

论文

间硝苯地平在Beagle犬体内的药代动力学

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摘要:

目的用反相高效液相色谱法研究间硝苯地平(m-nifedipine, m-Nif)在Beagle犬体内的药代动力学特征。方法正交设计优化色谱分离条件, Beagle犬分别iv给予m-Nif 0.288 mg·kg<sup>-1</sup>和ig m-Nif 1.152, 3.456, 10.370 mg·kg<sup>-1</sup>。用反相高效液相色谱法分析血浆中原型药物浓度, 血浆药物浓度-时间数据用3P97药代动力学软件分析。结果Beagle犬iv m-Nif, 其体内过程符合二室模型, T<sub>1/2β</sub>为116.8 min; ig给予m-Nif 后在Beagle犬体内的代谢符合一室模型, 其中低剂量(1.152 mg·kg<sup>-1</sup>)组C<sub>max</sub>为20 μg·L<sup>-1</sup>, T<sub>1/2</sub>(k<sub>e</sub>)为147 min; 中剂量(3.456 mg·kg<sup>-1</sup>)组C<sub>max</sub>为36 μg·L<sup>-1</sup>, T<sub>1/2</sub>(k<sub>e</sub>)为122 min; 高剂量(10.37 mg·kg<sup>-1</sup>)组C<sub>max</sub>为69 μg·L<sup>-1</sup>, T<sub>1/2</sub>(k<sub>e</sub>)为144 min。结论Beagle犬ig和iv m-Nif 后, 血浆中药物消除迅速, 口服绝对生物利用度较低。

关键词: 间硝苯地平 药物代谢动力学 高效液相色谱法

Pharmacokinetics of m-nifedipine in Beagle dogs

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Abstract:

AimTo study the pharmacokinetics of m-nifedipine (m-Nif) in Beagle dogs. Methods The Beagle dogs were divided into two groups. m-Nif was intravenously administered to the Beagle dogs in group 1 at the dose of 0.288 mg·kg<sup>-1</sup>, and it was orally administered to the Beagle dogs in group 2, 3 and 4 at the dose of 1.152, 3.456 and 10.370 mg·kg<sup>-1</sup>, respectively. m-Nif in plasma was detected by reversed phase high performance liquid chromatography. The pharmacokinetic parameters were calculated by 3P97 software. ResultsWhen m-Nif was intravenously administered, the plasma concentration-time curve was fit to a two-compartment model and T<sub>1/2β</sub> was 117 min. When m-Nif was orally administered, the plasma concentration-time curve was fit to a one-compartment model. T<sub>1/2</sub>(k<sub>e</sub>) and C<sub>max</sub> were 147 min and 20 μg·L<sup>-1</sup>; at the low dose of 1.152 mg·kg<sup>-1</sup>. T<sub>1/2</sub>(k<sub>e</sub>) was 122 min and C<sub>max</sub> was 36 μg·L<sup>-1</sup> at the middle dose of 3.456 mg·kg<sup>-1</sup>. T<sub>1/2</sub>(k<sub>e</sub>) was 144 min and C<sub>max</sub> was 69 μg·L<sup>-1</sup> at the high dose of 10.37 mg·kg<sup>-1</sup>, respectively. ConclusionIt was showed that the speed of elimination of m-Nif was high in Beagle dogs. The absolute bioavailability of m-Nif given orally was very low.

Keywords: pharmacokinetics; high performance liquid chromatography m-nifedipine

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