

## SUPPLEMENTAL INFORMATION

### Direct Automated MALDI Mass Spectrometry Analysis of Cellular Transporter Function: Inhibition of OATP2B1 Uptake by 294 Drugs

Melissa S. Unger<sup>1,2,3</sup>, Lena Schumacher<sup>1</sup>, Thomas Enzlein<sup>1</sup>, David Weigt<sup>1</sup>, Maciej J. Zamek-Gliszczynski<sup>4</sup>, Matthias Schwab<sup>5,6,7</sup>, Anne T. Nies<sup>5,6</sup>, Gerard Drewes<sup>3</sup>, Sandra Schulz<sup>1</sup>, Friedrich B.M. Reinhard<sup>3</sup>, Carsten Hopf<sup>1,2,\*</sup>

1. Center for Mass Spectrometry and Optical Spectroscopy (CeMOS), Mannheim University of Applied Sciences, Paul-Wittsack Str. 10, 68163 Mannheim, Germany
2. Institute of Medical Technology, Heidelberg University and Mannheim University of Applied Sciences, Theodor-Kutzer-Ufer 1-3, 68167 Mannheim, Germany
3. Cellzome – a GlaxoSmithKline company, Meyerhofstr. 1, 69177 Heidelberg, Germany
4. Drug Metabolism and Pharmacokinetics, GlaxoSmithKline, 1250 S Collegeville Rd, Collegeville, Pennsylvania 19426, United States
5. Dr. Margarete Fischer-Bosch Institute for Clinical Pharmacology, Auerbachstr. 112, 70376 Stuttgart, Germany
6. iFit Cluster of Excellence (EXC2180) “Image Guided and Functionally Instructed Tumor Therapies”, University of Tübingen, Tübingen, Germany
7. Departments of Clinical Pharmacology, Pharmacy and Biochemistry, University of Tübingen, Geschwister Scholl Platz, 72074 Tübingen, Germany

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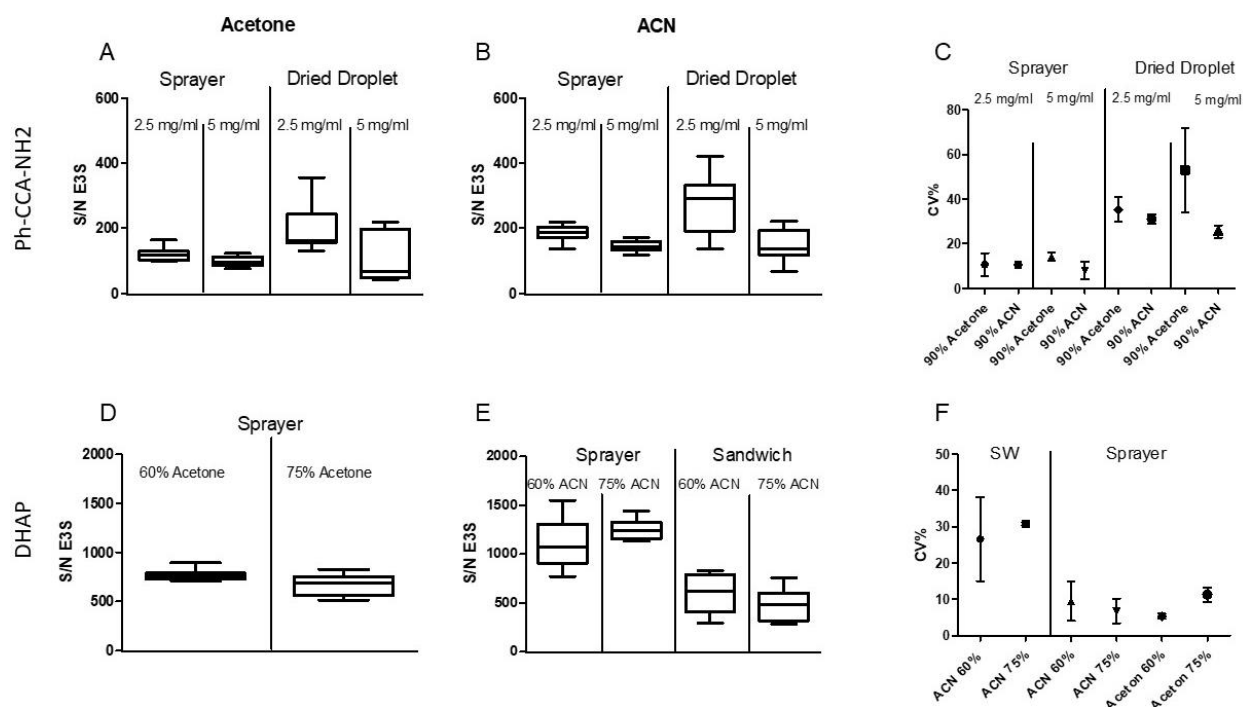
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## SUPPLEMENTARY METHODS

