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mda-7/IL-24 和 IL-24 delE5 诱导急性髓系白血病细胞向单核细胞分化 点此下载全文

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摘要:

目的:研究人黑素瘤分化相关基因-7(melanoma differentiation associated gene-7,mda-7 ;又称 IL-24)及其剪接体 IL-24 delE5 与急性髓系白血病(acute myeloid leukemia,AML)细胞分化的关系。方法:十四烷酰佛波醇-乙酯(12-O-tetradecanoylphorbol-13-acetate,TPA)处理人急性髓系白血病细胞系U937、HL-60,real-time PCR及Western blotting检测细胞中 mda-7/IL-24 和 IL-24 delE5 的表达,FACS检测细胞表面CD11b、CD1 4和CD115的表达。siRNA干扰U937、HL-60细胞中 mda-7/IL-24 和 IL-24 delE5 的表达后,再以TPA诱导细胞分化,FACS检测CD11b、CD14和CD115的表达。用前期构建的 mda-7/IL-24 和 IL-24 delE5 载体转染U937、HL-60及AML-M5患者的原代白血病细胞,观察异位过表达 mda-7/IL-24 和 IL-24 delE5 能否诱导白血病细胞向单核细胞分化。结果:TPA在诱导U937、HL-60细胞向单核细胞分化过程中显著促进 mda-7/IL-24 及其剪接体 IL-24 delE5 的表达,用siRNA干扰 mda-7/IL-24 及 IL-24 delE5 的表达有,TPA诱导U937、HL-60细胞向单核细胞分化的作用也被阻断。异位过表达 mda-7/IL-24 和 IL-24 delE5 可诱导U937、HL-60以及原代AML细胞向单核细胞分化。U937、HL-60细胞表面CD11b和CD14表达均显著升高,CD115表达仅在U937细胞有显著升高;3例AML-M5患者的原代AML细胞中CD11b、CD14及CD115表达均有不同程度升高,细胞呈现单核细胞的形态特征。结论:TPA可通过上调mda-7/IL-24 及其剪接体 IL-24 delE5 的表达诱导白血病细胞向单核细胞分化。

关键词:黑素瘤分化相关基因-7(mda-7/IL-24) <u>IL-24 delE5</u> 十四烷酰佛波醇-乙酯 急性髓系白血病 单核细胞 分化

mda-7/IL-24 and IL-24 delE5 induce myeloid leukemia cells to differentiate into monocytes Download Fulltext

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Abstract:

Objective: To investigate the effects of melanoma differentiation associated gene-7 (mda-7 , also named IL-24) and IL-24 delE5 , an mda-7/IL-24 splice variant, on differentiation of acute myeloid leukemia (AML) cells. Methods: The AML cell lines U937 and HL-60 were treated with 12-O-tetradecanoylphorbol-13-acetate (TPA); mda-7/IL-24 and IL-24 delE5 expressions were detected by real-time PCR and Western blotting assays. CD11b, CD14 and CD115 expressions on cell surface were examined by FACS. U937 and HL-60 cells were induced with TPA after siRNA interfering mda-7/IL-24 and IL-24 delE5 expressions, and then CD11b, CD14 and CD115 expressions were examined by FACS. U937 and HL-60 cell lines and primary leukemia cells from AML-M5 patients were transfected with mda-7/IL-24 or IL-24 delE5 plasmids, which were established in our previous study, in order to know whether ectopic overexpressions of mda-7/IL-24 and IL-24 delE5 can induce the monocytic differentiation of leukemia cells. Results: The expressions of mda-7/IL-24 and its splice variant, IL-24 delE5 , were significantly induced in U937 and HL-60 cells during TPA-mediated monocytic differentiation. Interfering mda-7/IL-24 and IL-24 delE5 expressions after siRNA treatment inhibited the monocytic differentiation ability of TPA on U937 and HL-60 cells. Ectopic overexpressions of mda-7/IL-24 and IL-24 delE5 induced differentiation of U937, HL-60, and primary AML leukemia cells into monocytes: CD11b, CD14 expressions on U937 and HL-60 cells and CD115 expression on U937 were significantly increased; CD11b, CD14 and CD115 expressions on primary leukemia cells from 3 AML patients were also increased with monocytic features. Conclusion: TPA can induce monocytic differentiation of leukemia cells by increasing mda-7/IL-24 and its splice variant IL-24 delE5 expressions.

Keywords: melanoma differentiation associated gene-7 (mda-7/IL-24) __IL-24 delE5 __12-O-tetradecanoylphorbol-13-acetate (TPA) acute myeloid leukemia monocyte differentiation

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