 0.32 & 12.02 ± 0.52 respectively. The effect of the rats' diet's odour, taste, and palatability may be the cause of the body weight gain that increases when costus levels rise.

According to [36], the injection of CCl4 is linked to reduced nutrient metabolism and absorption due to poor bile secretion as well as hunger loss that results in weight loss. According to [37], ccl4 significantly decreased feed intake, body weight gain, and feed efficiency rate.

Table (2): Effect of feeding hepatotoxicity rats on costus roots powder on body weight gain, feed intake and feed efficiency ratio.

Categories	Variables	Body weight Gain (%)	Feed intake (g/day)	Feed efficiency ratio
Control (-ve)		142.20±6.14a	11.20±0.70a	12.70±0.840a
Control (+ ve) ccl4 injection (1mg/kg body wt.) from the first day three times a week		108.60±5.41e	8.90±0.52c	12.20±1.02a
Curative groups	Control (+ve) +2.5%CS	117.60±1.81d	9.92±0.30b	11.80±0.37b
	Control (+ve) +5%CS	122.00±2.12d	11.02±0.41a	11.07±0.50b
Control (+ ve) ccl4 injection (1mg/kg body wt.) in the last week three times a week		130.00±5.24c	10.82±0.32b	12.02±0.52a
Protective Groups	Control (+ve) +2.5%CS	138.00±1.67b	9.80±0.21b	10.56±1.67c
	Control (+ve) +5%CS	140.00±1.87b	11.64±0.37a	12.00±0.46a

*Values are presented in terms of averages ± SE.

*Values numbers in the exact same column with different lettering vary considerably.

The findings presented in Table 3 demonstrated that there were statistically significant differences in the blood levels of triglyceride (TG) and total cholesterol (TC) among the rats in the negative control group and the rats in the positive control group (hepatotoxicity), the values were 99.80±3.11 & 150.60±9.31 vs 188.80±6.38 & 261.40±6.98 (curative group) and the values were 99.80±3.114 & 150.60±9.31 vs 134.80±3.49 & 163.00±2.73 (protective group) respectively.

Feeding curative groups with 2.5 and 5 percent of CS induced significant reduction ($p < 0.05$) in triglyceride & total cholesterol (106.20±4.43 & 159.60±4.72 (2.5% CS) & 124.00±6.04 & 129.60±6.30 (5% CS) contrasted with those of the control positive group 188.80±6.38 & 261.40±6.98 respectively. Moreover, with protective groups, the values were 117.80±8.34 & 162.60±5.32 (2.5% CS) and 100.60±1.81 & 162.20±5.63 (5% CS), contrasted with those of the control positive group 134.80±3.49 & 163.00±3.73 respectively. In fact, rats which fed on CS roots powder at 2.5% & 5% had lesser mean values of TC and TG contrasted with the positive control group.

Additionally, costus administration considerably controlled any changes in the blood lipid profile of hypercholesterolemic rats, which was likely attributable to the antidyslipidemic effect exhibited by the phytochemicals, particularly castanoside [38]. Our findings agree with [39] who reported that total cholesterol levels in the positive control group were clearly greater than those in the group with a negative control, with a statistically significant difference ($P < 0.05$).

Table (3): Effect of feeding hepatotoxicity rats on costus roots powder at different ratios on triglyceride and total cholesterol.

Variables	TG	TC
Categories	(mg/dl)	
Control (-ve)	99.80±3.114f	150.60±9.31d
Control (+ ve) ccl4 injection (1mg/kg body wt.) from the first day three times a week	188.80±6.38a	261.40±6.98a
Curative groups	Control (+ve) +2.5%CS	106.20±4.43e
	Control (+ve) +5%CS	124.00±6.04c
Control (+ ve) ccl4 injection (1mg/kg body wt.) in the last week three times a week	134.80±3.49b	163.00±2.73b
Protective Groups	Control (+ve) +2.5%CS	117.80±8.34d
	Control (+ve) +5%CS	100.60±1.81f

*Values are presented in terms of averages ± SE.