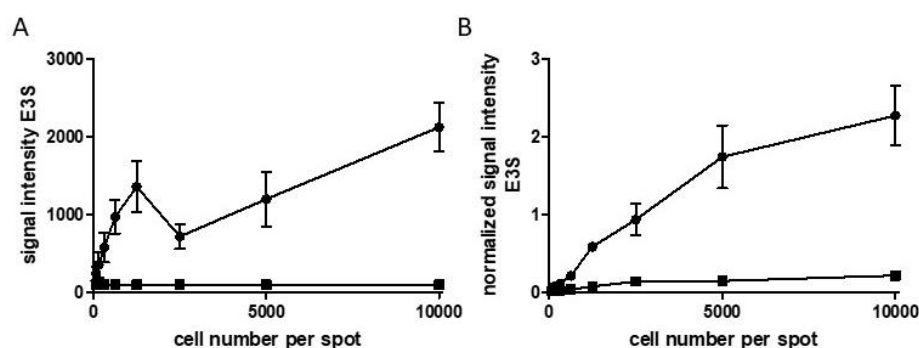
**CHEMICALS:** Dimethylsulfoxide (DMSO), ethanol, diammonium hydrogen citrate (DAHC), geneticin, poly-L-lysine and estrone-3-sulfate were obtained from Merck (Darmstadt, Germany). Milli-Q water (ddH<sub>2</sub>O; Millipore, Darmstadt, Germany) was prepared in-house. Acetone and acetonitrile (ACN) from VWR (Darmstadt, Germany). Phosphate buffered saline (PBS) was purchased from Biowest (Nuaille, France), trypsin from Corning (Kaiserslautern, Germany), Dulbecco's modified eagle medium (DMEM) from GE Healthcare (Freiburg, Germany) and fetal calf serum (FCS) from Thermo Fisher (Dreieich, Germany). CellTiter-Glo cell viability assay kit was obtained from Promega (Mannheim, Germany), methanol from Fisher Scientific (Schwerte, Germany), 2,5-dihydroxybenzoic acid (DHB) and sDHB (a 9:1 mixture of DHB and 2-hydroxy-5-methoxybenzoic acid) from Bruker Daltonics (Bremen, Germany), 2,5-dihydroxyacetophenone (DHAP) and sodium butyrate from Alfa Aesar (Haverhill, USA). Compounds were obtained from Tocris (Bristol, UK), Selleckchem (Houston, TX), Cayman (Ann Arbor, MI) or provided by GSK compound management (Stevenage, UK). Purity of all commercially sourced inhibitors was  $\geq 99\%$ ; all GSK-internally sourced compounds had  $\geq 95\%$  purity, except raloxifene (94%), tipranavir (91%), cetirizine (90%), diacerein (88%), and P-3004 unergol (87%).

