Interplay between Carboxylesterase 2 and Intestine Transporters Limits the Bioavailability of Allisartan, a prodrug of Exp3174 for Hypertension Treatment in Humans

Xiuli Li[1], Jingchao Sun[2], Zitao Guo[1], Dafang Zhong[1], and Xiaoyan Chen[1]

[1] *Shanghai Institute of Materia Medica, Chinese Academy of Sciences, Shanghai, China*

[2] *Shenzhen Salubris Pharmaceutical Co., Ltd, Guangdong, China*

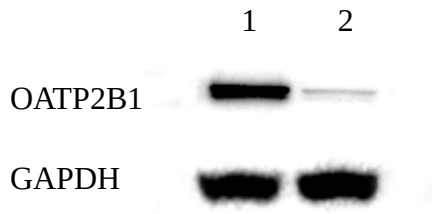
Drug Metabolism and Disposition

Figure legends

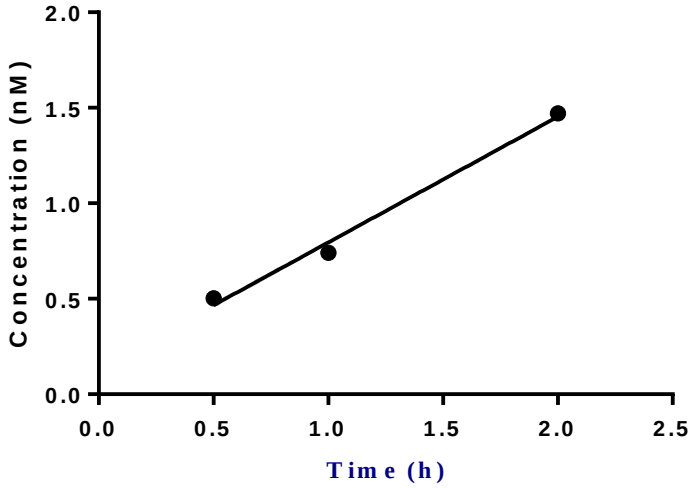
Supplemental Fig. 1. Western blot analysis of HEK293-OATP2B1 and HEK293-Mock cells for OATP2B1 detection. A total of 20 μg of cell proteins were subjected to SDS-PAGE and transferred to NC membrane for immunostaining. 1, HEK293-OATP2B1; 2, HEK293-Mock.

Supplemental Fig. 2. The time course of transport of Exp3174 in basolateral receiver side in Caco-2 monolayer cells. The concentration of Exp3174 was 50 μM.

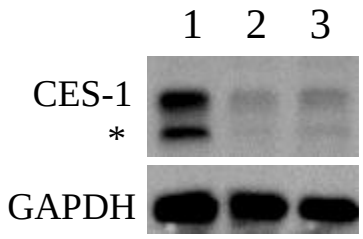
Supplemental Fig. 3. Western blot analysis of Caco-2, HEK293-OATP2B1, and MDCK-MDR1 cells for CES1 detection. A total of 8 μg of cell proteins were subjected to SDS-PAGE and transferred to PVDF membrane for immunostaining. 1, Caco-2; 2, HEK293-OATP2B1; 3, MDCK-MDR1. * monomeric form of CES1.



Supplemental Fig. 1



Supplemental Fig. 2



Supplemental Fig. 3