*Values numbers in the exact same column with different lettering vary considerably.

Table (4) demonstrates that when the blood levels of high-density lipoprotein, low density lipoprotein, and very low-density lipoprotein in the rats that were exposed to hepatotoxicity were contrasted with the negative control group, there were significant differences. Rats that were given a diet enriched with CS roots powder at 2.5 percent and 5 percent had significantly reduced levels of both LDL-C and VLDL-C on average when contrasted with the positive control group. All treatment groups (curative and protective) with enriched meals including various levels of CS demonstrated greater mean HDL-C values when contrasted with the positive control group.

Decline in HDL-C at all doses studied may not be functionally advantageous to the rats due to the reduced rate of blood cholesterol transfer to the organ of liver, the considerable reduction in triacylglycerol may be due to decreased lipolysis[40].

Table (4): Effect of feeding hepatotoxicity rats on costus roots powder at different ratios on HDL, LDL and VLDL.

Variables	HDL	LDL	VLDL
Categories	(mg/dl)		
Control (-ve)	35.40±1.67d	84.24±2.40d	19.96±0.62c,d
Control (+ ve) ccl4 injection (1mg/kg body wt.) from the first day three times a week	34.80±3.11a	180.04±8.15c	37.76±1.27a
Curative groups	Control (+ve) +2.5%CS	38.60±1.67c	88.76±7.68d
	Control (+ve) +5%CS	42.60±1.67b	92.20±4.51c
Control (+ ve) ccl4 injection (1mg/kg body wt.) in the last week three times a week	37.80±1.30a	105.08±6.04b	26.96±0.69b
Protective Groups	Control (+ve) +2.5%CS	41.40±1.67b	96.80±5.21C
	Control (+ve) +5%CS	43.00±1.22c	80.20±1.77e

*Values are presented in terms of averages ± SE.

*Values numbers in the exact same column with different lettering vary considerably.

Table (5) displays the influence of costus roots powder on the renal systems (urea nitrogen, uric acid & creatinine) in rats with hepatotoxicity. Carbon tetrachloride caused hepatotoxicity by increasing biochemical indicators for kidney not liver. Furthermore, the results revealed a significant difference between the positive control group and the other groups (negative control, curative, and protective groups) fed CS roots powder at 2.5% and 5%. While rats fed CS powder at 2.5% and 5% lowered their uric acid levels dramatically. The hepatic damage group had a higher mean value of urea nitrogen at 42.00 ± 2.12 mg/dl contrasted with the negative control group at 36.40 ± 2.40 mg/dl. In treatment groups provided on CS roots powder at 2.5% and 5%, the levels of urea nitrogen reached 38.80 ± 1.92 and 36.60 ± 2.30 mg/dl, respectively, contrasted with the positive control group 42.00 ± 2.12 mg/dl. In protective groups, the levels of urea nitrogen reached 39.00 ± 1.00 and 40.60 ± 1.51 mg/dl, respectively, compared to the positive control group 41.00 ± 1.00 mg/dl. Additionally, table (5) demonstrated that creatinine elevated with ccl4 injection, reaching 0.86 ± 0.05 mg/dl in contrast to 0.48 ± 0.08 mg/dl in the negative control group. Furthermore, results confirmed that the consequences of feeding on basal diet containing CS roots powder at 2.5% & 5% (curative groups), creatinine levels reached 0.60 ± 0.10 & 0.54 ± 0.05 mg/dl contrasted with positive control group 0.86 ± 0.05 mg/dl. When compared to the positive control group, the protective groups achieved levels of 0.70 ± 0.07 and 0.70 ± 0.05 mg/dl. Our findings corresponded with those of [41], who showed that the levels of creatinine, uric acid, and urea did not change in comparison to group one (control).

