

Research Article

Effect of Gum Arabic on Nephrotoxicity Induced by *Aristolochia bracteata* in Rats

Amani E. Omer¹, Ietimid A.M. Ayed², Samia M.A. El Badwi^{1*}

¹Department of Pharmacology and Toxicology, Faculty of Veterinary Medicine. University of Khartoum, Sudan

²Pharmacology Department, Faculty of Medicine and Health Sciences, Omdurman Islamic University, Sudan

*Corresponding author

Dr. Samia M.A. El Badawi

Email: samiaelbadwi@yahoo.com

Abstract: The present study was designed to investigate the effect of Gum Arabic (GA) on nephrotoxicity induced by *Aristolochia bracteata* aqueous extract. Thirty six Albino rats were used, divided into 6 groups. Group 1 rats were orally given 1ml distilled water daily for 10 days and served as the control group, group 2 were orally administered with 1ml distilled water for 10 days and from day 5 to day 10 given 1000mg/kg.bwt of *Aristolochia bracteata* aqueous extract. Group 3 and 4 were treated orally with Gum Arabic 250mg/kg and 500mg/kg respectively for 10 days and then given *Aristolochia bracteata* (1000mg/kg.bwt) from day 5 to day 10. Group 5 and 6 were given Gum Arabic alone at 250mg/kg and 500mg/kg respectively for 10 days. The concentration of urea and creatinine as well as the activity of AST and ALT was measured. RBCs, Hb, PCV, MCH and MCHC were determined and the kidney histopathology was microscopically examined. Administration of G.A. was significantly decreased the elevated urea, creatinine, AST and ALT levels caused by *Aristolochia bracteata*. These findings were strongly correlated with decreased damage of the kidney tubules when compared to the group treated with *Aristolochia bracteata*. This study indicates that, GA. Possesses constituents with nephroprotective activity.

Keywords: *Aristolochia bracteata*, Gum Arabic, Nephrotoxicity

INTRODUCTION

Aristolochia is a large plant genus with over 500 species, they are spread in the most diverse climates. *Aristolochia* is a genus of ever green woody vines and herbaceous perennials. The flowers have a specialized pollination mechanism, the plants are aromatic and their strong scent attract insects [1]. Every part of the plant is used, seeds, leaves, root and stem [2]. *Aristolochia bracteata* is known locally in Sudan as Umm Galagil, it is used in folk medicine as anti malarial, treat snake bites and vermifugal against guinea worms and anthelmintics although it was reported that, *Aristolochia* species are nephrotoxic and carcinogenic [3]. Gum Arabic is called Gum Acacia, Acacia trees are spiny shrubs or small trees preferring sandy or sterile regions. Chemically Gum Arabic is a branched chain complex mixture of polysaccharides and glycol proteins either neutral or slightly acidic, found as a mixed calcium, magnesium and potassium salts of polysaccharidic acid [4]. Gum Arabic was widely used by Eastern folk medicine practitioners as a restorative agent and is thought to be an excellent curative for renal failure patients [5].

MATERIALS AND METHODS

Plant material

Aristolochia bracteata (Umm Galagil) the whole plant, was collected from the river Nile valley and identified at the Medicinal and Aromatic

Plants Research Institute (MAPRI), Khartoum, Sudan by Botanist.

Preparation of the plant extract

The plant was air dried, the coarse plant was powdered in a blender and subjected to successive aqueous extraction. The powder was soaked in 400 ml of hot distilled water with continuous stirring till became cool, the extract was filtered through cotton filter. Extract was stored in deep freeze till freeze [6], yield percentage was calculated as 66 grams.

Animals

White albino rats weighing 90-120 grams were maintained in a room under standard environmental condition and controlled temperature (22±2.5°C), and relative humidity (60%) and maintained on standard animal pellets with free access to water.

