论著

白藜芦醇诱导鼻咽癌细胞CNE-2Z凋亡的线粒体机制

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摘要 目的 探讨白藜芦醇诱导鼻咽癌细胞CNE-22凋亡的线粒体机制。方法 用100 μ mol•L⁻¹白藜芦醇分别处理细胞0(对照),2,4,8,12,24 h和分别用0,25,50,100,200 μ mol•L⁻¹白藜芦醇处理CNE-2Z细胞8 h,采用Western印迹分析Bcl-2,Bax和胞浆中细胞色素C(Cyt C)的蛋白水平变化,罗丹明123荧光染色后经流式细胞术检测线粒体膜电位($\Delta\psi$ m)变化,比色法测定半胱天冬酶-9的活性改变。结果 ①100 μ mol•L⁻¹白藜芦醇处理细胞不同时间,Bcl-2蛋白表达和 $\Delta\psi$ m减少、 胞浆中的Cyt C和半胱天冬酶-9活性增高均呈时间依赖性(P0.01),但Bax蛋白表达无改变。除Bax以外的其他指标均自白藜芦醇处理2 h即有变化。Bcl-2的蛋白表达受抑和 $\Delta\psi$ m的丧失在4~8 h间最明显(与对照组比较P0.01),在24 h时已无意义。胞浆中Cyt C水平和半胱天冬酶-9活性在8 h达高峰(分别为0 h的3.0,5.4倍),24 h时仍明显高于对照组(P0.01)。②细胞经不同浓度的白藜芦醇处理8 h后,Bcl-2的蛋白表达受抑、 $\Delta\psi$ m的丧失、Cyt C的释放和半胱天冬酶-9活性的升高均具有剂量依赖性(P0.01),但Bax的蛋白表达未受影响。结论 白藜芦醇可能经一个线粒体/半胱天冬酶-9的特定途径诱导CNE-Z细胞凋亡,但此凋亡过程可能与Bax无关。

关键词 <u>白藜芦醇</u> <u>线粒体</u> <u>细胞色素C</u> <u>蛋白Bcl-2</u> <u>蛋白Bax</u> <u>半胱天冬酶类</u>

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Mitochondrial mechanism of apoptosis induced by resveratrol in nasopharyngeal carcinoma cells CNE-2Z

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Abstract

AIM To study whether resveratrol-induced apoptosis in nasopharyngeal carcinoma cells CNE-2Z is via a mitochondrial pathway. METHODS CNE-2Z cells were treated with 100 µmol•L-1 resveratrol for 0(control), 2, 4, 8, 12, and 24 h and treated with 0(control), 25, 50, 100, and 200 µmol•L⁻¹ resveratrol for 8 h. The Bcl-2, Bax and cytosolic cytochrome C protein levels were analyzed by Western blot. The mitochondrial membrane potential ($\triangle \psi m$) was detected by flow cytometry following rhodamine 123 fluorescence staining. The caspase-9 activity was determined by colorimetric assay. **RESULTS** ① After CNE-2Z cells were treated with 100 µmol•L⁻¹ resveratrol for 0(control), 2, 4, 8, 12, and 24 h, respectively, Bcl-2 protein expression and △ym were decreased, while cytosolic cytochrome C level and caspase-9 activity were increased in a time-dependent manner (n=3, P<0.01). But Bax protein expression had no change. All except Bax protein expression had been altered since 2 h. The inhibition of Bcl-2 protein expression and the loss of $\triangle \psi m$ were most obvious during 4-8 h (compared with control group P<0.01), but no significance at 24 h. Cytosolic cytochrome C level and caspase-9 activity reached the peak at 8 h and were still much higher than those in control group at 24 h (P<0.01). Cytosolic cytochrome C level and caspase-9 activity were about 3.0-fold and 5.4-fold of those at 0 h, respectively. ② After CNE-2Z cells were treated with 0(control), 25, 50, 100, and 200 µmol•L⁻¹ resveratrol for 8 h, respectively, the Bcl-2 protein expression inhibition, $\triangle \psi m$ loss, cytochrome C release, and caspase-9 activity increase showed in a dosedependent manner (n=3, P<0.01). However, Bax protein expression was not affected. **CONCLUSION** Resveratrol can induce apoptosis in CNE-2Z via a mitochondria/caspase-9-specific pathway, but this may not be related to Bax.

Key words resveratrol mitochondria cytochrome C

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