[42] demonstrated in related research that lipid peroxidation and a decrease in antioxidant status may be linked to a chain of events leading to methotrexate (MTX)-induced renal damage. Increased serum urea & creatinine levels can also indicate renal failure and the activation of apoptotic cell markers like PARP, which may play a role in the kidney damage caused by MTX. In lead-exposed populations, kidney impairment is related with distortions in several renal functions such as albuminuria, impaired glomerular filtration rate, and lower creatinine clearance [43, 44].

Table (5): Effect of feeding hepatotoxicity rats on costus at different ratios on kidney function.

Categories	Variables	Creatinine	Urea (mg/dl)	Uric acid
Control (-ve)		$0.48 \pm 0.08_{d,e}$	$36.40 \pm 2.40_c$	$3.68 \pm 0.08_b$
Control (+ve) ccl4 injection (1mg/kg body wt.) from the first day three times a week		$0.86 \pm 0.05_a$	$42.00 \pm 2.12_a$	$4.96 \pm 0.19_a$
Curative groups	Control (+ve) +2.5%CS	$0.60 \pm 0.10_c$	$38.80 \pm 1.92_b$	$3.94 \pm 0.23_{a,b}$
	Control (+ve) +5%CS	$0.54 \pm 0.05_d$	$36.60 \pm 2.30_c$	$3.88 \pm 0.08_{a,b}$
Control (+ve) ccl4 injection (1mg/kg body wt.) in the last week three times a week		$0.74 \pm 0.05_b$	$41.00 \pm 1.00_a$	$4.16 \pm 0.26_a$
Protective Groups	Control (+ve) +2.5%CS	$0.70 \pm 0.07_b$	$39.00 \pm 1.00_b$	$3.88 \pm 0.31_{a,b}$
	Control (+ve) +5%CS	$0.70 \pm 0.05_b$	$40.60 \pm 1.51_{a,b}$	$4.12 \pm 0.39_a$

*Values are presented in terms of averages \pm SE.

*Values numbers in the exact same column with different lettering vary considerably.

Results in Table (6) displayed that serum AST, ALT, and ALP activity considerably elevated ($P < 0.05$) in the positive group contrasted with the negative control group. Our findings support [39] findings from, which show how costus roots affect the levels of ALT, AST, and ALP in rats with liver fibrosis. The results revealed a statistically significant difference among the ALT liver enzyme levels in the positive control rat group and the negative control rat group. Rats were given calcium carbide, and [45, 46] found that the ALT and AST greatly increased. According to [47] *Costus speciosus* could modify plasma enzyme levels (AST, ALT, LDH, and ALP) to be quite close to normal. Additionally, rats that had been poisoned with carbon tetrachloride [48, 49] to determine whether the ethanolic extract of the *Costus speciosus* rhizomes showed hepatoprotective properties. The serum levels of urea and creatinine were considerably greater in the hypothyroidism and hyperthyroid mice as contrasted with the control & costus groups, according to the most recent research ($p < 0.05$). According to [50] the cholesterol-treated group (+Ve) significantly outperformed the negative control group (-Ve) and all other groups in terms of the activities of liver function (tissue injury markers). The costus, on the other hand, significantly decreased all tissue injury markers in the hypercholesterolemic rats. This effect may be attributable to the costus' free radical scavenging ability, which can maintain the integrity of the cell membrane [51]. [52,53] both declared that costus (*S. lappa*) root extract had hepatoprotective action towards liver damage brought on by contact with deltamethrin. [54, 55, 56] demonstrated that *Costus speciosus* might alter plasma enzyme levels to close to their usual levels and function as a hepatoprotective agent regarding carbon tetrachloride. These enzymes included aspartate aminotransferase, alanine aminotransferase, alkaline phosphatase, lactate dehydrogenase, alkaline phosphatase, and acid phosphate.