Effect of Gum Arabic on Nephrotoxicity induced by *Aristolochia bracteata*

Thirty six healthy adult albino rats weighing 90-100 grams were divided into 6 groups each of 6 rats. Rats were housed under standard conditions in cages at the Department of Pharmacology and Toxicology, Faculty of Veterinary Medicine, University of Khartoum. Rats were identified by colored marks on tails and left for one week adaptation period. After the adaptation period, rats were

divided according to weight into 6 groups each of 6 rats. Group 1 given orally 1ml distilled water daily for 10 days and act as a control group, group 2 were orally given 1ml distilled water for 10 days and from day 5 to day 10 given 1000mg/kg.bwt *Aristolochia bracteata* extract. Group 3 and 4 were treated with Gum Arabic 250mg/kg and 500mg/kg respectively for 10 days and *Aristolochia* extract from day 5 to day 10. Group 5 and 6 were given Gum Arabic 250mg/kg and 500mg/kg respectively for 10 days

Statistical analysis

All results were expressed as mean \pm S.E. One Way Analysis of Variance (ANOVA) was used for the analysis of data. Duncan's multiple range tests was used for determining the significance. A probability of $p < 0.05$ was considered as significant [7].

RESULTS

Biochemical findings:

The effect of GA on serum constituents of rats intoxicated with *Aristolochia bracteata* are presented in Table 1. The concentration of urea, creatinine, ALT

and AST were significantly higher in the group treated with *Aristolochia bracteata* (group 2) when compared to the control, while groups 3 and 4 recorded lower concentration of the same parameter when compared to group 2. Groups 5 and 6 that treated with GA alone showed lower concentrations in urea, creatinine, AST and ALT.

Heamatological results:

Heamatological findings are presented in Table 2. The values of RBCs, PCV, Hb, MCV, MCH and MCHC were significantly reduced in the intoxicated group (group 2) while these values were significantly improved in the groups treated with GA on *Aristolochia bracteata* and in the groups treated with GA alone.

Histopathological findings:

Histopathological sections of the kidneys of rats treated with GA on *Aristolochia bracteata* are presented in Fig 1(a-d). Fig 1-a showed necrosis, hemorrhage and shrinkage of glomeruli. Fig 1-b showed mild necrosis and hemorrhage while Fig 1-c showed lymphocyte infiltration of the glomeruli and Fig 1-d showed no abnormalities.

Table -1: Biochemical changes in rats treated with GA. on nephrotoxicity induced by *Aristolochia bracteata*

Group	Urea (mg/dl)	Creatinine (mg/dl)	ALT i.u./l	AST i.u./l
G1	42.65 \pm 1.28 ^a	0.63 \pm 0.04 ^a	61.50 \pm 1.49 ^a	73.94 \pm 2.09 ^a
G2	46.62 \pm 1.81 ^b	0.88 \pm 0.04 ^b	62.80 \pm 2.12 ^b	82.97 \pm 2.41 ^b
G3	42.48 \pm 0.35 ^{a_b}	0.52 \pm 0.07 ^{a_b}	59.30 \pm 2.34 ^{a_b}	77.47 \pm 2.71 ^{a_b}
G4	43.60 \pm 2.46	0.53 \pm 0.04 ^f	59.63 \pm 6.12 ^e	76.29 \pm 2.14 ^f
G5	40.65 \pm 1.17 ^f	0.56 \pm 0.07 ^{a_b}	53.10 \pm 3.27 ^f	74.14 \pm 2.07 ^f
G6	36.75 \pm 1.22 ^g	0.53 \pm 0.05 ^g	61.30 \pm 0.32 ^g	76.28 \pm 2.13 ^g

Means with same letter are not significantly different ($p > 0.05$)

G1 : Control, G2 : Saline + *Aristolochia bracteata* 1000 mg/kg, G3 : *Aristolochia bracteata* 1000 mg/kg + Gum Arabic 250 mg/kg., G4 : *Aristolochia bracteata* 1000 mg/kg + Gum Arabic 500 mg/kg., G5 : Gum Arabic 250 mg/kg., G6 : Gum Arabic 500 mg/kg.

