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基于黄连解毒汤药动学的差异性优化中药复方制备工艺

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中文摘要:目的: 以黄连解毒汤为例,基于中药复方药物动力学原理优化中药复方制备工艺。 方法: 大鼠分别灌胃给予不同制备工 艺所得黄连解毒汤全方提取物 I, II,III,IV,于不同时间点采集血浆样品,以黄连解毒汤全方中含量较高的栀子苷为检测指标,采用HP LC测定不同时间点的血药浓度,绘制药-时曲线,应用DAS 2.0软件拟合药动学参数。 结果: 不同制备工艺所得黄连解毒汤全方提取 物中栀子苷的药动学特征相差较大。全方水提取沉淀物组的药-时曲线下面积(AUC)最小且消除最快;全方水提去沉淀物组的AUC  $_{(0\sim\infty)}$ 与全方水提物组及全方水提醇沉精制物组的 $\mathrm{AUC}_{(0\sim\infty)}$ 相比存在显著性差异 (P<0.05);全方水提醇沉精制物组中栀子苷的药-时曲线呈现平缓的趋势;全方水提物组的最大血药浓度( $C_{max}$ )最大,但是消除较全方水提醇沉精制物组快。结论:通过不同工艺产 物中栀子苷的药代动力学特征的比较,提示中药复方的制备工艺可根据临床给药剂型、给药时间间隔的需求而采用不同的制备工 艺,为初步评价中药复方全方的制备工艺研究提供了一种新思路。

中文关键词:黄连解毒汤 制备方法 栀子苷 药代动力学 中药复方

## Evaluation of Muti-herbs Remedy Prepared Differently based on Pharmacokinetic Variations of Huanglian Jiedu Decoction

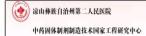
Abstract: Objective: To investigate the preparation progresses of muti-herbs remedy, based on pharmacokinetic theory, taken Huanglian Jiedu decoction (HLJDT) example. Method: The plasma samples were collected at different time points, after oral administration of products of HLJDT by four different preparation methods, RP-HPLC was used to determine plasma levels of geniposide which was an index component and higher levels of HLJDT, then concentration-time curve was drawn. And DAS 2.0 software was used to simulate the corresponding pharmacokinetic parameters. Result: Specificity and accuracy of the method were in line with the requirements of biological sample analysis. The analytical results showed that plasma concentration was a larger difference between the oral products made from the same formula HLJDT, but prepared in a different way. The AUC of sediment group of aqueous extract of the multi-herbs remedy (III) was minimum and elimination of that was fastest; there was a significant difference comparing the  $AUC_{(0\sim\infty)}$  between aqueous extract of the multi-herbs remedy without sediment (II) and aqueous extract of the multi-herbs remedy ( I ) and the whole side of water extraction and alcohol precipitation refined extract group (IV) (P<0.05); the concentration-time curve of geniposide in sample IV showed flat trend; the  $C_{\rm max}$  of geniposide in aqueous extract of the multi-herbs remedy was the largest, but elimination of it was more faster than that of sample IV. Conclusion: comparing the pharmacokinetics of the four products prepared by different ways, prompted that the preparation processes of multi-herbs remedy on pharmacokinetics studies could be based on the demands of clinical drug formulations and that of dosing interval, which preliminarily provided a new idea to evaluate preparation processes of multi-herbs remedy.

keywords:Huanglian Jiedu decoction preparation means geniposide pharmacokinetic muti-herbs remedy

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