

Title: Modeling Human Nonalcoholic Steatohepatitis-Associated Changes in Drug Transporter Expression Using Experimental Rodent Models

Authors: Mark J. Canet, Rhiannon N. Hardwick, April D. Lake, Anika L. Dzierlenga, John D. Clarke, Nathan J. Cherrington

Journal Title: Drug Metabolism and Disposition

Supplemental Table 1

Gene	Human G Δ	Model	Rat G Δ
BCRP	0.9668	Athero	1.3248
		fa/fa	0.9749
		MCD	4.3729
MDR1A	0.8501	Athero	-1.0674
		fa/fa	0.3448
		MCD	2.3089
MDR1B		Athero	3.8287
		fa/fa	2.3772
		MCD	4.0775
MRP1	1.9526	Athero	1.4940
		fa/fa	-0.0244
		MCD	2.1080
MRP2	0.1726	Athero	1.8500
		fa/fa	-2.5933
		MCD	2.3534
MRP3	0.9156	Athero	3.2695
		fa/fa	0.6078
		MCD	4.2171
MRP4	2.3751	Athero	1.5168
		fa/fa	-0.6889
		MCD	2.8226
OATP1B1	0.0118	Athero	-0.0744
		fa/fa	-0.9347
		MCD	-2.7656
OATP1B3	-0.6837	Athero	-0.0744
		fa/fa	-0.9347
		MCD	-2.7656

Supplemental Table 1. Glass's Δ in mRNA Expression of Human NASH and Rat NASH Models. . Effect size of human transporter mRNA expression compared to effect sizes tabulated for mRNA expression of rat NASH models. Positive effect changes reflect induction of mRNA expression whereas negative effect changes reflect repression in mRNA expression. The magnitude of the values calculated represents the power of that particular model in detecting a difference in gene expression over control.

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Supplemental Table 2

Gene	Human G Δ	Model	Mice G Δ
BCRP	0.9668	Athero	0.6643
		db/db	1.1539
		MCD	2.0073
		ob/ob	0.8274
MDR1A	0.8501	Athero	1.1007
		db/db	14.3532
		MCD	5.6094
		ob/ob	7.4741
MDR1B		Athero	0.0624
		db/db	-0.5437
		MCD	0.2012
		ob/ob	-0.0176
MRP1	1.9526	Athero	-0.4296
		db/db	1.7775
		MCD	1.5931
		ob/ob	0.9164
MRP2	0.1726	Athero	5.4235
		db/db	3.8863
		MCD	4.6428
		ob/ob	4.4081
MRP3	0.9156	Athero	1.0547
		db/db	3.7532
		MCD	1.0913
		ob/ob	4.0013
MRP4	2.3751	Athero	1.1133
		db/db	4.4777
		MCD	5.8187
		ob/ob	7.4106
OATP1B1	0.0118	Athero	0.0525
		db/db	-4.1034
		MCD	-1.1830
		ob/ob	-1.7444
OATP1B3	-0.6837	Athero	0.0525
		db/db	-4.1034
		MCD	-1.1830
		ob/ob	-1.7444

Supplemental Table 2. Glass's Δ in mRNA Expression of Human NASH and Mouse NASH Models. . Effect size of human transporter mRNA expression compared to effect sizes tabulated for mRNA expression of mouse NASH models. Positive effect changes reflect induction of mRNA expression whereas negative effect changes reflect repression in mRNA expression. The magnitude of the values calculated represents the power of that particular model in detecting a difference in gene expression over control.

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Supplemental Table 3

Gene	Human G Δ	Model	Rat G Δ
BCRP	1.2878	Athero	-0.2987
		fa/fa	-1.2807
		MCD	0.7470
MRP2	0.9153	Athero	0.6560
		fa/fa	-0.0205
		MCD	3.3401
MRP3	1.0801	Athero	-0.8147
		fa/fa	0.7428
		MCD	4.3816
MRP4	1.1151	Athero	0.7763
		fa/fa	1.9884
		MCD	3.2338
OATP1B1	1.552	Athero	-1.9403
		fa/fa	2.0365
		MCD	-3.2784
OATP1B3	-1.861	Athero	-1.9403
		fa/fa	2.0365
		MCD	-3.2784
PGP	1.2755	Athero	1.7635
		fa/fa	-0.1348
		MCD	2.2404

Supplemental Table 3. Glass's Δ in Protein Expression of Human NASH and Rat NASH Models. Effect size of human transporter protein expression compared to effect sizes tabulated for protein expression of rat NASH models. Positive effect changes reflect induction of protein expression whereas negative effect changes reflect repression in protein expression. The magnitude of the values calculated represents the power of that particular model in detecting a difference in protein expression over control

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Supplemental Table 4

Gene	Human G Δ	Model	Mice G Δ
BCRP	1.2878	Athero	1.1488
		db/db	-0.4542
		MCD	-2.2921
		ob/ob	0.7902
MRP2	0.9153		
MRP3	1.0801	Athero	-0.1936
		db/db	1.7243
		MCD	0.3446
		ob/ob	1.6854
MRP4	1.1151	Athero	-0.6641
		db/db	1.8107
		MCD	2.0786
		ob/ob	1.7429
OATP1B1	1.552	Athero	0.4204
		db/db	-4.9064
		MCD	-5.0646
		ob/ob	-3.5524
OATP1B3	-1.861	Athero	0.4204
		db/db	-4.9064
		MCD	-5.0646
		ob/ob	-3.5524
PGP	1.2755	Athero	0.9611
		db/db	1.8784
		MCD	0.0050
		ob/ob	2.1669

Supplemental Table 4. Glass's Δ in Protein Expression of Human NASH and Mouse NASH Models. . Effect size of human transporter protein expression compared to effect sizes tabulated for protein expression of mouse NASH models. Positive effect changes reflect induction of protein expression whereas negative effect changes reflect repression in protein expression. The magnitude of the values calculated represents the power of that particular model in detecting a difference in protein expression over control.