

387~391.Tautomycetin诱导乳腺癌耐药细胞MCF-7/ADR的凋亡及其机制[J].牛铭山,孙岩,唐莉,邱荣国.中国肿瘤生物治疗杂志,2012,19(4)

Tautomycetin诱导乳腺癌耐药细胞MCF-7/ADR的凋亡及其机制 [点此下载全文](#)

[牛铭山](#) [孙岩](#) [唐莉](#) [邱荣国](#)

大连理工大学 化工与环境生命学部 分子药物中心,辽宁 大连 116023;大连理工大学 化工与环境生命学部 分子药物中心,辽宁 大连 116023;大连理工大学 化工与环境生命学部 分子药物中心,辽宁 大连 116023;大连理工大学 化工与环境生命学部 分子药物中心,辽宁 大连 116023

基金项目: 国家杰出青年科学基金资助项目 (No. 30688003)

DOI: 10.3872/j.issn.1007-385X.2012.4.008

摘要:

目的: 研究tautomycetin对乳腺癌耐药细胞MCF-7/ADR增殖及凋亡的影响及其机制。方法: MTT法检测tautomycetin对MCF-7/ADR细胞增殖的影响,流式细胞术检测MCF-7/ADR细胞的凋亡,Western blotting法检测tautomycetin对MCF-7/ADR细胞caspase相关蛋白、Bcl-2、Cyto-C、P53蛋白表达和Akt磷酸化的影响。结果: Tautomycetin可剂量($0.01\sim100\text{ }\mu\text{mol/L}$)依赖性地抑制MCF-7/ADR细胞的增殖($P < 0.05$), IC_{50} 值为($1.26\pm0.12\text{ }\mu\text{mol/L}$;与对照组相比,tautomycetin($1\text{ }\mu\text{mol/L}$)可诱导MCF-7/ADR细胞凋亡,早期凋亡比例由($0.67\pm0.18\%$)升高至($17.2\pm3.8\%$),晚期凋亡比例由($0.96\pm0.23\%$)升高至($28.4\pm5.7\%$)($P < 0.05$)。Tautomycetin可活化MCF-7/ADR细胞中caspase-7和caspase-9,降低Bcl-2蛋白的表达,促进线粒体释放Cyto-C,降低p-Akt的水平,但对caspase-8和P53的表达没有影响。结论: Tautomycetin可阻断Akt活化,以P53非依赖的方式通过Cyto-C介导的通路诱导MC F-7/ADR细胞凋亡。

关键词: [乳腺癌](#) [tautomycetin](#) [增殖](#) [凋亡](#) [caspase](#) [细胞色素C](#)

Tautomycetin induces apoptosis of human breast cancer cell line MCF-7/ADR and its mechanism [Download Fulltext](#)

[NIU Ming-shan](#) [SUN Yan](#) [TANG Li](#) [QIU Rong-guo](#)

Research Center for Molecular Medicine, Faculty of Chemical, Environmental and Biological Science and Technology, Dalian University of Technology, Dalian 116023, Liaoning, China; Research Center for Molecular Medicine, Faculty of Chemical, Environmental and Biological Science and Technology, Dalian University of Technology, Dalian 116023, Liaoning, China; Research Center for Molecular Medicine, Faculty of Chemical, Environmental and Biological Science and Technology, Dalian University of Technology, Dalian 116023, Liaoning, China; Research Center for Molecular Medicine, Faculty of Chemical, Environmental and Biological Science and Technology, Dalian University of Technology, Dalian 116023, Liaoning, China

Fund Project:Project supported by the National Foundation for Distinguished Young Scientists of China (No.30688003)

Abstract:

Objective: To investigate the effects of tautomycetin on the proliferation and apoptosis of human breast cancer cell line MCF-7/ADR and the related mechanism. Methods: The effect of tautomycetin on the proliferation of MCF-7/ADR cells was examined by MTT assay; its effect on apoptosis of MCF-7/ADR cells was assessed by flow cytometry; and its effects on expressions of caspase-related proteins, Bcl-2, cytochrome C (Cyto-C), P53 and Akt in MCF-7/ADR cells were detected by Western blotting. Results: Tautomycetin inhibited the proliferation of MCF-7/ADR cells in a dose-dependent manner ($0.01\sim100\text{ }\mu\text{mol/L}$, $P < 0.05$), with the IC_{50} value of ($1.26\pm0.12\text{ }\mu\text{mol/L}$). Compared with the control group, the early apoptosis rate of MCF-7/ADR cells after $1\text{ }\mu\text{mol/L}$ tautomycetin treatment was increased from ($0.67\pm0.18\%$) to ($17.2\pm3.8\%$), and the late apoptosis rate from ($0.96\pm0.23\%$) to ($28.4\pm5.7\%$), ($P < 0.05$); tautomycetin activated caspase-9 and caspase-7, decreased Bcl-2 expression, promoted Cyto-C secretion and decreased p-Akt levels in MCF-7/ADR cells, while showed no obvious effect on caspase-8 and P53 expressions. Conclusion: Tautomycetin can inhibit the phosphorylation of Akt, and induce the Cyto-C-mediated apoptosis of MCF-7/ADR cells in a P53-independent pathway.

Keywords:[breast cancer](#) [tautomycetin](#) [proliferation](#) [apoptosis](#) [caspase](#) [cytochrome C](#)

[查看全文](#) [查看/发表评论](#) [下载PDF阅读器](#)