

论文

α-硫辛酸对糖尿病大鼠肝脏损伤保护作用

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摘要:

目的 探讨α-硫辛酸对糖尿病肝病大鼠肝纤维化的保护作用及其机制。方法 SD大鼠随机分为对照组($n=10$)、糖尿病肝病组($n=8$,腹腔注射链脲佐菌素60 mg/kg+皮下注射40%四氯化碳0.2 mL/100 g)及α-硫辛酸低、中、高剂量组($n=9$,灌胃给予15、30、60 mg/kg α-硫辛酸,连续12周),检测血糖和肝组织中丙二醛、谷胱甘肽过氧化物酶(GSH-Px)、超氧化物歧化酶(SOD)水平、肝组织中基质金属蛋白酶9(MMP-9)和金属蛋白酶组织抑制因子1(TIMP-1)蛋白表达。结果 与对照组比较,糖尿病肝病组大鼠血糖[(25.45±3.24)mmol/L]和丙二醛[(11.90±3.38)μg/mg·pro]含量均升高,SOD、GSH-Px活性[(38.45±6.73)、(19.54±2.24) μg/mg·pro]均降低,MMP-9和TIMP-1蛋白表达量及MMP-9/TIMP-1比值均降低($P<0.01$);与糖尿病肝病组比较,α-硫辛酸中剂量组大鼠血糖[(14.25±3.23)mmol/L]及丙二醛含量[(8.05±2.13) μg/mg·pro]均降低,SOD、GSH-Px活性[(46.95±6.13)、(25.14±3.23) μg/mg·pro]均升高($P<0.01$),MMP-9和TIMP-1蛋白表达量及MMP-9/TIMP-1比值均升高($P<0.01$)。结论 α-硫辛酸对糖尿病肝病大鼠肝脏组织具有保护作用,其机制可能与抗氧化过程有关。

关键词: α-硫辛酸 糖尿病肝纤维化 基质金属蛋白酶9(MMP-9) 金属蛋白酶组织抑制因子1(TIMP-1)

Protective effect of α-lipoic acid on liver injury among diabetes mellitus rats

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Abstract:

Objective To investigate protective effect and mechanism of α-lipoic acid on liver disease and hepatic fibrosis in diabetic rats. Methods The Sprague-Dawley rats were randomly divided into 5 groups: normal control group ($n=10$), model group of diabetic liver disease induced by streptozotocin (60 mg/kg) intraperitoneal injection + tetrachloride carbon ($n=8$), low-dose α-lipoic acid group (15 mg/kg · d intragastric administration for 12 weeks, $n=9$), medium-dose α-lipoic acid group (30 mg/kg · d intragastric administration for 12 weeks, $n=9$), and high-dose α-lipoic acid group (60 mg/kg · d intragastric administration for 12 weeks, $n=9$). Levels of malondialdehyde, glutathione peroxidase (GSH-Px), superoxide dismutase (SOD), and blood glucose in blood and liver tissue of the rats were measured. Expression of matrix metalloproteinase 9 (MMP-9) and tissue inhibitor of metalloproteinase factor-1 (TIMP-1) protein expression were determined with immunohistochemistry and western blot. Results Compared with the normal control group, blood glucose (25.45±3.24 mmol/L) and malondialdehyde (11.90±3.38 μg/mg · pro) were increased, while SOD (38.45±6.73 mmol/L), GSH-Px (19.54±2.24 mmol/L), MMP-9, TIMP-1, and MMP-9/TIMP-1 were reduced ($P<0.01$ for all). Compared with the model group, blood glucose (14.25±3.23 mmol/L) and malondialdehyde (8.05±2.13 mmol/L) of medium-dose α-lipoic acid group were reduced ($P<0.01$), and SOD (46.95±6.13 mmol/L), GSH (25.14±3.23 mmol/L), MMP-9, TIMP-1, and MMP-9/TIMP-1 were increased ($P<0.01$ for all). Conclusion The study shows that α-lipoic acid has protective effect on liver tissue of diabetic rat. The mechanism probably relates to the reduce of oxide stress, attenuation of extracellular matrix, and inhibition of liver fibrosis.

Keywords: α-lipoic acid diabetic liver fibrosis matrix metalloproteinase-9 tissue inhibitors of matrix metalloproteinase-1

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参考文献:

[1] Hickman IJ,Macdonald GA.Impact of diabetes on the severity of liver disease[J].Am J Med,2007,120(10) :829-834.

[2] Li CJ,Lv L,Li H,et al.Cardiac fibrosis and dysfunction in experimental diabetic cardiomyopathy are ameliorated by alpha-lipoic acid[J].Cardiovasc Diabetol,2012,11(1): 73.

[3] 董媛,蔡美琴.α-硫辛酸对外周器官氧化损伤的保护作用[J].中华临床营养杂志,2010,18(3): 187-190.

[4] 王俐,蔡美琴.α-硫辛酸在糖尿病治疗中的临床应用价值[J].中华临床营养杂志,2010,18(5): 323-326.

[5] Son SM.Reactive oxygen and nitrogen species in pathogenesis of vascular complications of diabetes[J].Diabetes Metab J,2012,36(3): 190-198.

[6] 刘友良,黄平.基质金属蛋白酶-2与肿瘤关系的研究进展[J].现代肿瘤医学,2006,14(1):109-111.

[7] Guo M,Wu MH,Korompai F,et a1.Upregulation of PKC gene and isozymes in cardiovascular tissues during early stages of experimental diabetes[J].Physiol Genomics,2003,12(2): 139-146.

[8] Peart JN,Headrick JP.Sustained cardioprotection: exploring unconventional modalities[J].Vascul Pharmacol,2008,49(3): 63-70.

[9] Eyibilen A,Cayli S,Aladag I,et al.Distribution of matrix metalloproteinases MMP-1,MMP-2,MMP-8 and tissue inhibitor of matrix metalloproteinases-2 in nasal polyposis and chronic rhinosinusitis[J].Histol Histopathol,2011,26(5): 615-621.

[10] 万为国,蒋学俊,李晓艳,等.糖尿病大鼠不同阶段肝脏内MMP-9与TIMP-1表达及全反式维甲酸的干预作用[J].疑难病杂志,2008,7(6): 323-326.

[11] Nie QH,Duan GR,Iao XD,et a1.Expression of TIMP-1 and TIMP-2 in rats with hepatic fibrosis[J].World J Gasnmnterol,2004,10(1):86-90.

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