

论文

不同甲状腺机能状态对地西洋药代动力学的影响

许风云;张远;楼雅卿

北京医科大学药理学系,北京100083

摘要:

用大鼠建立甲状腺机能低下(甲低)和甲状腺机能亢进(甲亢)的动物模型。以HPLC法测定地西洋血药浓度,研究甲低和甲亢对地西洋药代动力学的影响。结果表明,甲低组大鼠地西洋血药浓度显著高于对照组(P<0.05),Cmax升高,AUC增大,吸收T<sub>1/2</sub>延长,Vd减小,消除减慢。轻度甲亢组地西洋血药浓度、Cmax及AUC与对照组相比无差异,但随着甲亢程度加重,上述指标逐渐增高。轻、中度甲亢组地西洋吸收基本不变,Vd减小,消除加快;重度甲亢组吸收明显加快,Vd减小,消除减慢。提示不同甲状腺机能状态对地西洋药代动力学的影响不同。

关键词: 甲状腺机能低下 甲状腺机能亢进 地西洋 药代动力学

EFFECTS OF DIFFERENT THYROID STATUS ON THE PHARMACOKINETICS OF DIAZEPAM

Xu Fengyun; Zhang Yuan and Lou Yaqing

Abstract:

Experimental models of hypothyroidism and hyperthyroidism in Sprague-Dawley rats were established in this study. Diazepam was given to rats at a single oral dose of 30~40 mg·kg<sup>-1</sup> and the plasma concentration of diazepam was detected by HPLC. The results showed that the plasma concentration of diazepam was significantly higher in hypothyroid rats than that in controls (P<0.05). The Cmax, AUC and T<sub>1/2</sub>(Ka) were increased. The Vd was decreased and the elimination was slowed. Mild hyperthyroidism showed nearly no effect on the plasma concentration, Cmax and AUC of diazepam in the rats. But when the rats became more heavily hyperthyroid, the plasma concentration, Cmax and AUC of diazepam were increased gradually. The absorption of diazepam was changed slightly in mild and moderate hyperthyroid rats, the Vd was decreased and the elimination was accelerated. In heavily hyperthyroid rats, however, the absorption of diazepam was obviously accelerated. The Vd was decreased and the elimination was slowed. Therefore, we conclude that different thyroid status may have different effects on the pharmacokinetics of diazepam.

Keywords: Hyperthyroidism Diazepam Pharmacokinetics Hypothyroidism

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