**AUTOFLUORESCENCE AND QUENCHING MEASUREMENT:** For exclusion of possible false negatives and false positives due to autofluorescence and quenching effects during the fluorescence-based assay, the 294 drug set was analyzed and compared with the fluorescence of DBF itself. For testing the autofluorescence, the compounds were dissolved in HBSS to a final concentration of 10  $\mu\text{M}$ . DBF in a concentration of 10  $\mu\text{M}$  served as a comparison. The fluorescence intensities were compared at the excitation wavelength of 485 nm and emission wavelength of 535 nm. For evaluating if there is a quenching effect, DBF was given to each of the compound wells and it was checked if there was a decrease in the fluorescence compared to DBF alone

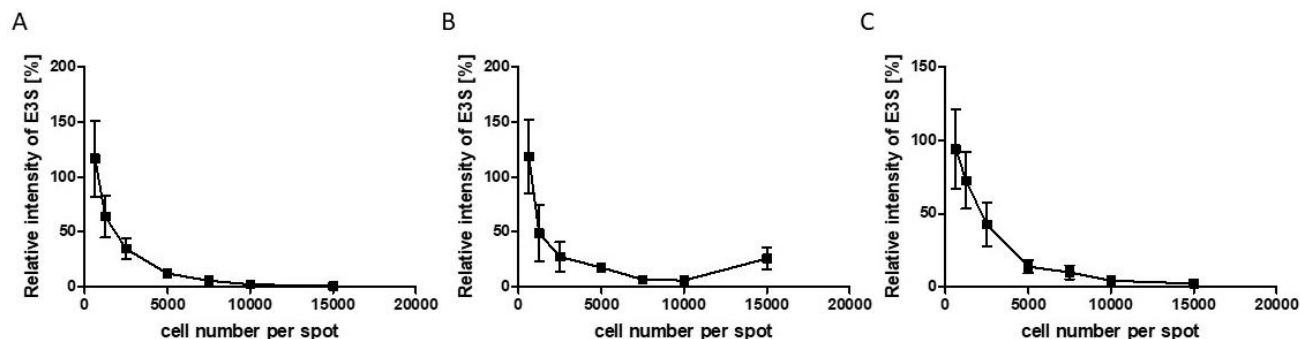
# SUPPLEMENTARY DATA:



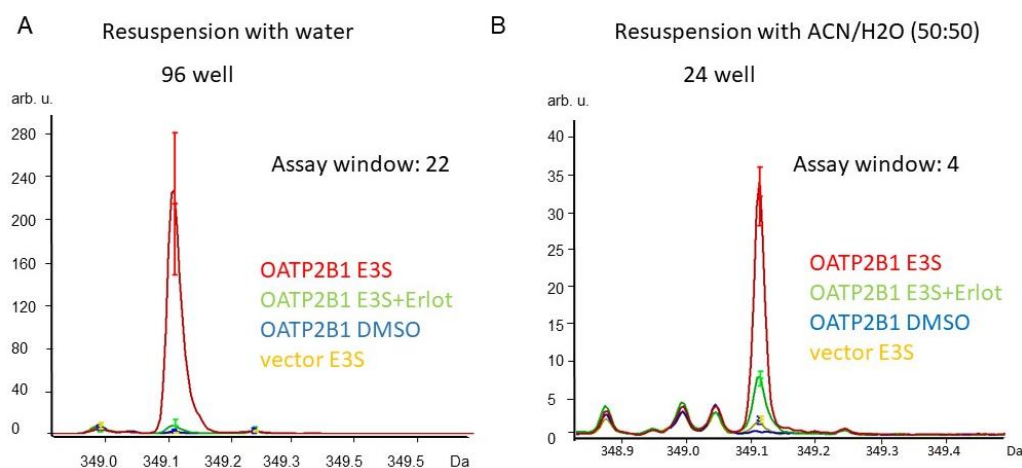
**Figure S1:** Method optimization with Ph-CCA-NH<sub>2</sub> and DHAP including CV% and S/N analysis shows possibility of both matrices: Cell pellets of HEK293 were resuspended at 2500 cells per  $\mu$ L in acetonitrile/ddH<sub>2</sub>O. One microliter of the suspension was applied to a MALDI target plate, which was spray coated with Ph-CCA-NH<sub>2</sub>. For the Dried Droplet application, cell pellets of HEK293 were resuspended at 5000 cells per  $\mu$ L in acetonitrile/ddH<sub>2</sub>O. The cells were mixed 1:1 with matrix before the manual application onto the target plate. E3S was spiked to the cell pellet in the final concentration 1  $\mu$ M. For a control, there were just cells measured without spiked E3S. Samples were measured using a rapifleX MALDI Tissue typer. (A,B,D,E) S/N of the E3S peak is shown in a boxplot of 9 measurement replicates each with median, standard deviation and whiskers of 5-95 percentile. (C,F) CV% of E3S peak of two biological replicates with 9 measurement replicates each. SW, sandwich.