Table- 2: Heamatological changes in rats treated with GA. on nephrotoxicity induced by *Aristolochia bracteata*

Group	RBCs (10/mm) ³	PCV %	HGB (g/dl)	MCV (fl)	MCH (pg)	MCHC (g/dl)
G1	6.92 \pm 0.11 ^b	51.27 \pm 2.16 ^b	12.90 \pm 0.30 ^b	76.17 \pm 0.82 ^b	18.88 \pm 0.30 ^b	24.90 \pm 0.39 ^b
G2	6.51 \pm 0.13 ^a	49.78 \pm 0.95 ^a	12.50 \pm 0.33 ^a	74.78 \pm 2.94 ^a	19.00 \pm 0.30 ^a	24.52 \pm 0.78 ^a
G3	6.88 \pm 0.20 ^{a_b}	53.22 \pm 1.45 ^{a_b}	12.58 \pm 0.37 ^{a_b}	77.42 \pm 1.18 ^{a_b}	18.28 \pm 0.04 ^{a_b}	23.63 \pm 0.34 ^{a_b}
G4	6.53 \pm 0.22 ^e	52.20 \pm 1.63 ^d	12.34 \pm 0.29 ^e	79.76 \pm 1.09 ^d	19.08 \pm 0.15 ^e	23.94 \pm 0.25 ^e
G5	6.77 \pm 0.17 ^f	51.18 \pm 1.57 ^f	12.70 \pm 0.33 ^f	75.55 \pm 0.93 ^f	18.75 \pm 0.14 ^f	24.83 \pm 0.42 ^f
G6	6.53 \pm 0.13 ^g	53.15 \pm 0.96 ^g	12.35 \pm 0.30 ^g	81.48 \pm 0.88 ^g	18.92 \pm 0.20 ^g	23.25 \pm 0.31 ^g

Means with same letter are not significantly different ($p > 0.05$)

G1 : Control, G2 : Saline + *Aristolochia bracteata* 1000 mg/kg., G3 : *Aristolochia bracteata* 1000 mg/kg + Gum Arabic 250 mg/kg., G4 : *Aristolochia bracteata* 1000 mg/kg + Gum Arabic 500 mg/kg., G5 : Gum Arabic 250 mg/kg., G6 : Gum Arabic 500 mg/kg.

