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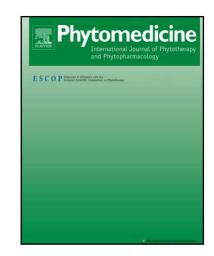
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ACCEPTED MANUSCRIPT

Repeated administration of Sailuotong, a fixed combination of *Panax* ginseng, Ginkgo biloba, and Crocus sativus extracts for vascular dementia, alters CYP450 activities in rats

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ABSTRACT

Background: Sailuotong (SLT) is a standard Chinese preparation made from extracts of *Panax Ginseng* (ginseng), *Ginkgo biloba* (ginkgo), and *Crocus sativus* (saffron). Preliminary clinical trials and animal experiments have demonstrated that SLT could improve cognition of vascular dementia (VD).

Purpose: To avoid incident drug-drug interaction which is easily encountered in patients of VD, the potential influence of SLT on main drug-metabolic cytochromes P450 enzymes (CYP450) was investigated.

Method: A "cocktail probes" approach was employed to evaluate the activities of CYP450. A rapid and selective analysis method was developed to examine 5 CYP probe drugs and their specific metabolites in plasma by using online SPE followed by a single LC-MS/MS run. After pretreatment for 2 weeks with SLT, ginseng, gingko, saffron or water (control), a cocktail solution containing caffeine, losartan,

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