

Pharmacokinetics of Isoniazid and Rifampicin-Loaded Bovine Serum Albumin Nanoparticles in Rabbits

Zhaohui GE ^{1,2} #, Rong MA ² #, Guangxian XU ², Jing BAI ², Siming LIANG ²,
Zhen CHEN ², Qian WANG ³, & Wei MA ¹ *

¹ The First Affiliated Hospital of Xi'an Jiaotong University, Xi'an 710061, China

² General Hospital of Ningxia Medical University, Yinchuan 750004, China

³ University of South Florida College of Pharmacy, Tampa, FL, USA

SUMMARY. The aim of this study was to evaluate the pharmacokinetics of isoniazid and rifampicin-loaded bovine serum albumin nanoparticles (INH-RFP-BSA-NPs) in rabbits after intravenous administration. Sixteen New Zealand rabbits were randomly divided into experimental group (administration of INH-RFP-BSA-NPs) and control group (administration of equal amounts of INH and RFP). INH and RFP blood plasma concentrations and pharmacokinetic parameters (C_{max} , $t_{1/2}$, AUC, MRT, CL, V_{ss}) were observed. Plasma INH and RFP concentrations remained relatively stable and could be detected in serum until 96 and 48 h after administration, respectively. Significant differences in pharmacokinetic parameters of RFP were observed between the INH-RFP-BSA-NP and control group ($P < 0.05$). Except C_{max} or CL of INH, significant differences in other indices existed between these two groups ($P < 0.05$). Intravenous administration of INH- and RFP-loaded nanoparticles can maintain effective blood plasma concentrations of INH and RFP for an extended period of time. INH-RFP-BSA-NPs have good and sustained *in vitro* release effects.

RESUMEN. El objetivo de este estudio fue evaluar la farmacocinética de nanopartículas de albúmina de suero bovino cargadas de isoniazida y rifampicina (INH-RFP-BSA-NP) en conejos después de la administración intravenosa. Dieciséis conejos de Nueva Zelanda se dividieron aleatoriamente en grupos experimentales (administración de INH-RFP-BSA-NP) y grupo de control (administración de cantidades iguales de INH y RFP). Se observaron las concentraciones plasmáticas de INH y RFP y los parámetros farmacocinéticos (C_{max} , $t_{1/2}$, AUC, MRT, CL, V_{ss}). Las concentraciones plasmáticas de INH y RFP se mantuvieron relativamente estables y pudieron detectarse en el suero hasta 96 y 48 h después de la administración, respectivamente. Se observaron diferencias significativas en los parámetros farmacocinéticos de RFP entre INH-RFP-BSA-NP y el grupo control ($P < 0.05$). Excepto C_{max} o CL de INH, existieron diferencias significativas en otros índices entre estos dos grupos ($P < 0.05$). La administración intravenosa de nanopartículas cargadas con INH y RFP puede mantener concentraciones efectivas de INH y RFP en plasma sanguíneo durante un período de tiempo prolongado. INH-RFP-BSA-NP tiene buenos y sostenidos efectos de liberación *in vitro*.

KEY WORDS: albumin, isoniazid, nanoparticles, pharmacokinetic processes, rifampicin.

These authors contributed equally to this work

* Author to whom correspondence should be addressed. E-mail: drmawei60@163.com