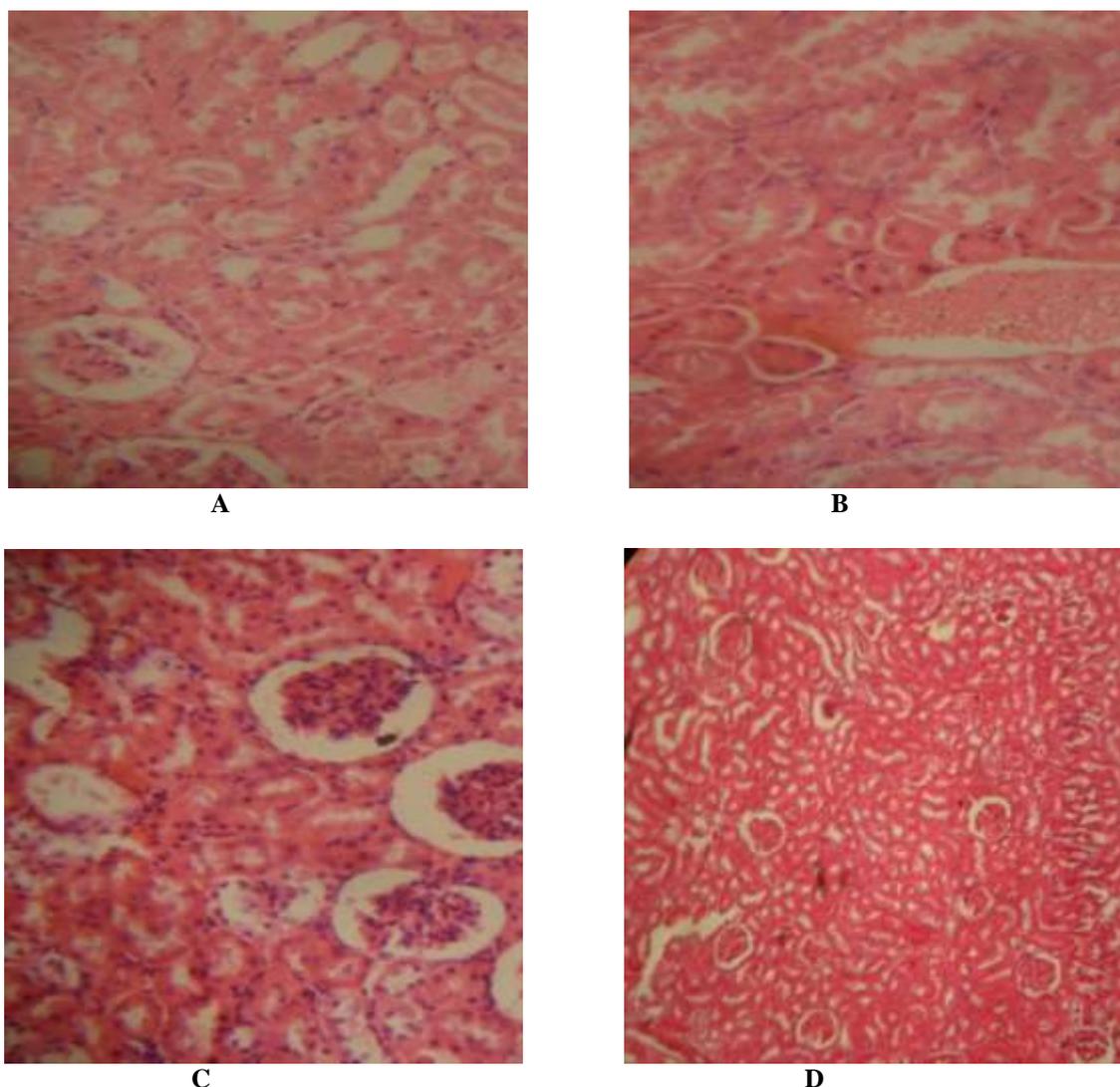


Fig 1: Sections of kidney of rats given *Aristolochia bracteata* extract

- A- Necrosis, hemorrhage and Shrinkage of the glomeruli (*Aristolochia* 1000mg/kg-G 2)**
- B- Mild necrosis of glomeruli and kidney tubules (*Aristolochia* 1000mg/kg +GA 250mg/kg-G3)**
- C- Lymphocytic infiltration of the glomeruli (*Aristolochia* 1000mg/kg +GA 500mg/kg-G4)**
- D- Normal kidney tubules and glomeruli (Normal control-G1)**

DISCUSSION

In this study oral administration of *Aristolochia bracteata* aqueous extract significantly increase the values of urea, creatinine, Ast and ALT. The Same results were found when *Aristolochia* extract given to goats [8], similarly *Aristolochia* extract in goats revealed an increase in urea concentration that correlated with pathological and histopathological changes of kidneys [9]. In the present study, oral administration of Gum Arabic to rats intoxicated with *Aristolochia* extract showed a marked decrease in concentration of urea and creatinine and in another study Gum Arabic was found to attenuated renal dysfunction in adenine induced chronic renal failure [10]. It was also reported that, Gum Arabic could be beneficial adjuvant therapy in patients with acute and chronic renal failure to prevent disease

progression and delay the need for renal replacement therapy and could conceivably alleviate adverse effects of chronic renal failure [11], [12]. Medicinal plants was found to produced nephroprotective activity as it was reported in studies of *Khaya senegalensis* [13], *Zingiber officinale* [14] and *Nigella sativa* and *Allium sativum* [15].

