

Supplemental table S1: Atorvastatin metabolite concentrations in the time-series experiment on primary human hepatocytes of Individual 1

	AS	ASL	ASpOH	ASoOH	ASLpOH	ASLoOH				
time [min]	Extracellular Concentrations [pmol ml ⁻¹]									Recovery [%]
0	8797.1	30.5	n.d.	n.d.	n.d.	n.d.				
10	8258.3 ±405.1	28.5 ±4.9	n.d.	3.3 ±0.2	n.d.	n.d.				97.6
30	8086.6 ±518.3	31.3 ±5.3	11.0 ±0.5	41.0 ±0.9	n.d.	n.d.				96.8
60	7648.4 ±1473.8	41.4 ±2.6	29.8 ±3.3	167.4 ±17.5	5.2 ±0.1	9.0 ±0.2				94.8
120	6132.4 ±171.2	54.1 ±1.8	241.6 ±7.5	899.9 ±19.1	10.1 ±0.5	21.5 ±0.4				88.8
180	5592.2 ±254.3	61.7 ±0.6	337.1 ±19.7	1205.0 ±37.8	11.4 ±0.8	24.1 ±1.9				86.5
240	5190.3 ±189.3	62.0 ±4.6	516.2 ±11.9	1675.5 ±63.5	13.7 ±0.6	25.5 ±0.9				89.0
300	4404.2 ±230.1	55.9 ±4.4	723.4 ±30.2	2201.0 ±66.3	16.2 ±1.0	29.3 ±0.4				89.0
360	3285.1 ±352.2	51.3 ±7.3	868.4 ±61.0	2523.1 ±76.3	15.4 ±0.9	36.3 ±2.4				80.8
480	n.o.	n.o.	n.o.	n.o.	n.o.	n.o.				
600	2073.0 ±184.9	42.0 ±4.9	1245.7 ±38.2	3146.5 ±91.1	21.2 ±1.4	45.8 ±6.2				78.2
time [min]	Intracellular Concentrations [pmol ml ⁻¹]									
10	39727.4 ±2185.2	418.6 ±9.7	379.7 ±18.4	516.1 ±78.1	194.1 ±11.1	n.d.	n.d.			
30	41668.0 ±2317.2	566.8 ±72.0	2740.9 ±167.4	3029.1 ±186.8	237.9 ±18.7	n.d.	n.d.			
60	49772.4 ±1749.1	709.0 ±23.5	4737.3 ±233.1	6526.6 ±657.6	356.6 ±38.4	213.7 ±49.7				
120	44509.8 ±2992.8	766.6 ±25.4	8344.8 ±177.9	9949.9 ±834.8	527.6 ±34.1	340.5 ±39.4				
180	34790.4 ±3072.2	780.7 ±83.4	7872.6 ±964.4	8711.4 ±1062.5	502.5 ±35.1	339.5 ±6.2				
240	29716.1 ±3144.4	757.2 ±34.4	8576.3 ±1190.3	8411.1 ±861.5	531.8 ±34.1	385.8 ±35.4				
300	35702.4 ±1663.2	654.0 ±71.6	9728.0 ±931.4	8887.3 ±809.0	588.1 ±47.4	280.8 ±115.7				
360	27020.0 ±909.3	625.0 ±142.6	9767.9 ±439.4	8665.5 ±394.5	529.4 ±53.4	286.6 ±26.3				
480	25491.5 ±1835.9	622.7 ±13.9	13298.5 ±309.4	11235.2 ±787.7	596.5 ±25.7	442.7 ±53.8				
600	18839.0 ±1056.5	520.2 ±60.1	13176.6 ±1601.2	9775.0 ±987.2	539.7 ±38.1	410.3 ±54.1				

Extracellular concentrations (upper part) and intracellular concentrations (lower part) of Atorvastatin metabolites, Atorvastatin acid and lactone (AS and ASL) and corresponding para- and ortho-hydroxy-metabolites (acids: ASpOH and ASoOH; lactones: ASLpOH and ASLoOH) at the defined time-points with mean and standard deviation (n=3) from measurements per LC-MS/MS (n.d.: not determinable; n.o.: not observed). The recovery calculated from material balance equations is defined as the sum of intracellular and extracellular metabolite amounts divided by initial AS amount.