

Renal Protective Activity of Hsian-tsao Extracts in Diabetic Rats¹

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Objective To investigate the renal protective activity of Hsian-tsao *Mesona procumbens* Hemsl. water extracts in diabetic rats. **Methods** Thirty Sprague-dawley female rats were randomly divided into three groups ($n=10$ each), "control group" with intraperitoneal saline injection, "diabetic group" with 60 mg of intraperitoneal streptozotocin injection per kg of body weight and "Hsian-tsao group" with intragastric administration of Hsian-tsao extraction everyday for 4 weeks after intraperitoneal streptozotocin injection. The body weight and blood sugar were measured before and after model induction in the three groups. Thrombospondin-1 (TSP-1) expressions in the kidney were monitored by immunohistochemistry. Kidney ultrastructural changes were also analyzed by using transmission electron microscopy. **Results** Before diabetic model induction, there were no significant differences among the three groups in body weight and blood sugar. Four weeks after the induction of diabetes, the differences became statistically significant. Electron microscopy also revealed disruption of the foot processes of the podocytes and other damages in diabetic group. These damages were significantly less severe in Hsian-tsao group when compared with the diabetic group. TSP-1 expressions in the kidney were significantly increased in both the diabetic group and Hsian-tsao group, but it was relatively lower in Hsian-tsao group than in diabetic group. **Conclusion** Our results showed that Hsian-tsao treatment in the diabetic rats effectively prevented the pathological alterations in the kidney and decreased the TSP-1 expression. It was suggested that Hsian-tsao had protective effect on the kidneys of the diabetic rats.

Key words: Rat; *Mesona procumbens* Hemsl.; Diabetic nephropathy; Thrombospondin-1

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REFERENCES

1. Witowski J, Breborowicz A (1999). The role of cellular glucose transporters in pathogenesis of diabetic nephropathy. *Przegl Lek* **56**, 793-799.
2. Yokozawa T, Nakagawa T, Wakaki K *et al.* (2001). Animal model of diabetic nephropathy. *Exp Toxicol Pathol* **53**, 359-363.
3. Chen H, Herndon M E, Lawler J (2000). The cell biology of thrombospondin-1. *Matrix Biol* **19**, 597-614.
4. Hugo C (2003). The thrombospondin 1-TGF-beta axis in fibrotic renal disease. *Nephrol Dial Transplant* **18**, 1241-1245.
5. Zhang X M, Shen F, Xv Z Y, *et al.* (2005). Expression changes of thrombospondin-1 and neuropeptide Y in myocardium of STZ-induced rats. *Int J Cardiol* **105**, 192-197.
6. Wahab N A, Schaefer L, Weston B S, *et al.* (2005). Glomerular expression of thrombospondin-1, transforming growth factor beta and connective tissue growth factor at different stages of diabetic nephropathy and their interdependent roles in mesangial response to diabetic stimuli. *Diabetologia* **48**, 2650-2660.
7. Sheu S Y, Liu C, Chiang H C (1984). The hypoglycemic principle of *Mesona procumbens* and *Orthosiphonstamineus*. *T'ai-wan K'o Hsueh* **38**, 26-31.
8. Hung C Y, Yen G C (2002). Antioxidant activity of phenolic compounds isolated from *Mesona procumbens* Hemsl. *J Agric Food Chem* **50**, 2993-2997.
9. Yen G C, Hung Y L, Hsieh C L (2000). Protective effect of extracts of *Mesona procumbens* Hemsl. on DNA damage in human lymphocytes exposed to hydrogen peroxide and UV irradiation. *Food Chem Toxicol* **38**, 747-754.
10. Valentovic M A, Alejandro N, Carpenter A B, *et al.* (2006). Streptozotocin (STZ) diabetes enhances benzo(a)pyrene induced renal injury in Sprague Dawley rats. *Toxicology Letters* **164**, 214-220.
11. Kawada J (1992). New hypotheses for the mechanisms of

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- streptozotocin and alloxan inducing diabetes mellitus. *Yakugaku Zasshi* **112**, 773-791.
12. Schena F P, Gesualdo L (2005). Pathogenetic mechanisms of diabetic nephropathy. *J Am Soc Nephrol* **16**(Suppl 1), S30-33.
 13. Thomas M C, Burns W C, Cooper M E (2005). Tubular changes in early diabetic nephropathy. *Adv Chronic Kidney Dis* **12**, 177-186.
 14. Bonnefoy A, Hantgan R, Legrand C, *et al.* (2001). A model of platelet aggregation involving multiple interactions of thrombospondin-1, fibrinogen, and GPIIb/IIIa receptor. *J Biol Chem* **276**, 5605-5612.
 15. Wang S, Skorczewski J, Feng X, *et al.* (2004). Glucose up-regulates thrombospondin 1 gene transcription and transforming growth factor-beta activity through antagonism of cGMP-dependent protein kinase repression via upstream stimulatory factor 2. *J Biol Chem* **279**, 34311-34322.
 16. Yang Y L, Chuang L Y, Guh J Y, *et al.* (2004). Thrombospondin-1 mediates distal tubule hypertrophy induced by glycated albumin. *Biochem J* **379**, 89-97.
 17. Obineche E N, Mensah-Brown E, Chandranath S I, *et al.* (2001). Morphological changes in the rat kidney following long-term diabetes. *Arch Physiol Biochem* **109**, 241-245.
 18. Lai L S, Chou S T, Chao W W (2001). Studies on the antioxidative activities of Hsian-tsao (*Mesona procumbens* Hemsl) leaf gum. *J Agric Food Chem* **49**, 963-968.

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