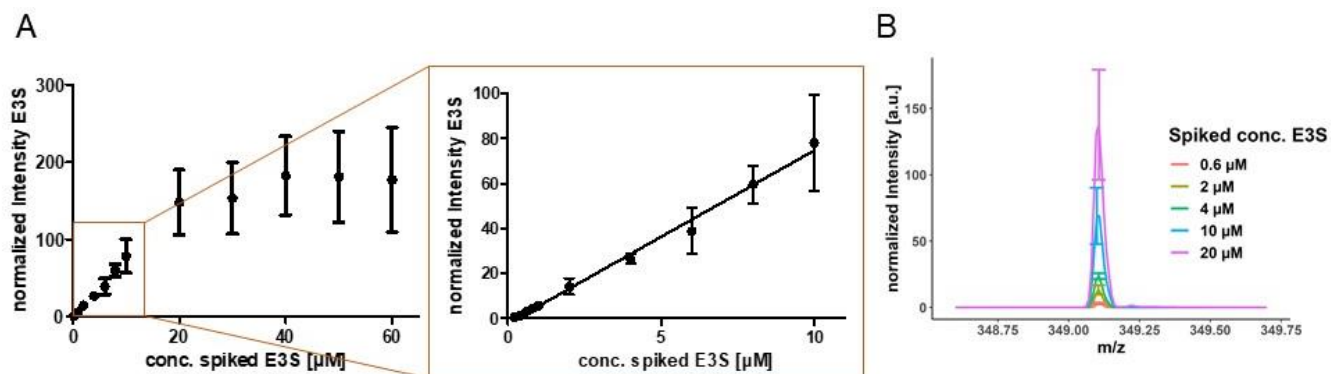
**Figure S2:** Evaluation of best cell number per spot: Comparison of cells treated with E3S (circles) versus vehicle DMSO (squares). Comparison of non-normalized data (A) with data normalized on D<sub>4</sub>-E3S (B). Mean  $\pm$  std. dev. of 9 measurement replicates.



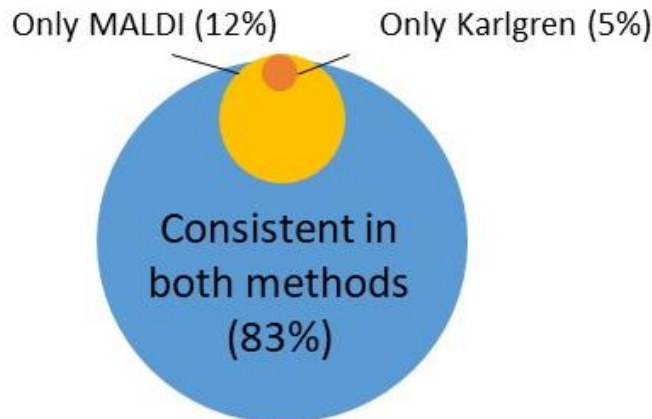
**Figure S3:** Evaluation of matrix suppression effect: Comparison of matrix suppression effect in three different cell passages. Analysis of differing cell numbers per spot. Cells were pelleted and resuspended in different amounts of suspension solution (ddH<sub>2</sub>O/matrix (50/50)). As matrix, 2.5 mg/mL Ph-CCA-NH<sub>2</sub> in 70% ACN was used. Final amount of 0.3  $\mu$ M D4-E3S and 1  $\mu$ M E3S were spiked to the matrix solvent. Cells were spotted manually and measured using a rapifleX MALDI. Shown are 18 measurement replicates with standard deviation. Intensities were normalized through setting 312 cells as 100 %.



