#### 论著

CTGF反义寡核苷酸对血管紧张素 II 诱导的心肌成纤维细胞增殖、胶原 合成的抑制效应

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目的:研究结缔组织生长因子(CTGF)反义寡核苷酸对血管紧张素II诱导的心肌成纤维细胞增殖、胶原 合成及CTGF表达的抑制作用。 方法: 差速贴壁法分离新生SD大鼠心肌成纤维细胞(CFs)。反义、正义、错义 CTGF寡核苷酸分别经阳离子脂质体介导转染CFs,并与血管紧张素Ⅱ(AngⅡ)10-6mol/L共培养48 h,采用 MTT法测CFs生长数目,羟脯氨酸法测胶原蛋白含量,RT-PCR法及Western blotting法分别测CTGF mRNA及 ▶ 复制索引 蛋白水平的表达。 结果: Ang II 可以在mRNA及蛋白水平明显促进CFs CTGF的表达(P<0.01),且呈浓度-时 间依赖性。CTGF反义链组的MTT反应A值、胶原蛋白含量、CTGF mRNA及蛋白水平的表达量均明显低于Ang II 组(P<0.01),而CTGF正义链组和错义链组与Ang II 组无显著差异(P>0.05)。 结论: Ang II 刺激下产生的 CFs CTGF mRNA和蛋白水平的表达上调,可能是心肌纤维化传导通路中的关键环节,CTGF反义寡核苷酸可以 序列特异性地阻断CTGF的介导,从而抑制Ang II 对心肌成纤维细胞增殖、胶原合成及CTGF表达的诱导作用,进 相关信息 一步证实CTGF可能是拮抗心肌纤维化的特异性靶位。

心肌; 纤维化 结缔组织生长因子 寡核苷酸类,反义 关键词

分类号 R363

Inhibitory effects of antisense oligonucleotides targeting connective

tissue growth factor mRNA on the proliferation and collagen synthesis of cardiac fibroblasts

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#### Abstract

<FONT face=Verdana>AIM: To observe the effect of Ang II on the expression of connective tissue growth factor (CTGF) in cultured cardiac fibroblast (CFs), the antisense oligonucleotides (ASOND) was used to investigate whether CTGF is necessary for the proliferation and collagen synthesis of CFs. METHODS: CFs of SD rats were isolated and cultured. The proliferation and collagen synthesis of CFs were assessed by MTT assay and measuring hydroxyproline, respectively. The expression of CTGF mRNA and protein were detected by reverse-transcription PCR (RT-PCR) and Western blotting, respectively. RESULTS: Ang II could significantly increase the expression of CTGF both at mRNA and protein level in a dose and timedependent manner (P<0.01). The MTT A value, the synthesis of collagen and the expression of CTGF in ASOND group were all obviously lower than in Ang II group (P<0.01). There was no significantly difference among Ang II group, sense OND aroup and missense OND group (P>0.05), CONCLUSION: It indicates that CTGF is involved in the process of myocardial fibrosis induced by Ang II, which may be a target for the treatment of cardiac fibrosis. </FONT>

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