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### 论文

蝙蝠葛苏林碱及异构体对PC12细胞缺血性损伤的保护作用

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摘要:

用细胞培养术,以NaCN加缺糖引起PC12细胞损伤作为细胞缺血性损伤的模型,观察蝙蝠葛苏林碱(DS)及其3个光学 异构体对PC12细胞缺血性损伤的保护作用:以Fura-2/AM为Ca<sup>2+</sup>的荧光探针,用AR-CM-MIC阳离子测定系统观察 DS及异构体对NaCN引起的 $Ca^{2+}$ 内流的抑制作用。结果表明DS及3个异构体在 $10^{-8}\sim 10^{-5}$  mol· $L^{-1}$  的浓度范围内 对PC12细胞损伤有较明显的保护作用,其机制之一与抑制NaCN加缺糖诱发的细胞内游离Ca<sup>2+</sup>异常升高有关。

关键词: 蝙蝠葛苏林碱 嗜铬细胞癌PC12细胞 细胞培养 细胞内游离Ca<sup>2+</sup> 兴奋性氨基酸

## EFFECTS OF DAURISOLINE AND ITS THREE OPTICAL ISOMERS ON ISCHEMIC INJURY IN CULTURED PHEOCHROMOCYTOMA (PC12) CELLS

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#### Abstract:

In this study, the protective effects of (-)R·R-daurisoline and its three optical isomers on ischemic injury in cultured PC12 cells induced by treating cells with NaCN in glucose-free medium were investigated. Cell viability was measured using MTT assay. The results indicated that these compounds, especially (-)-S.R and (+)-R.S isomers were found substantially to attenuate ischemic injury in PC12 cells in a dosedependent manner. The IC<sub>50</sub> values of (-)- $R \cdot R$ , (-)- $S \cdot R$ , (+)- $R \cdot S$  and (+)- $S \cdot S$  isomers were shown to be  $18.6 \times 10^{-6}$ ,  $2.4 \times 10^{-6}$ ,  $5.9 \times 10^{-6}$  and  $90 \times 10^{-6}$  mol·L<sup>-1</sup>, respectively. Intracellular free Ca<sup>2+</sup> concentration in PC12 cells was measured using AR-CM-MIC cation measurement system with Fura-2/AM  $\blacktriangleright$  刘国卿 as  $Ca^{2+}$  fluorecent indicator.(-)- $R\cdot R$ -daurisoline and its three optical isomers: (-)- $S\cdot R$ , (+)- $R\cdot S$  and (-)- $S \cdot S$  were found to markedly inhibit the increase of cytosolic free  $Ca^{2+}$  concentration induced by NaCN (20 mmol·L<sup>-1</sup>) in a dose-dependent manner. Their  $IC_{50}$  were found to be  $3.55 \times 10^{-6}$ ,  $0.59 \times 10^{-6}$ ,  $1.29 \times 10^{-6}$  and  $24.3 \times 10^{-6}$  mol L<sup>-1</sup> respectively. It is suggested that the cytoprotective effects of daurisoline and its isomers were mediated by blocking Ca<sup>2+</sup> influx into cells.

Keywords: Exicitatory amino acid Cell culture Pheochromocytoma cells(PC12) Intracellular free Ca<sup>2+</sup> Daurisoline

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