

Fig. S1. Effects of CSC on OCT2 and MATE2-K activity.

(A) HEK-OCT2 and HEK-MATE2-K cells were incubated with the reference substrate TEA for 5 min, in the absence (control) or presence of the reference inhibitors amitriptyline (for OCT2) or verapamil (for MATE2-K) or of CSC used at 80 and 320 µg/mL. After washing, intracellular accumulation of substrate was determined by scintillation counting. Data are expressed as percentages of substrate accumulation found in untreated control cells, arbitrarily set at 100 %. They are the means \pm SEM of at least three independent determinations. *, $p < 0.05$ when compared to control cells. (B) HEK-OCT2 and HEK-MATE2-K cells were incubated with TEA for 5 min in the absence or presence of various concentrations of CSC (from 1 to 640 µg/mL) or of reference inhibitors. OCT2 and MATE2-K activities were next calculated as described in Materials and Methods and are expressed as percentages of those found in control cells not exposed to CSC, arbitrarily set at 100 %. Data are the means \pm SEM of at least three independent assays. IC₅₀ values are indicated on the top of each graph.

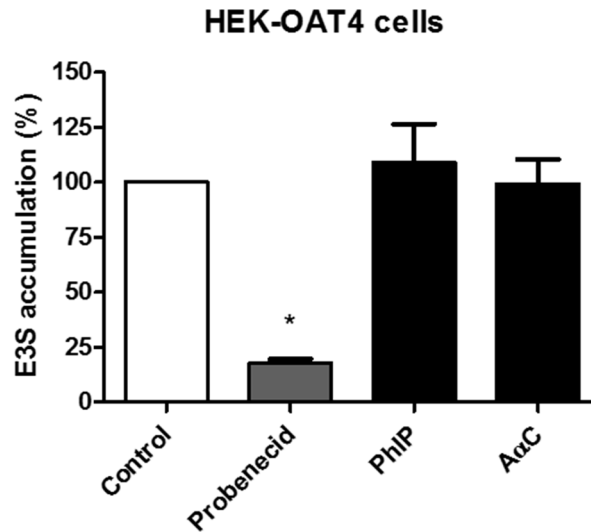


Fig. S2. Effects of the heterocyclic amines PhIP and AαC on OAT4 activity.

HEK-OAT4 cells were incubated with the reference OAT4 substrate E3S for 5 min, in the absence (control) or presence of the reference inhibitor probenecid or of 100 μ M PhIP or 100 μ M A α C. After washing, intracellular accumulation of substrate was determined by scintillation counting. Data are expressed as percentages of substrate accumulation found in untreated control cells, arbitrarily set at 100 %. They are the means \pm SEM of at least three independent determinations. *, $p < 0.05$ when compared to control cells.