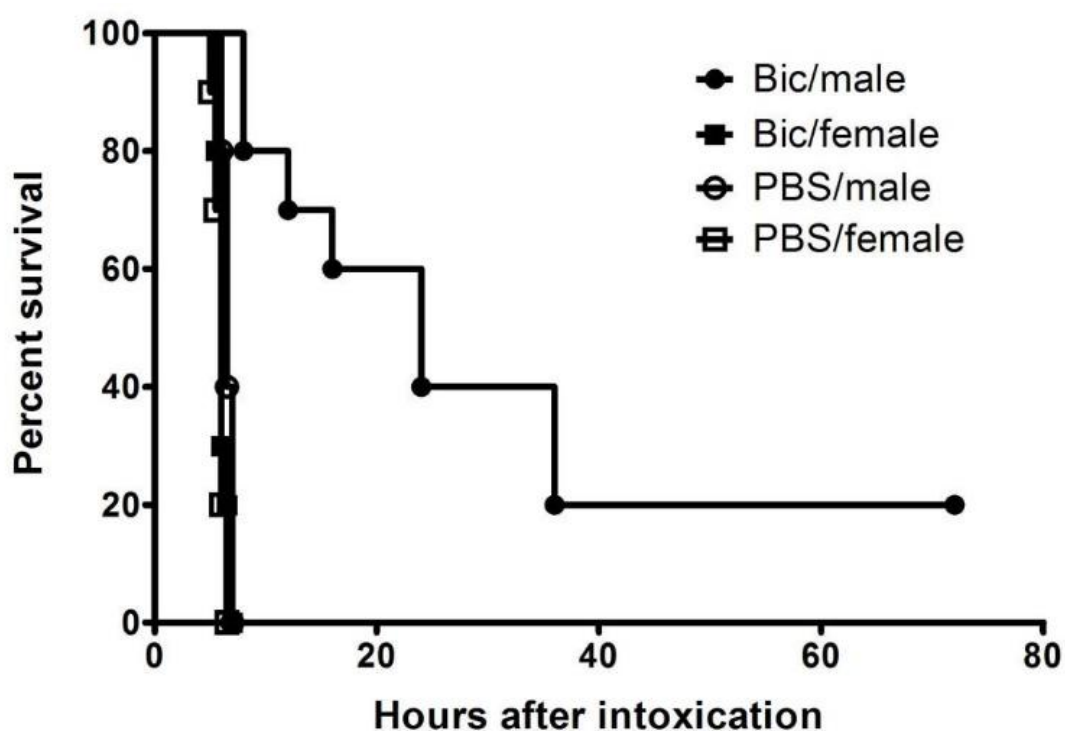


A Sexual Dimorphism Influences Bicyclol-Induced Hepatic Heat Shock Factor 1 Activation and Hepatoprotection

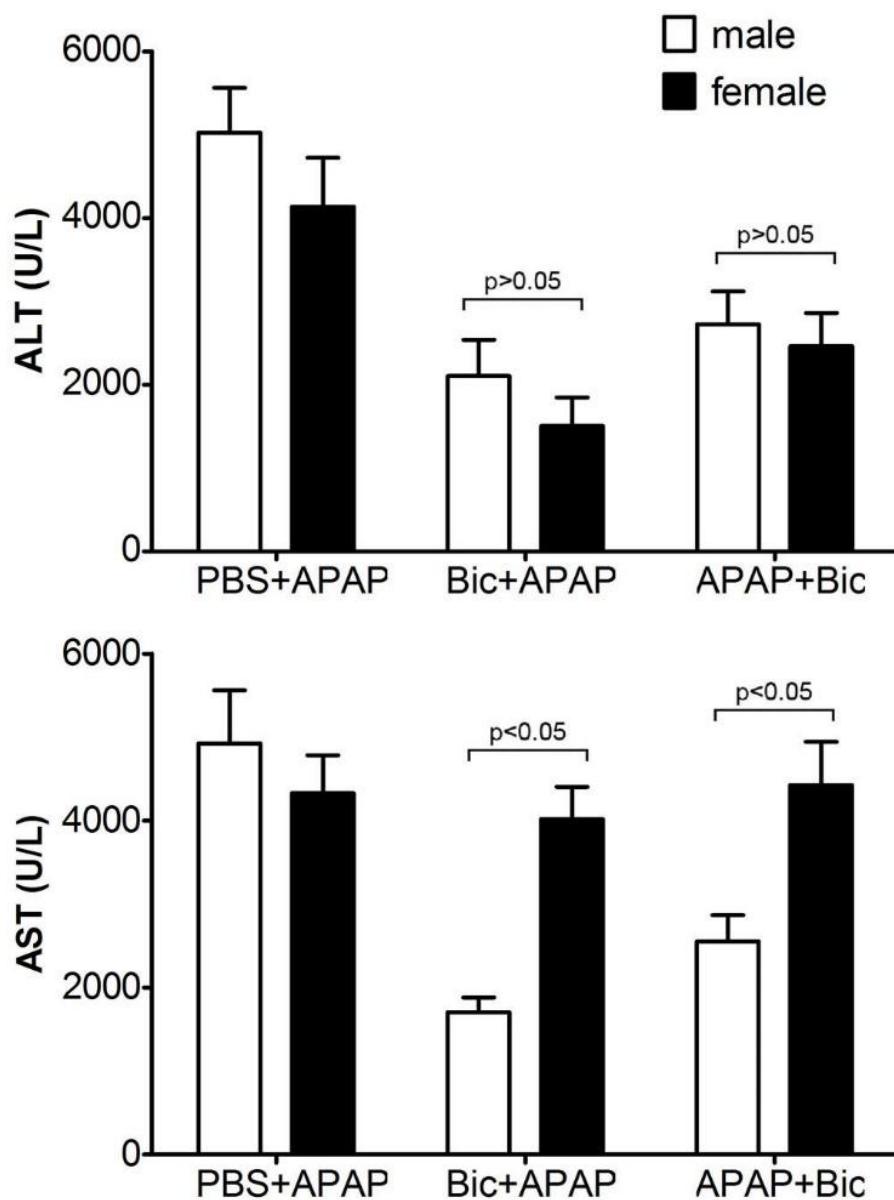
Xiaosong Chen, Jianjian Zhang, Conghui Han, Huijuan Dai, Xianming Kong,

