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论文

高灵敏度LC/MS/MS法同时测定人血浆中麻黄碱和氯苯那敏

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摘要:

关键词: 麻黄碱 氯苯那敏 液相色谱-串联质谱法 药代动力学

Simultaneous determination of ephedrine and chlorpheniramine in human plasma by a highly sensitive liquid chromatography-tandem mass spectrometric method

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Abstract:

AimTo develop and validate a liquid chromatography-tandem mass spectrometric (LC/MS/MS) method for the simultaneous quantification of ephedrine and chlorpheniramine in human plasma after oral administration of a compound preparation. MethodsThe analytes and the internal standard, diphenhydramine, were isolated from plasma by protein precipitation with methanol, then chromatographied on a Zorbax SB-C $_{18}$ column (150 mm \times 4.6 mm ID) using a mobile phase consisted of methanol-water-formic acid (80:20:0.5, v/v), at a flow rate of 0.5 mL·min⁻¹. A tandem mass spectrometer equipped with electrospray ionization source was used as detector and was operated in the positive ion mode. Selected reaction monitoring (SRM) using the precursor to produce ion combinations of m/z 166 \rightarrow 115, m/z 275 \rightarrow 230 and m/z 256 \rightarrow 167 were used to quantify ephedrine, chlorpheniramine and the internal standard, respectively. ResultsThe linear concentration ranges of the calibration curves for ephedrine and chlorpheniramine were 0.50-200 $\mu g \cdot L^{-1}$ and 0.050-20.0 $\mu g \cdot L^{-1}$, respectively. The lower limits of quantification were 0.50 $\mu g \cdot L^{-1}$ for ephedrine and 0.050 $\mu g \cdot L^{-1}$ for chlorpheniramine, individually. The intra- and inter-day relative standard deviation (RSD) across three validation runs over the entire concentration range was less than 9.3% for both ephedrine and chlorpheniramine. The interday accuracy (RE) was within ±3.4% for the analytes. Each sample was chromatographied within 3.3 min. The method was successfully used in pharmacokinetics study of ephedrine and chlorpheniramine in human plasma after oral administration of a compound preparation containing 5 mg ephedrine hydrochloride, 1 mg chlorpheniramine maleate, 50 mg phenytoin, 12.5 mg theophylline, 12.5 mg theobromine and 7.5 mg caffeine. No interaction among the six components was observed on their pharmacokinetic parameters. ConclusionThe method was proved to be highly sensitive, selective, and suitable for pharmacokinetics investigations of different compound preparations containing low dosage of both ephedrine and chlorpheniramine.

Keywords: chlorpheniramine LC/MS/MS pharmacokinetics ephedrine

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