



IL-15上调NKG2D表达对CIK细胞杀伤活性的增强效应

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Enhancement of Cytotoxicity against IL-15 Up-regulate NKG2D Expression on Cytokine-induced Killer Cells

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摘要

目的观察IL-15对细胞因子诱导的杀伤细胞(Cytokine-induced killer cells, CIK)NKG2D受体表达及其对食管癌EC9706细胞杀伤活性的影响。方法体外分离外周血单个核细胞,分为两组。对照组:干扰素- γ 、白细胞介素-2、CD3单抗诱导培养CIK细胞。IL-15组:加用IL-15培养。流式细胞仪检测细胞免疫表型及CD3+细胞、CD56+细胞表面NKG2D的表达,LDH法测定第14天细胞在效靶比20:1、30:1时对EC9706细胞的杀伤活性;效靶比30:1时,观察NKG2D单抗封闭细胞表面NKG2D分子后对两组细胞杀伤活性的影响。结果随着培养时间的延长,CIK群体细胞及CD56+细胞表面NKG2D表达逐渐增强,IL-15组与对照组相比差异有统计学意义($P<0.05$);效靶比20:1、30:1时,IL-15组细胞对EC9706细胞的杀伤活性均较对照组明显增强,差异均有统计学意义($P<0.05$);效靶比30:1时NKG2D单抗封闭CIK细胞表面NKG2D分子后,对照组细胞、IL-15组细胞对EC9706细胞的杀伤活性均较阻断前明显下降,差异均有统计学意义($P<0.05$)。结论IL-15上调CIK细胞表面NKG2D分子表达,增强CIK细胞对EC9706细胞的杀伤活性,CIK细胞通过NKG2D发挥作用。

关键词: 关键词: 细胞因子诱导的杀伤细胞 NKG2D 食管癌 细胞治疗 IL-15

Abstract: ObjectiveTo analyse the effects of IL-15 on the expression of NKG2D and the cytotoxicity of cytokine-induced killer (CIK) cells against human esophagus carcinoma cell EC9706 in vitro. MethodsPeripheral blood mononuclear cells were isolated from healthy donors then divided into two groups: the control group (cells were cultured in the presence of IFN- γ , anti-CD3 antibody and IL-2) and IL-15 group (cells were cultured in the presence of IFN- γ , anti-CD3 antibody, IL-2 and 10 ng/ml IL-15). Phenotypic characteristics of CIK cells and the expression of NKG2D were analyzed by flow cytometry. After 14 days culture, cytotoxicity of CIK cells against EC9706 cells was measured by using a standard LDH releasing assay. Effector cells were added to target cells at E:T ratios of 20:1, 30:1. In blocking experiments, NKG2D monoclonal antibody was added to CIK cells for 15 min before plating at 30:1 E:T ratio. ResultsNKG2D molecules were significantly up-regulated during the culture period on CD3+ cells (the CIK cells population) and the CD56+cells. The differences of NKG2D expressions were significant between IL-15 group and the control group ($P<0.05$). Cytotoxicity of CIK cells treated by the IL-15 group was higher than that of the control group, both at 20:1 E:T ratios and at 30:1 E:T ratios ($P<0.05$). When the NKG2D molecules on CIK cell ular membrane were blocked by the NKG2D monoclonal antibody, the cytolytic activity in the control group cells and IL-15 group cells was significantly inhibited. ConclusionIL-15 up-regulated the expression of NKG2D on CIK cells, which enhanced

the NKG2D mediated cytotoxicity against EC9706 cells.CIK cells played a role through the NKG2D molecules.

Key words: Key words: Cytokine-induced killer cells NKG2D Esophagus Carcinoma Cell therapy IL-15

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