

Effect of Corn Silk Extract on Renal Functions in Acetaminophen Induced Nephrotoxicity in Mice

FAIZA MEHBOOB¹, SAEED KANWAL², MUHAMMAD KAMRAN AMEER³, NAVEED AKHTER⁴, NADIA MAJEED⁵, SHAMSA IJAZ⁶

ABSTRACT

Aim: To evaluate the effects of corn silk on renal functions in acetaminophen induced nephrotoxicity

Study design: Experimental Study.

Settings: Anatomy Department, University of Health & Sciences Lahore. From February, 2012 to February, 2013.

Methods: Twenty seven male albino mice of age 6-8 weeks and weight 30 ± 5 gm were used. The control Group A was treated with single injection of 16.6ml/kg normal saline on day one of the study via intraperitoneal route and was sacrificed on day ten. Group B was treated with single intraperitoneal injection of acetaminophen (600 mg/kg) and was sacrificed after 2 days. Group C was administered single intraperitoneal injection of acetaminophen (600 mg/kg) on day one of the experiment followed by the oral gavage of corn silk (CS) extract 400 mg/kg for next 8 days and then sacrificed on day ten. For blood sampling cardiac puncture was performed.

Results: A statistically significant raised ($p < 0.005$) levels of serum urea and creatinine in group B were observed. In groups C serum urea and creatinine levels were significantly reduced.

Conclusion: The results of present study suggest CS extract provides protection against acetaminophen induced nephrotoxicity.

Keywords: Acetaminophen, Nephrotoxicity, Corn Silk, Mice, Serum urea and creatinine.

INTRODUCTION

Acetaminophen is used for management of pain, fever and osteoarthritis¹ owing to its antipyretic and analgesic effects². Prostaglandins are inflammatory mediators that are blocked by acetaminophen by inhibiting cyclooxygenases². Acetaminophen is readily absorbed after ingestion³. First phase of metabolism is completed in liver. Approximately 15 percent of drug is oxidized into a toxic metabolite, N-acetyl-p-benzo-quinone imine (NAPQI) by cytochrome P-450 system in liver².

Over dosage of acetaminophen causes acute renal tubular necrosis through consumption of glutathione (an antioxidant system) and covalently binding the reactive metabolite to vital cell constituents⁴.

The long silky hairs of stigma and styles of the maize plant are the corn silk. It had long been used for the treatment of edema, gouty arthritis, urinary bladder infection, kidney stones, nephritis, prostatitis and diabetes mellitus^{5,6}. It is a mild diuretic and demulcent. CS constituents include amines, saponins, pigments, fixed oils, resins, flavanoids, alkaloids, chlorogenic acid, tannins, phytoesters, allantoin⁷.

MATERIAL AND METHODS

Acetaminophen was procured from Merck Pharmaceuticals. 600mg of acetaminophen powder was dissolved in 16.6ml normal saline. CS extract was prepared

by a rotating evaporator at PCSIR Laboratories complex, Lahore. 100gm CS was dissolved in 1000ml methanol (80%). 2000mg of CS extract was dissolved in 50ml of distilled water. Mouse Dose of CS extract was calculated 400mg/kg/day with the help of dose conversion formula⁸. Three experimental groups (A, B & C) consisting of nine animals were randomly selected.

3ml of blood was collected via cardiac puncture exchanged to vacutainer. Blood was permitted to remain for one hour before centrifuging it at a speed of 3000 cycles per minute. The clear serum was gathered and stored at -20°C . Serum urea and creatinine levels were estimated by utilizing commercially accessible kits.

Statistical analysis:

Data was analyzed through SPSS version 18. Mean and standard deviation was calculated for quantitative variables. One way ANOVA was applied to compare variables and afterward post Hoc Tukey Test. p value < 0.05 was considered as statistically significant.

RESULTS

Serum urea levels of acetaminophen injected group B were significantly raised as compared to control group A ($p=0.001$), whereas serum urea levels were significantly reduced ($p=0.001$) in the treatment group C when compared to group B (Table 2). Mean serum creatinine level of toxic group B was significantly increased when compared with control group A ($p=0.001$), while mean serum creatinine level of group C receiving CS extract was significantly reduced when compared with that of group B ($p=0.001$).

¹Assistant Professor Multan Medical & Dental College Multan

²Associate Professor ABWA Medical & Dental College Faisalabad

³Assistant Professor Multan Medical & Dental College Multan

⁴Associate Professor Rai Medical College Sargodha

⁵Assistant Professor Niazi Medical College Sargodha

⁶Assistant professor Islam Medical College Sialkot

Correspondence to Dr. Saeed Kanwal Email: kashalpal@gmail.com Cell: 0323-4225122

Table 1 Experimental Groups

Group	Intervention	Dosage/day	Route	Schedule	Sacrificed
A	0.9% N. Saline	17ml/kg	Intraperitoneal	1 st day	10 th day
B	Acetaminophen	600 mg/kg	Intraperitoneal	1 st day	After 48 hours
C	Acetaminophen CS extract	600mg/kg 400mg/kg	Intraperitoneal Oral	1 st Day 2 nd to 9 th Day	10 th day

Table 2. Comparison of Mean serum urea & Mean serum creatinine level b/w groups A, B, & C.

Factor	Group A (Mean±SD)	Group B (Mean±SD)	Group C (Mean±SD)	p-value
Serum urea (mg/dl)	44.49±7.52	83.15±6.25	60.97±5.46	0.001*
Serum creatinine (mg/dl)	0.46±0.11	1.98±0.26	1.18±0.25	0.001*

* p≤0.05 is considered statistically significant.

DISCUSSION

CS constituents, owing to its antioxidant activity, are broadly reported to defend several organs against oxidative damage⁶. The present work is designed to assess the consequences of CS extract on renal damage produced by toxic dose of acetaminophen in mice.

Serum urea and serum creatinine (renal profile) of group B micewere significantly increased in comparison to the mice in group A. This phenomenon is proven to be dueoxidative stress provoked by acetaminophen in past studies⁹.Khosandi and Orazizadeh¹⁰ also reported impaired renal status after ingestion of 500mg/kg of acetaminophen in mice. The decline inmean serum urea of group Cwas statistically significant when compared with group B which showed that CS extract has some protective role against acetaminophen induced renal damage. Mean value of serum creatinine in group Cwas statistically significant when compared with group B. It showed that CS treatment has lowered the serum creatinine value which suggests that CS has a protective role against acetaminophen induced nephrotoxicity. Sepehri et al⁸ also documented that methanolic extract of CS alleviated the nephrotoxic action of gentamicin in the adult male rats.

Acetaminophen over dosage raises the NAPQI concentration due to saturation of sulfate and glucuronide pathways for its routine clearance so it is shunted to cytochrome P-450 system. NAPQI spontaneously traps cellular proteins and leads to the oxidative stress induced cell death⁹.

Eugenol, R-terpineol and citronellol are volatile oxygenated chemicals present in CS which act as antioxidants. Antioxidant, anti-inflammatory and clearanceof free radical properties of CS clarifies its nephroprotectiveactivity¹¹.

Present study revealed that CS extract significantly reduced renal biochemical parameters i.e., plasma urea and creatinine. This points outs that considerable protection is offered by CS extract.

CONCLUSION

The results of present study suggest that acetaminophen at a dose of 600mg/kg body weight leads to renal function

impairment while CS extract protects acetaminophen induced renal injury.

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