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麝香及西黄方中麝香酮在正常、乳腺癌癌前病变大鼠体内药动学的 比较

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摘要:目的 比较麝香及西黄方中麝香酮在正常、乳腺癌癌前病变大鼠体内药动学。方法 7,12-二甲基苯并蒽 (DMBA) 联合雌激素、孕激素建立大鼠乳腺癌癌前病变模型。大鼠随机分为麝香组、西黄方正常组、西黄方模型组,灌胃给药(含 2.73 mg/kg 麝香酮)30 d,于 0.083、0.25、0.5、1、1.5、2、4、6、8、24 h 采血,GC-MS 法测定麝香 酮血药浓度,计算主要药动学参数。结果 与麝香组比较,西黄方正常组 AUC_{0-1} 、 AUC_{0-2} 、 MRT_{0-1} 、 MRT_{0-2} 、 $t_{1/22}$ 升高 (P<0.01), CL_z/F 降低 (P<0.05);与西黄方正常组比较,西黄方模型组 AUC_{0-1} 、 AUC_{0-2} 、 MRT_{0-1} 升高 (P<0.05), CL_z/F 降低 (P<0.05)。结论 西黄方中麝香酮生物利用度高于麝香中,体现了复方配伍的协同作用。在乳腺癌癌前病变大鼠中,麝香酮消除减慢,体内作用时间延长。

关键词:麝香:西黄方:麝香酮:乳腺癌癌前病变:体内药动学:GC-MS

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Comparison of *in vivo* pharmacokinetics of muscone from *Moschus* and Xihuang Decoction in normal rats and rats with precancerous lesions of breast cancer

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ABSTRACT: AIM To compare the *in vivo* pharmacokinetics of muscone from *Moschus* and Xihuang Decoction in normal rats and rats with precancerous lesions of breast cancer. METHODS The rat models of precancerous lesions of breast cancer were established by 7, 12-dimethyl-benzoanthracene (DMBA) combined with estrogen and progesterone. Rats were randomly assigned into *Moschus* group, Xihuang Decoction normal group, Xihuang Decoction model group and given intragastric administration (containing 2.73 mg/kg muscone) for 30 d, after which blood collection was performed at 0.083, 0.25, 0.5, 1, 1.5, 2, 4, 6, 8, 24 h, GC-MS was adopted in the plasma concentration determination of muscone, and main pharmacokinetic parameters were calculated. **RESULTS**

Compared with the *Moschus* group, the Xihuang Decoction normal group demonstrated increased $\mathrm{AUC}_{0-\iota}$, $\mathrm{AUC}_{0-\infty}$, $\mathrm{MRT}_{0-\iota}$, $\mathrm{MRT}_{0-\infty}$, $t_{1/2z}$ (P < 0.01) and decreased CL_z/F (P < 0.05). Compared with the Xihuang Decoction normal group, the Xihuang Decoction model group displayed increased $\mathrm{AUC}_{0-\iota}$, $\mathrm{AUC}_{0-\infty}$, $\mathrm{MRT}_{0-\iota}$ (P < 0.05) and decreased CL_z/F (P < 0.05). **CONCLUSION** The bioavailability of muscone in Xihuang Decoction is higher than that in *Moschus*, exhibiting the synergistic effect of compound compatibility. In the rats with precancerous lesions of breast cancer, muscone shows slowed down elimination and prolonged *in vivo* action time.

KEY WORDS: *Moschus*; Xihuang Decoction; muscone; precancerous lesions of breast cancer; *in vivo* pharmacokinetics; GC-MS

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