

沉默HOXC10基因促进骨肉瘤MG-63细胞凋亡及其机制

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Title: HOXC10 gene silencing promotes apoptosis of osteosarcoma MG-63 cells and its mechanism

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摘要: 目的: 探讨沉默同源框C10 (Homeobox C10, HOXC10) 基因表达对骨肉瘤细胞MG-63细胞增殖和凋亡的影响, 并初步探讨可能的作用机制。方法: 采用脂质体转染法将特异性针对HOXC10基因的shRNA转入骨肉瘤MG-63细胞 (HOXC10-shRNA组), 同时以转染Scramble-shRNA作为对照组 (Scramble-shRNA组), 分别应用实时荧光定量PCR及蛋白质印迹法检测HOXC10 mRNA及蛋白的表达水平。转染处理后, 分别采用CCK-8法及FCM法检测骨肉瘤MG-63细胞的增殖抑制率及凋亡情况, 采用蛋白质印迹法检测HOXC10、ATM和核因子-κB (nuclear factor kappa B, NF-κB) 及凋亡相关蛋白Survivin和Caspase-3的表达水平。结果: HOXC10-shRNA转入MG-63细胞后, HOXC10 mRNA和蛋白的表达水平均明显下调 ($P < 0.05$)。与Scramble-shRNA组细胞活力 (0.95 ± 0.12) %和凋亡率 (4.35 ± 0.52) %相比, HOXC10-shRNA组细胞活力 (0.38 ± 0.06) %明显下降, 细胞凋亡率为 (18.19 ± 3.76) %明显升高 ($P < 0.05$) ; ATM/NF-κB信号通路中相关蛋白ATM、NF-κB及Survivin的表达水平均明显下调 ($P < 0.05$), 而Caspase-3表达明显上调 ($P < 0.05$)。结论: 沉默HOXC10基因表达后可抑制骨肉瘤MG-63细胞增殖并促进其凋亡, 其机制可能与调控ATM/NF-κB信号通路的活化相关。

Abstract: Objective: To investigate the effect of Homeobox C10 (HOXC10) gene silencing on proliferation and apoptosis of osteosarcoma cell line MG-63, and to explore its possible mechanism. Methods: The specific sequence targeting HOXC10 gene (HOXC10-shRNA) was transfected into MG-63 cells, and Scramble-shRNA as control group, then the expression levels of HOXC10 mRNA and protein were detected by real-time fluorescent quantitative PCR and Western blotting, respectively. After transfection, the proliferation and apoptosis of MG-63 cells were detected by CCK-8 and FCM assay, respectively. Furthermore, the expression levels of ATM, nuclear factor-kappa B (NF-κB), Survivin and Caspase-3 proteins were detected by Western blotting. Results: The expression levels of HOXC10 mRNA and protein in MG-63 cells transfected with HOXC10-shRNA were significantly down-regulated (both $P < 0.05$). Compared with Scramble-shRNA group cell viability (0.95 ± 0.12) % and apoptotic rate (4.35 ± 0.52), cell viability in MG-63 cells transfected with HOXC10-shRNA (0.38 ± 0.06) % were significantly inhibited ($P < 0.05$), while its apoptotic rate (18.19 ± 3.76) % were markedly increased ($P < 0.05$). The expression levels of ATM, NF-κB, and Survivin proteins were significantly down-regulated (all $P < 0.05$), while the expression level of Caspase-3 was significantly up-regulated ($P < 0.05$) in HOXC10-shRNA group. Conclusion: HOXC10 gene silencing can inhibit proliferation and promote apoptosis of osteosarcoma cell line MG-63. The mechanism may be involved in inactivation of ATM/NF-κB signal pathway.

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备注/Memo: -

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