CONCLUSION

The aqueous extract of *Aristolochia bracteata*, whole plant at a dose of 1000 mg/kg, given orally to albino rats has a nephrotoxic effect. Gum Arabic in a dose of 250 and 500 mg/kg dissolved in distilled water and given orally to rats, make a considerable effect in lowering concentrations of urea and creatinine levels and improving the kidney damage caused by *Aristolochia bracteata*.

REFERENCES

1. Depierreux M, VanDamme B, Vanden Houte, K., Vanherweghen JL; Pathological aspects of a newly described nephropathy related to the prolonged use of Chinese herbs. *American Journal of Kidney Disease*; 1994; 24(2): 172-180.
2. Grollman AP, Shibutani S, Moriya M, Miller F, Wu L, Moll U, Suzuki N; Aristolochic acid and the Etiology of Endemic (Balkan) Nephrotoxicity, *Proceeding of the National Academy of Sciences*. 2007;104: 12129-12134. www.pnas.org/104/29/12129/suppl/DC.
3. Dickman KG, Moriya M, Zavadil J, Sidorenko VS, Edwards KL; Aristolochic acid associated Urothelial cancer in Taiwan. *Proceeding of the National Academy of Sciences, USA*, 2012; 109:841-6.
4. Ali BH, Ziada A, Blunden G; Biological effects of Gum Arabic: a review of some recent research. *Food Chem. Toxicol.* 2009; 47: 1-8.
5. Russell W, Chesney, Andrea B, Patters; *The Future of Pediatric Nephrology. Therapy*, 2006; 3(2):183-185.
6. Harborne J.B. *Phytochemical method*. 2nd. Edition. 1934 .
7. Snedecor GW, Cochran WG; *Statistical Methods* 8th. edition. 1989.
8. Adam SEI, Barakat SEM, Wasfi IA, The toxicity of *Aristolochia bracteata* in Goats. *Vet. Pathol.* 1983; 20:611-616.
9. Barri MES, Onsa TO, Elawad AA, Elsayed NY, Wasfi IA, Abdul-Bari EM, Adam SEI; Toxicity of five Sudanese plants to young ruminants. *J. Comp. Pathol*, 1983; 93:559-575.
10. Ali BH, Al Hussein I, Kayed RR, Al Mansoori N, Al Harthi T, Al Zaabi M, Nenmar A; Effect of Gum Arabic in rats with adenine-induced chronic renal failure. *Exp. Biol. Med. (Maywood)* .2010; 235(3):373-82.
11. Mahmoud MF, Diaai AA, Ahmed F; Evaluation of the efficacy of Ginger, Gum Arabic and *Boswellia* in acute and chronic renal failure. *Ren.Fail*, 2012; 34(1):73-82.
12. Ali AA, Ali K.E, Fadel alla AE, Khalid KE; The effect of Gum Arabic treatment on metabolic profile of CRF patients under regular hemodialysis in central Sudan. *Nat. Prod. Res.*, 2008; 22 (1): 12-21.
13. El Badwi SMA, Bakhiet AO, Abdel Gadir EH; Hemato-biochemical Effect of Aqueous Extract of *khaya senegalensis* Stem bark on Gentamicin –Induced Nephrotoxicity in Wistar Rats. *Journal of Biological Sciences* . 2012; 12 (6): 361-366.
14. Lakshmi BVS, Sudkakar M; Protective effect of *Zingiber officinale* on gentamicin – induced nephrotoxicity in rats. *Int.J. Pharmacol* , 2010; 6: 58-62.
15. Abelaziz I, kandeel M; The protective effect of *Nigella sativa* and *Allium sativum* extract on Amikacin –induced nephrotoxicity. *Int.J. Pharmacol* , 2011;7: 697-703.