**Figure S4:** Comparison of automated vs non-automated method shows increase in signal window due to automation: HEK293-OATP2B1 cells were treated with 20  $\mu$ M E3S for 30 minutes. In case of 24 well assay, the cell pellet was resuspended at 2500 cells per  $\mu$ l in acetonitrile/ddH<sub>2</sub>O. The cell pellets were resuspended at 5000 cells per  $\mu$ l in acetonitrile/ddH<sub>2</sub>O. Before the manual application onto the plate, the cells were mixed 1:1 with Ph-CCA-NH<sub>2</sub> (2.5 mg/ml in acetonitrile/ddH<sub>2</sub>O (90/10)) and thus applied Dried Droplet. In case of 96 well assay, the cells were resuspended at 10000 cells per  $\mu$ l in ddH<sub>2</sub>O and then mixed 1:1 with Ph-CCA-NH<sub>2</sub> in the above mentioned concentration. Dried Droplet application was done automatically with the CyBio FeliX pipetting platform. Samples were measured using a rapifleX MALDI. (A,B) Plots were done in GraphPadPrism, comparison of cells treated with E3S and DMSO; standard deviation of 9 measurement replicates.



**Figure S5:** Relationship between E3S intensity and concentration: **(A)** HEK293-vec cells were seeded out in a 96 well plate and snap-frozen in liquid nitrogen after harvest. E3S was spiked in different concentrations to the matrix. D4-E3S was spiked to the matrix at the usual concentration of 0.3  $\mu\text{M}$  final. 2.5 mg/mL Ph-CCA- $\text{NH}_2$  in 70 % ACN was used as matrix. D4-E3S-normalized intensities and standard deviation of 4 technical replicates with 4 measurement replicates each are shown. **(B)** Original E3S spectra (not normalized to D4-E3S).



**Figure S6:** Comparison of Karlgren publication (ref. 29) with MALDI data: Overlap of 83% of consistently identified inhibitors/non-inhibitors. Compounds that showed an inhibition of  $50\% \pm 5\%$  in both methods were considered as inhibitor.

