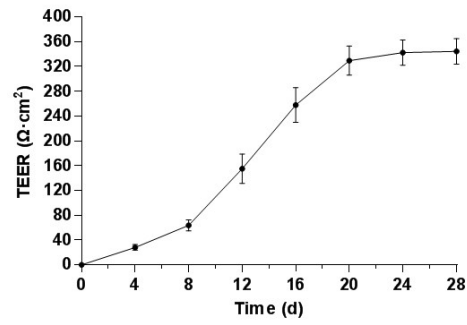


Supplement Table 1 The transepithelial electrical resistance (TEER) values ($\Omega \cdot \text{cm}^2$) before and after transport experiments (Mean \pm SD, n = 3)

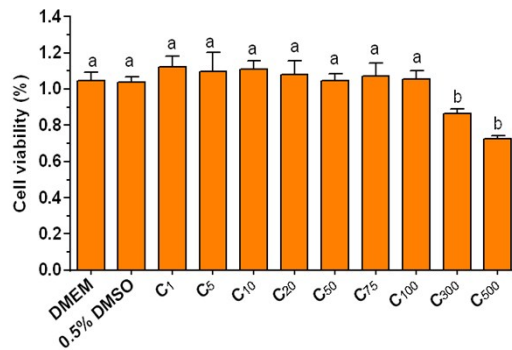
Group	AP \rightarrow BL		BL \rightarrow AP	
	Before	After	Before	After
7,8-DHF (pH=7.4)	369.8 \pm 12.3	354.3 \pm 13.6	345.6 \pm 16.3	334.5 \pm 15.3
7,8-DHF (pH=6.0)	354.3 \pm 12.6	350.2 \pm 16.3	342.3 \pm 34.1	339.6 \pm 23.6
7,8-DHF (pH=6.7)	350.3 \pm 16.1	342.6 \pm 11.2	326.1 \pm 14.9	308.6 \pm 11.2
7,8-DHF (37 °C)	364.2 \pm 5.60	354.3 \pm 18.9	332.1 \pm 11.8	326.3 \pm 10.3
7,8-DHF (4 °C)	326.9 \pm 15.8	325.3 \pm 16.8	336.2 \pm 15.2	335.4 \pm 14.9
7,8-DHF+ NaN ₃	385.6 \pm 28.9	316.6 \pm 25.6	356.3 \pm 15.6	316.9 \pm 21.3
7,8-DHF+ NaN ₃ +2-DOG	402.3 \pm 14.3	326.3 \pm 18.9	356.9 \pm 14.7	316.4 \pm 20.1
7,8-DHF+ Verapamil	336.5 \pm 12.8	303.2 \pm 11.9	345.3 \pm 21.6	311.8 \pm 14.9
7,8-DHF+ MK 571	335.9 \pm 23.5	321.8 \pm 21.4	347.2 \pm 17.9	340.7 \pm 14.6
7,8-DHF+ Probenecid	315.6 \pm 12.6	302.6 \pm 10.8	371.5 \pm 41.2	352.8 \pm 33.9
7,8-DHF+ Lesinurad	354.7 \pm 25.7	347.9 \pm 21.7	423.6 \pm 38.1	387.5 \pm 41.2
7,8-DHF+ ZnSO ₄	368.3 \pm 26.6	357.9 \pm 24.3	367.9 \pm 42.7	346.8 \pm 33.9
7,8-DHF+ Rifampicin	389.3 \pm 53.6	362.1 \pm 43.1	400.3 \pm 21.7	374.9 \pm 18.7
7,8-DHF+ Quinidine	313.5 \pm 11.4	304.5 \pm 10.3	326.8 \pm 11.6	314.2 \pm 17.8
7,8-DHF+ Levofloxacin	326.8 \pm 14.8	323.8 \pm 12.4	345.7 \pm 16.3	340.8 \pm 12.7
7,8-DHF+ Cimetidine	362.1 \pm 26.9	360.4 \pm 25.6	326.7 \pm 21.8	323.8 \pm 18.8
7,8-DHF+ Phloretin	351.9 \pm 42.3	342.6 \pm 33.9	328.7 \pm 14.9	324.8 \pm 15.7
7,8-DHF+Phloridzin	350.1 \pm 11.4	335.2 \pm 8.60	387.9 \pm 26.9	374.2 \pm 18.7

Supplement Table 2 Percentage recovery of 7,8-DHF at the different incubation time in HBSS and Caco-2 cells with or without inhibits (Mean \pm SD, n = 3)

