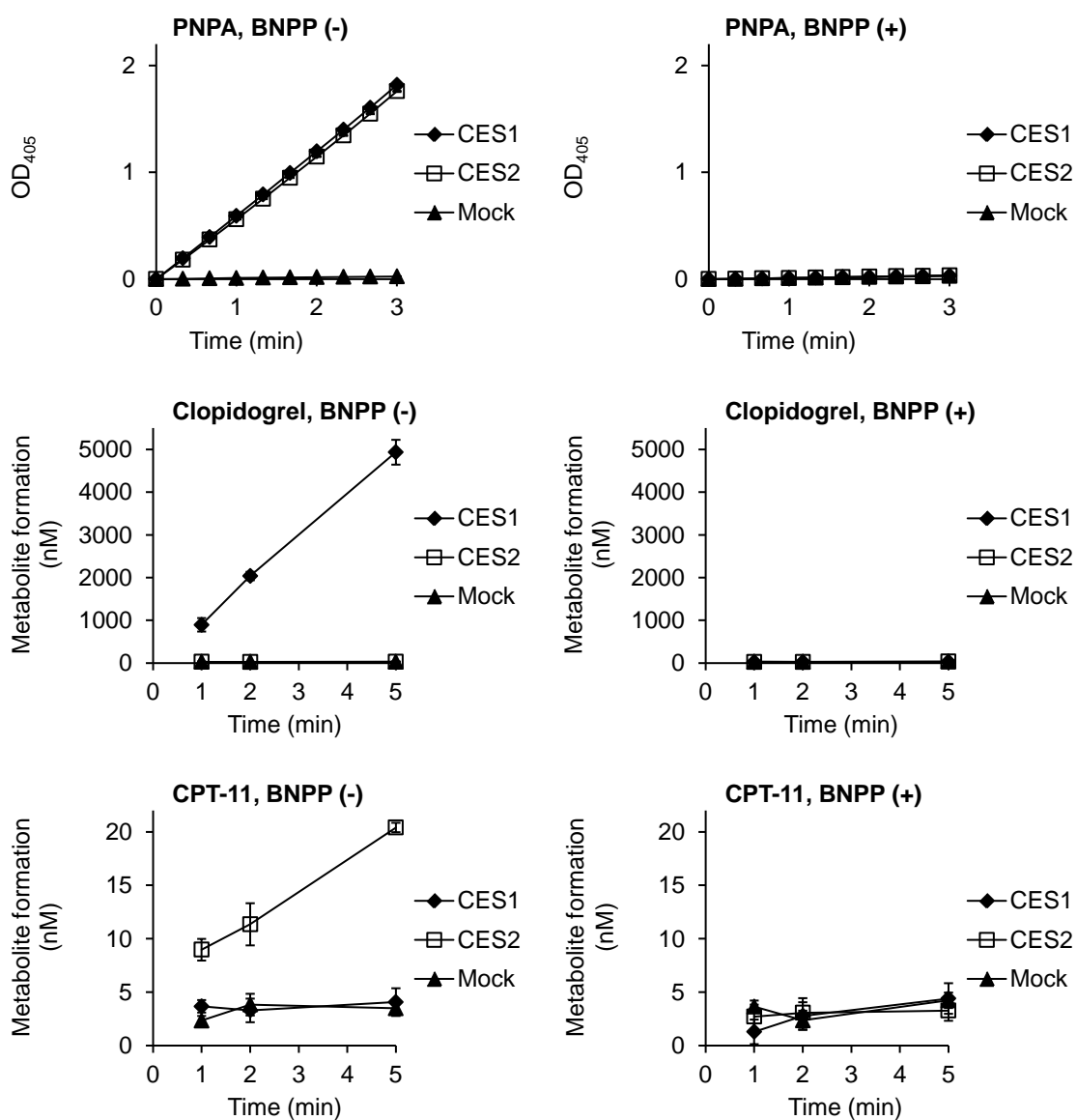


Different hydrolases involved in bioactivation of prodrug-type angiotensin receptor blockers: carboxymethylenebutenolidase and carboxylesterase 1

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Drug Metabolism and Disposition

Supplemental Figure S1



Supplemental Figure S1 Hydrolysis of *p*-nitrophenyl acetate (PNPA), clopidogrel, and CPT-11 by recombinant human CES1 and CES2.

PNPA (1 mM), clopidogrel, and CPT-11 (10 μ M each) were incubated in 50 mM Tris-HCl buffer (pH 7.5) with 5% (v/v) conditioned media from human CES1- or CES2-overexpressed cell culture as an enzyme source under the conditions with or without 1 mM bis-*p*-nitrophenylphosphate (BNPP). PNPA-hydrolase activity at 30 °C was determined by monitoring the absorbance at 405 nm. Clopidogrel- and CPT-11-hydrolase activities at 37 °C were determined by measuring formed metabolites, clopidogrel carboxylic acid and SN-38, respectively, by LC-MS/MS. Data represent the means \pm SD of triplicate determinations.