Table (6): Effect of feeding hepatotoxicity rats on costus at different ratios on liver function.

Categories	Variables	ALT	AST (U/L)	ALP
Control (-ve)		30.60±4.61e	40.80±0.83d	809.00±7.28e
Control (+ve) ccl4 injection (1mg/kg body wt.) from the first day three times a week		66.40±2.07a	116.00±7.48a	862.20±4.76a
Curative groups	Control (+ve) +2.5%CS	36.80±3.76d	39.00±1.00d	834.00±3.80c
	Control (+ve) +5%CS	31.60±2.07e	36.60±2.30e	825.20±4.32d
Control (+ve) ccl4 injection (1mg/kg body wt.) in the last week three times a week		54.60±5.89b	86.60±5.12b	843.60±3.64b
Protective Groups	Control (+ve) +2.5%CS	37.00±2.55d	55.80±4.43c	842.40±5.50b
	Control (+ve) +5%CS	41.40±1.67c	51.40±5.50c	838.00±5.56c

*Values are presented in terms of averages \pm SE.

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As can be shown in Table (7), rats in the positive control group had significantly greater serum MDA levels than rats in the healthy group (153.40±6.95 mM/g vs 78.00±7.17 mM/g).

Rats in other groups, as opposed to the treatment group, saw a considerably ($p < 0.05$) lower level of serum MDA.

In addition, the results showed that supplying "curative groups" to liver-toxic rats lead to a considerable decline in the mean values of serum MDA; these values were 94.60 ± 5.81 mM/g (for 2.5%) and 91.20 ± 2.58 mM/g (for 5%), as opposed to 153.40 ± 6.95 mM/g for the positive control group, respectively. Additionally, the mean serum MDA levels in the current study's "protective groups" were significantly lesser than those in the "positive control group" (129.60 ± 5.41 mM/g), at 100.40 ± 3.20 mM/g (for 2.5%) and 94.60 ± 5.81 mM/g (for 5%), respectively.

According to the study's findings, rats in the control positive group had serum glutathione levels that were substantially lower than rats in the negative control group.

According to [41], the levels of MDA in red blood cells (RBCs) were statistically considerably reduced while GSH dramatically increased. MDA levels in the liver homogenate of the ccl4 treated group 2 were significantly greater than those in the untreated group 1 ($P < 0.05$), according to the comparison. This can be explained by the fact that ccl4 is changed into the dangerous metabolite trichloromethyl (ccl3) radical by CYP2E1 in hepatocytes. The binding of ccl3 to lipoproteins causes the lipids in the endoplasmic reticulum to undergo peroxidation, which increases the level of MDA in the liver [56]. GSH levels in the liver homogenate of the ccl4-treated group 2 significantly decreased in comparison to the negative control (-ve), demonstrating that oxidative stress mediated by ccl4-induced hepatocyte damage led to the depletion of reduced GSH. This conclusion is in line with those reached by the investigations by [57, 58].

Table (7): Effect of feeding hepatotoxicity rats on costus at different ratios on malondialdehyde and glutathione.

Categories	Variables	MDA (mM/g)	GSH (μ M/g)
Control (-ve)		$78.00 \pm 7.17f$	$4.96 \pm 0.19a$
Control (+ve) ccl4 injection (1mg/kg body wt.) from the first day three times a week		$153.40 \pm 6.95a$	$3.00 \pm 0.56c$
Curative groups	Control (+ve) +2.5%CS	$94.60 \pm 5.81d$	$3.94 \pm 0.23a,b$
	Control (+ve) +5%CS	$91.20 \pm 2.58d,e$	$3.72 \pm 0.40b$
Control (+ve) ccl4 injection (1mg/kg body wt.) in the last week three times a week		$129.60 \pm 5.41b$	$4.16 \pm 0.26a$
Protective Groups	Control (+ve) +2.5%CS	$100.40 \pm 3.20c$	$4.12 \pm 0.39a$
	Control (+ve) +5%CS	$94.60 \pm 5.81d$	$3.88 \pm 0.31a,b$

*Values are presented in terms of averages \pm SE.