Longmei Xu, Qiang Xia, Ming Zhang and Jianjun Zhang

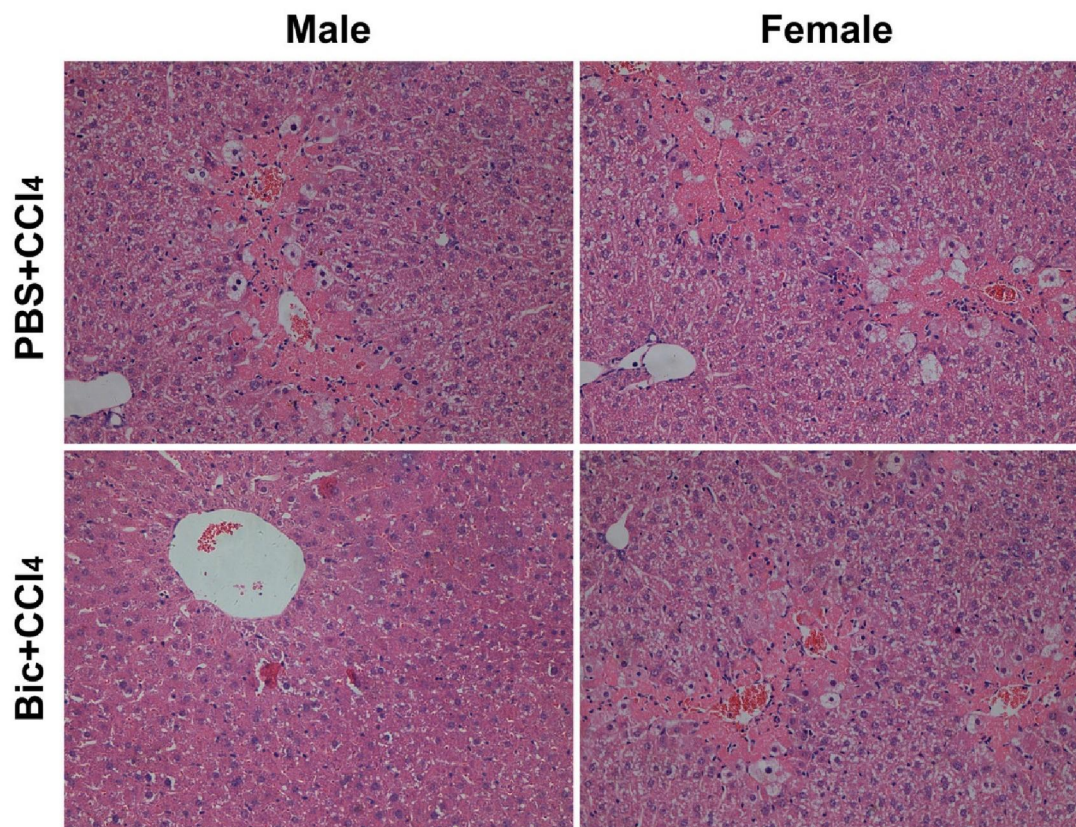
Molecular Pharmacology



Supplemental Figure 1. The sex difference remained when bicyclol was administered after GalN/LPS intoxication. A single dose of bicyclol (300 mg/kg) was administered at 1 h after GalN/LPS intoxication and survival of mice was observed thereafter (n=10 per group). The treatment led to a significant survival advantage only in male mice by Kaplan-Meier analysis (log-rank test, $P < 0.05$ between Bic/male and other groups).



Supplemental Figure 2. The sex difference in bicyclol-induced hepatoprotection against APAP toxicity. Male or female mice were orally administered with three doses of bicyclol (300 mg/kg) or PBS, followed by intraperitoneal injection of 300 mg/kg of APAP (Bic+APAP). Separate groups received one dose of bicyclol at 1 h after APAP treatment (APAP+Bic). Serum ALT and AST concentrations at 6 h after APAP injection were shown (n=4 per group). Statistical significance or insignificance were indicated.



Supplemental Figure 3. The sex difference in bicyclol-induced hepatoprotection against CCl₄ toxicity. Male or female mice were orally administered with three doses of bicyclol (300 mg/kg) or PBS, followed by intraperitoneal injection of carbon tetrachloride (0.4% diluted in olive oil) at a single dose of 10 ml/kg. Representative HE-stained sections from post-intoxication livers harvested at 48 h. (original magnification, $\times 200$). There were 4 mice in each group and different individuals in the same group showed consistent results.