**Table S1:** Autofluorescence and Quenching of whole dataset

	Quenching		Autofluorescence	
	Relative values		Relative values	
Compound	10 $\mu$ M	100 $\mu$ M	10 $\mu$ M	100 $\mu$ M
Acamprosate	40.6	53.7	1.4	1.8
Acemetacin	85.2	80.9	1.3	1.4
Acetaminophen	107.3	108.7	1.3	1.4
Acyclovir	118.3	114.4	1.6	1.6
Albuterol	105.3	103.0	1.4	1.3
Albuterol Sulfate	90.8	101.2	1.6	1.4
Alclofenac	111.9	110.6	1.4	1.6
Alendronate Sodium	125.6	116.6	1.4	1.6
Allopurinol	104.8	112.9	1.3	1.5
Alpha Tocopherol	110.6	105.8	1.4	1.5
Alpiropide	76.8	68.8	1.5	1.6
Amiodarone Hydrochloride	75.9	116.4	0.9	1.0
Amitriptyline	110.3	106.3	1.4	1.6
Amlodipine	70.5	69.1	1.4	1.2
Amlodipine	104.0	81.7	1.2	1.3
Amoxicillin	121.3	93.5	1.4	1.5
Ampiroxicam	107.1	107.4	1.4	1.4
Amsacrine	77.7	48.7	1.2	1.1
Anastrozole	112.7	111.8	1.4	1.7
Aripiprazole	113.6	112.3	1.0	1.0
Aspirin	111.2	110.1	1.4	1.6
Astemizole + Pseudoephedrine	86.7	94.1	1.5	1.5
Atenolol	112.7	109.6	1.4	1.6
Atomoxetine Hydrochloride	110.6	107.7	1.3	1.5
Atorvastatin	99.2	78.2	0.9	1.0
Azathioprine	114.0	111.2	1.4	1.6
Azithromycin	113.6	110.5	1.3	1.1
Azulene sodium sulfonate	107.6	100.3	0.8	1.1
Baclofen	116.9	114.1	1.4	1.6
Beclomethasone	86.3	81.4	1.3	1.0
Benazepril Hydrochloride	121.8	120.2	1.4	1.5
Benzbromarone	116.2	108.5	1.3	1.0
Benzonatate	118.2	107.4	1.3	1.4
Benzoyl peroxide	128.7	119.3	1.4	1.6
Bicalutamide	113.3	91.2	1.3	0.9
Bisoprolol Fumarate	109.7	108.8	1.4	1.6
Brimonidine Tartrate	110.5	105.5	1.4	1.7
Budesonide	108.7	95.2	1.4	1.4
Bumetanide	109.3	108.9	1.3	1.5
Bupropion	116.7	80.0	1.4	1.7
Buspirone Hydrochloride	91.4	88.5	1.4	1.5
Calcitriol	107.8	93.3	0.8	1.0
Carbamazepine	93.9	99.6	1.4	1.6
Carbidopa	118.0	112.1	1.6	1.7
Carvedilol	90.5	84.5	1.0	1.0
Cefdinir	85.7	93.1	1.3	1.3
Celecoxib	112.2	97.2	1.3	1.0
Cephalexin	106.9	115.2	1.3	1.2
Ceritinib	89.4	43.1	0.9	1.1
Cetirizine Hydrochloride	118.3	111.4	1.4	1.1
Chlorhexidine	107.9	59.2	1.3	1.2
Chlorthalidone	106.7	107.8	1.4	1.5
Cholecalciferol	114.9	115.5	1.0	1.1
Cicloxic acid	109.3	116.5	1.4	1.6
Ciprofloxacin	117.1	109.1	1.5	1.6
Citalopram	111.9	108.4	1.4	1.5
Clavulanate Potassium	121.5	116.4	1.3	1.6
Clindamycin hydrochloride	110.8	107.4	1.3	1.4
Clobetasol Propionate	109.0	100.7	1.0	1.0
Clonidine	83.3	80.6	1.5	1.8
Clopidogrel Bisulfate	93.8	91.7	1.4	1.4
Colchicine	109.2	99.6	1.3	1.4
Colforsin daropate	52.1	60.3	1.5	1.5
Conjugated Estrogens	118.8	116.3	1.4	1.7
Cromoglicic acid	111.6	116.1	1.4	1.5
Cyclobenzaprine Hydrochloride	100.3	110.2	1.4	1.6
Cyclosporine	84.8	80.9	0.8	0.8
Cysteamine	81.6	86.6	1.5	1.7
Desogestrel	65.8	92.6	1.3	1.4
Desonide	107.5	108.6	1.3	1.4
Desvenlafaxine	108.4	115.1	1.4	1.4
Diacerein	108.7	107.8	1.3	1.4
Diclofenac	106.8	111.1	1.3	1.5