*Values numbers in the exact same column with different lettering vary considerably.

The aforementioned chemical analysis of the costus was crucial in reducing lipid peroxidation and oxidative stress, which are a result of an imbalance between the production of ROS (reactive oxygen species) & the antioxidant enzymes' capacity to scavenge free radicals. Excessive ROS production results in this imbalance and antioxidant exhaustion [59 & 38].

According to [60] study, rats treated with lead acetate had considerably lower levels of total glutathione, superoxide dismutase, and glutathione peroxidase. *Costus afer*, which likewise considerably decreased the value of the lead acetate-induced indicator of lipid peroxidation, malondialdehyde, significantly reversed these lead acetate-induced decreases. This outcome is consistent with that which [61] reported. This was consistent with the earlier study, which found that *C. speciosus* rhizomes have preventive potential due to their antioxidant effects in preventing degenerative disorders with obvious oxidative damage from ROS or free radicals [62].

Conclusion

Diets enriched with 2.5% and 5% CS roots powder improved blood lipid levels and reduced kidney and liver function risks.

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التأثير الوقائي والعلاجي لجذور القسط الهندي على التسمم الكبدي في ذكور الفئران

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الملخص العربي:

الغرض الأساسي من هذه التجربة دراسة تأثير تركيزات مسحوق جذور القسط الهندي (2.5% و5%) على التسمم الكبدي لذكور الفئران البيضاء. تم استخدام 42 من ذكور الفئران البيضاء في التجربة. وتم تقسيم الفئران إلى سبعة مجاميع، بما في ذلك المجموعات العلاجية والوقائية، بعد تغذيتهم على النظام الغذائي الأمثل لمدة أسبوع واحد (سنة فئران لكل منهما) المجموع العلاجية: تم تقديم النظام الغذائي الأساسي للمجموعة الأولى كمجموعة ضابطة سالبة. المجموعة الثانية كانت مجموعة ضابطة موجبة التي تغذت على نظام غذائي أساسي مع ثلاث حقنات اسبوعيا من رابع كلوريد الكربون (1 ملجم/كجم من وزن الجسم) للتسبب في تسمم الكبد. المجموعة الثالثة تم تغذيتها بنفس طريقة تغذية المجموعة (2) بالإضافة إلى 2.5% من الجذور القسط الهندي. المجموعة (4) تم تغذيتها بنفس طريقة تغذية المجموعة (2) بالإضافة إلى 5% من مسحوق القسط الهندي. بعد ثمانية أسابيع، أعطيت المجموعة (5) مجموعة ضابطة موجبة نظام غذائي بالإضافة إلى حقنة رابع كلوريد الكربون (1 ملجم/كجم من وزن الجسم) تم تغذية المجموعة (5) بالإضافة إلى 2.5% من جذور القسط البودر كمجموعة (6). تلقت المجموعة (7) نفس غذاء المجموعة (5) بالإضافة إلى 5% من مسحوق جذور القسط الهندي. تم تقدير المركبات الفعالة في جذور القسط الهندي. تم أخذ عينات دم من الشريان الأورطي للفئران في نهاية التجربة. اتضح من النتائج أن مسحوق جذور القسط الهندي عند مستوى (2.5% و5%) في المجموعات العلاجية والوقائية أدى إلى زيادة وزن الجسم. يمكن تلخيص النتائج على النحو التالي: ساعد النظام الغذائي المدعم بنسبة (2.5% و5%) من مسحوق جذور القسط الهندي على تعزيز مستويات الدهون في الدم وكذلك تقليل المخاطر على وظائف الكلى والكبد.

الكلمات المفتاحية: التسمم الكبدي، جذور القسط الهندي، وظائف الكبد، وظائف الكلى.