



Biopharmaceutical *In Vitro* and *In Vivo* Characterization of Isopropylidene Shikimic Acid

Hui KONG^{1,2}, Hai C. QU¹, Pei YANG¹, Yan H. WU¹, Fei L. LIN¹,
Li S. CAO¹, Jun L. FENG¹, Jing FU¹ & Jian NI^{1*}

¹ *School of Chinese Materia Medica, Beijing University of Chinese Medicine, Beijing 100102, China*

² *School of Preclinical Medicine, Beijing University of Chinese Medicine, Beijing 100102, China*

SUMMARY. Biopharmaceutical properties of isopropylidene shikimic acid (ISA) were investigated in rats after oral administration. Our results showed that 27.35% ISA was bioavailable after oral administration (AUC, 836.13 mg/L.min) in comparison with intravenous injection (AUC, 3057.02 mg/L.min). In addition, we demonstrated that ISA was stable in artificial intestinal solutions, and hardly degraded in liver, intestinal microsomes and hepatocytes. However, ISA was unstable in artificial gastric juice with about 20-50% degraded in 2 h. Following oral administration, 22.9, 31.66, and 0.19% intact ISA were excreted in urine, feces, and bile, respectively. These findings suggest that low oral bioavailability of ISA might result from overdose and poor stability in stomach acids.

RESUMEN. Las propiedades biofarmacéuticas del ácido shikímico isopropilideno (ISA) fueron investigadas en ratas después de la administración oral. Nuestros resultados mostraron que el 27,35% de ISA era biodisponible después de la administración oral (AUC, 836,13 mg/L.min) en comparación con la inyección intravenosa (AUC, 3057.02 mg/L.min). Además, hemos demostrado que ISA era estable en solución intestinal artificial y casi no se degrada en hígado, los microsomas intestinales y hepatocitos. Sin embargo, ISA resultó inestable en el jugo gástrico artificial con aproximadamente 20-50% de degradación en 2 h. Tras la administración oral, 22.9, 31.66, y 0.19% a ISA intacta se excretan en la orina, las heces y la bilis, respectivamente. Estos hallazgos sugieren que la baja biodisponibilidad oral de ISA podría ser el resultado de la sobredosis y la poca estabilidad en los ácidos estomacales.

KEY WORDS: Bioavailability, Isopropylidene shikimic acid, Metabolism, Oral dosage, Pharmacokinetics.

* Author to whom correspondence should be addressed. *E-mail:* njtcm@263.net