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Wnt5a 在肺癌A549细胞球增殖和上皮间质转化中的作用



分享到:

《第三军医大学学报》[ISSN:1000-5404/CN:51-1095/R] 卷: 36 期数: 2014年第18期 页码: 1890-1894 栏目: 论著 出版日期: 2014-09-30

Title: Role and mechanism of Wnt5a regulating proliferation and epithelial mesenchymal transition in A549 spheres

作者: [胡欣春](#); [徐建军](#); [喻东亮](#); [彭金华](#); [魏益平](#)
南昌大学第二附属医院心胸外科; 江西省胸科医院胸外科

Author(s): [Hu Xinchun](#); [Xu Jianjun](#); [Yu Dongliang](#); [Peng Jinhua](#); [Wei Yiping](#)

Department of Cardiothoracic Surgery, the Second Affiliated Hospital of Nanchang University, Nanchang, Jiangxi Province, 330006; Department of Chest Surgery, Jiangxi Chest Hospital, Nanchang, Jiangxi Province, 330006, China

关键词: [Wnt5a](#); [肺癌干细胞](#); [增殖](#); [上皮间质转化](#)

Keywords: [Wnt5a](#); [cancer stem cells](#); [proliferation](#); [epithelial mesenchymal transition](#)

分类号: R341; R730.23; R734.2

文献标志码: A

摘要: 目的 研究Wnt5a在A549细胞球增殖和EMT中的作用及相关机制。方法 利用无血清培养出A549细胞球后应用流式细胞术对其进行CD133⁺CD44⁺表达检测。实验分为对照组、Wnt5a组、p38组和Wnt5a+p38组。应用Western blot观察各组Wnt5a、p-p38、B-catenin、Vimentin和E-cadherin蛋白的表达变化,并利用MTT和裸鼠成瘤实验观察Wnt5a shRNA慢病毒感染前后A549细胞球的生长抑制率和成瘤能力。结果 流式细胞术检测A549细胞球中CD133⁺CD44⁺阳性率为60.5%; Wnt5a蛋白在Wnt5a组和Wnt5a+p38组中的表达较对照组明显下调($P<0.05$), p-p38蛋白、B-catenin蛋白、Vimentin蛋白在Wnt5a组、p38组和Wnt5a+p38组的表达明显低于对照组($P<0.05$), 而 E-cadherin蛋白在上述3组中的表达显著高于对照组($P<0.05$)。Wnt5a shRNA慢病毒感染A549细胞球12、24 h和48 h时抑制率显著高于对照组($P<0.05$), 并且该细胞的成瘤能力较对照组显著下调。结论 下调Wnt5a表达可抑制A549细胞球的增殖及EMT能力, 可望成为治疗非小细胞肺癌的靶点。

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更新日期/Last Update: 2014-09-18

Abstract: **Objective** To determine the role and mechanism of Wnt5a regulating the proliferation and epithelial mesenchymal transition (EMT) in lung cancer A549 spheres. **Methods** A549 spheres were cultured in serum-free medium for 2 weeks. CD133⁺CD44⁺ rate of those cells was detected by FACS. Four experimental groups including a control group, a Wnt5a group, a p38 group and a Wnt5a+ p38 group were designed. The protein expression of Wnt5a, p-p38, β -catenin, vimentin and E-cadherin among the 4 groups was detected by Western blotting. The cell inhibitory rate and tumorigenicity were detected after Wnt5a shRNA lentivirus transfection of A549 spheres. **Results** CD133⁺CD44⁺ rate was 60.5% in A549 spheres and 1.65% in A549 cells. Wnt5a protein expression was significantly lower in the Wnt5a and Wnt5a+ p38 groups than in the control group ($P<0.05$). Compared to the control group, the protein expression of p-p38, β -catenin and vimentin was obviously lower but that of E-cadherin was higher in the Wnt5a, p38 and Wnt5a+ p38 groups ($P<0.05$). The cell inhibitory rate was significantly higher and tumor size in nude mice was remarkably smaller at 12, 24 and 48 h after Wnt5a shRNA lentivirus transfection ($P<0.05$). **Conclusion** Wnt5a expression down-regulation inhibits the proliferation and EMT of A549 spheres, and Wnt5a might be a novel target for treatment of non-small-cell lung cancer.

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