

论文

蛋白激酶抑制剂staurosporine增强抗癌药对肿瘤细胞的杀伤

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摘要:

蛋白激酶抑制剂staurosporine 5 ng·ml⁻¹阻断人胚肺2BS细胞于G₁/S边界,而不影响人胃癌BGC-823细胞的周期运行。细胞周期时相特异药物阿霉素、阿糖胞苷或博来霉素A₅与staurosporine合用,2BS细胞和BGC-823细胞的IC₅₀均发生改变,显示低剂量staurosporine增强抗癌药对肿瘤细胞的杀伤。用谷胱甘肽(GSH)的荧光探针mBCL测定不同细胞周期时相的GSH,发现staurosporine使2BS细胞中GSH含量显著增高,而使BGC-823细胞中GSH含量显著下降。Staurosporine对正常和肿瘤细胞周期行进及胞内GSH水平的不同影响,可能是它增强抗癌药物对肿瘤细胞杀伤作用的原因。

关键词: 蛋白激酶抑制剂(staurosporine) 抗肿瘤药物 细胞周期 谷胱甘肽

PROTEIN KINASE INHIBITOR STAUROSPORINE ENHANCES CYTOTOXICITY OF ANTI TUMOR DRUGS TO CANCER CELLS

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Abstract:

Treated with low dosage(5 ng·ml⁻¹)of staurosporine for 18 h, human embryolung 2BS cells were blocked at the G₁/S boundary, but human gastric carcinoma BGC-823 cells still kept their cell cycle. In comparison with IC₅₀ of 2BS and BGC-823 cells treated with cell cycle phasespecific antitumor drugs adriamycin, Ara-C and BLM A₅ alone or combined with staurosporine(5ng·ml⁻¹), the IC₅₀ values increased from 0.325 μg·ml⁻¹, 5 μg·ml⁻¹ and 6.5 μg·ml⁻¹ to 0.45 μg·ml⁻¹, 10 μg·ml⁻¹ and 6.5 μg·ml⁻¹, respectively in 2BS cells; but decreased from 0.325 μg·ml⁻¹, 25 μg·ml⁻¹ and 1.1 μg·ml⁻¹ to 0.07 μg·ml⁻¹, 6.25 μg·ml⁻¹ and 0.4 μg·ml⁻¹, respectively in BGC-823 cells. These results suggest that combination of staurosporine 5 ng·ml⁻¹ with antitumor drugs showed different effects on tumor cells and normal cells. With the GSH fluorescent probe mBCL, we found that GSH contents increased in 2BS cells treated with staurosporine 5ng·ml⁻¹.

Keywords: Cell cycle Antitumor drug Glutathione(GSH) Staurosporine

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