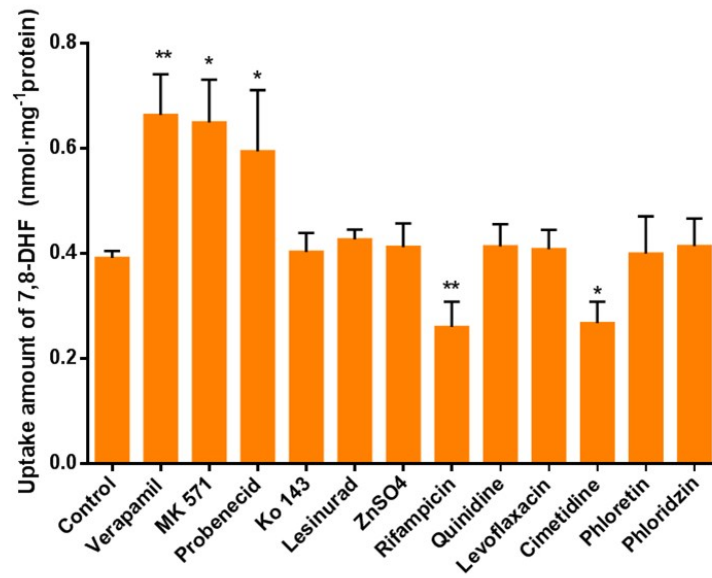
Group	Recovery of 7,8-DHF (%)			
	60 min	120 min	180 min	240 min
HBSS	99.51 \pm 2.35	98.43 \pm 2.16	97.94 \pm 1.47	96.56 \pm 1.74
HBSS (Caco-2)	98.92 \pm 1.63	97.83 \pm 1.54	97.41 \pm 3.26	96.21 \pm 3.31
HBSS (Caco-2)+Verapamil	99.21 \pm 1.25	98.41 \pm 0.39	97.76 \pm 1.44	96.44 \pm 2.01
HBSS (Caco-2)+MK 571	99.63 \pm 3.01	98.56 \pm 3.14	97.66 \pm 1.69	96.59 \pm 3.04
HBSS (Caco-2)+Probenecid	98.72 \pm 1.26	97.89 \pm 1.45	97.86 \pm 1.45	96.50 \pm 1.63
HBSS (Caco-2)+Ko 143	99.03 \pm 2.69	98.49 \pm 2.63	98.01 \pm 4.56	96.55 \pm 0.96
HBSS (Caco-2)+Lesinurad	99.40 \pm 3.05	99.03 \pm 3.16	98.52 \pm 4.01	97.01 \pm 3.63
HBSS (Caco-2)+ZnSO ₄	98.78 \pm 2.78	98.07 \pm 1.23	97.98 \pm 2.74	96.44 \pm 1.74
HBSS (Caco-2)+Rifampicin	99.45 \pm 1.69	99.23 \pm 1.41	98.41 \pm 1.74	96.89 \pm 2.59
HBSS (Caco-2)+Quinidine	98.46 \pm 1.74	98.03 \pm 3.61	97.55 \pm 2.86	96.32 \pm 0.29
HBSS (Caco-2)+Levofloxacin	99.32 \pm 2.34	98.33 \pm 0.34	97.82 \pm 1.48	96.44 \pm 2.41
HBSS (Caco-2)+Cimetidine	98.70 \pm 2.30	97.56 \pm 1.75	97.58 \pm 3.04	96.31 \pm 3.01
HBSS (Caco-2)+Phloretin	98.63 \pm 2.01	97.41 \pm 2.69	96.97 \pm 2.14	96.03 \pm 1.06
HBSS (Caco-2)+Phloridzin	99.26 \pm 2.67	98.56 \pm 3.06	97.86 \pm 2.31	97.03 \pm 3.33



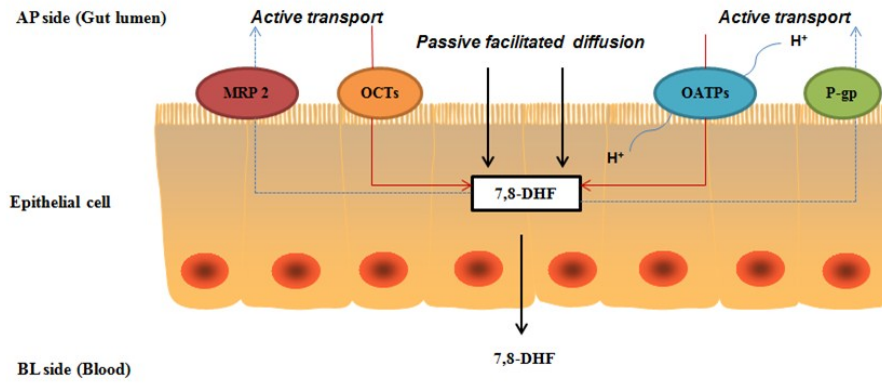
Supplement Fig.1 TEER value of the Caco-2 cell monolayers during culture. Data were shown as mean±SD (n=12 per group)



Supplement Fig.2 Cytotoxicity of 7,8-DHF on Caco-2 cells as determined using the CCK-8 assay. Data were shown as mean±SD (n=6 per group), graph bars with different letters on top correspond to statistically significant results ($p < 0.05$) based on one-way ANOVA analysis followed by Tukey's honest significant difference post hoc tests.



Supplement Fig.3 Uptake amount of 7,8-DHF in the Caco-2 cells at the end of transport time with various inhibitors (nmol · mg⁻¹ protein). **p* < 0.05, ***p* < 0.01, vs Control, the significant differences were analysed by unpaired two-tailed Student's *t* test.



Supplement Fig.4 Possible routes for 7,8-DHF transport across Caco-2 cell monolayers.