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In Vitro Evidence of Tacrolimus-Zidovudine (AZT) Interaction

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SUMMARY. With the improvement on the number of solid organ transplantation in the HIV-infected patients, the potential risk between immunosuppressants and anti-HIV drugs will increase. The aim of the present study is to evaluate the tacrolimus-zidovudine (AZT) interaction using in vitro human liver microsomes (HLMs) incubation system. The results showed that tacrolimus exhibited concentration-dependent inhibition towards HLMs-catalyzed AZT glucuronidation. Both Dixon plot and Lineweaver-Burk plot showed the competitive inhibition of AZT glucuronidation by tacrolimus. The plot with the slopes from the Lineweaver-Burk plot versus the concentrations of tacrolimus was the most common plot method to calculate the inhibition kinetic parameter (K_i), and the present study used this plot method to calculate the Ki value to be 379 μ M. All these results indicated possible tacrolimus-AZT interaction, which will furtherly broaden the inhibition profiles between clinical drugs and AZT.

KEY WORDS: Tacrolimus, Zidovudine (AZT), Drug-drug interaction (DDI).

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