Dicyclomine Hydrochloride	106.7	102.3	1.4	1.6
Diethylstilbestrol	92.8	89.4	1.4	1.7
Diflunisal	112.8	108.5	1.3	1.6
Diltiazem Hydrochloride	43.8	57.2	1.5	1.6
Diphenhydramine Hydrochloride	110.5	106.8	1.4	1.5
Dipyridamole	103.8	92.6	0.8	1.2
Dithranol	111.7	105.3	1.2	0.9
Divalproex Sodium	111.7	105.8	1.3	1.6
Donepezil Hydrochloride	50.3	58.1	1.5	1.6
Dorzolamide Hydrochloride	104.9	96.5	1.4	1.5
Doxazosin Mesylate	94.3	68.2	1.0	1.0
Doxepin Hydrochloride	83.7	82.9	1.5	1.6
Doxycycline	74.3	63.6	0.9	1.3
Doxycycline	127.0	76.1	1.0	1.2
Dronedarone	109.8	77.3	1.0	1.0
Drospirenone	107.8	95.2	1.2	0.9
Duloxetine	85.4	88.3	1.4	1.6
Efalith	110.1	116.3	1.4	1.6
Efamol	111.4	107.2	1.3	0.9
Efavirenz	90.5	78.3	1.3	1.3
Enalapril Maleate	109.3	109.6	1.3	1.5
Enprostil	63.9	91.2	1.6	1.8
Ergocalciferol	79.5	90.5	1.2	1.4
Erlotinib	109.1	75.8	1.4	0.9
Erythromycin	110.6	101.2	1.4	1.2
Escitalopram Oxalate	92.6	96.2	1.4	1.5
Estradiol	111.4	108.2	1.3	1.5
Ethanolamine oleate	106.0	119.4	1.4	1.7
Ethinyl Estradiol	84.8	91.7	1.5	1.6
Etodolac	105.5	104.0	1.4	1.6
Etretinate	108.2	110.4	1.3	1.4
Ezetimibe	106.1	90.8	0.8	0.8
Famotidine	121.6	113.9	1.5	1.5
Felbinac	109.2	111.1	1.4	1.5
Felodipine	91.4	84.0	1.1	1.0
Fenbufen	64.3	98.5	1.4	1.6
Fenofibrate	111.8	108.2	1.3	1.4
Finasteride	105.0	104.3	1.3	1.0
Fluconazole	113.0	111.7	1.3	1.5
Fluocinonide	114.4	101.6	1.3	1.1
Fluoxetine Hydrochloride	66.7	90.7	1.5	1.8
Flurbiprofen	110.5	106.3	1.4	1.6
Flutamide	95.6	92.2	1.3	1.5
Fluticasone	108.8	101.0	1.4	1.3
Fluticasone Propionate	110.2	89.9	1.1	1.3
Fluvastatin	116.3	94.2	1.0	1.3
Folic Acid	45.0	58.9	1.5	1.7
Foscarnet	122.4	120.9	1.5	1.7
Fosfosal	101.7	103.5	1.4	1.6
Furosemide	112.4	113.4	1.3	1.5
Gabapentin	101.1	90.6	1.3	1.6
Gemfibrozil	113.1	113.7	1.3	1.5
Genistein	116.2	106.7	1.5	1.5
Glimepiride	116.4	108.6	1.4	1.3
Glipizide	78.5	76.9	1.5	1.7
Gliquinone	105.4	86.4	0.9	0.9
Glyburide	111.9	101.3	1.3	0.8
Guacetasal	117.3	109.0	1.4	1.6
Guaifenesin	111.7	111.7	1.3	1.6
Guanfacine	117.4	110.9	1.4	1.6
Haloperidol	117.0	107.4	1.4	1.4
Hydralazine Hydrochloride	104.6	106.8	1.3	1.5
Hydrochlorothiazide	115.7	112.6	1.4	1.6
Hydrocortisone	111.2	110.9	1.6	1.8
Hydroxychloroquine Sulfate	112.5	100.6	1.4	1.5
Hydroxyzine	90.6	86.2	1.4	1.3
Ibuprofen	112.2	114.3	1.4	1.6
Indapamide	110.8	109.2	1.4	1.2
Indomethacin	117.0	110.9	1.3	1.5
Iornoxicam	94.2	88.7	1.4	1.6
Ioxoprofen	116.1	117.7	1.4	1.6
Ipriflavone	114.9	110.1	1.3	1.3
Irbesartan	119.7	111.1	1.4	1.3
Isoorbide Mononitrate	106.8	107.5	1.3	1.6
itraconazole	92.9	81.5	1.1	1.6
Ketoconazole	110.7	112.9	0.9	1.5
Ketoprofen	92.2	98.5	1.4	1.5

Compound	Quenching		Autofluorescence	
	Relative values		Relative values	
	10 $\mu$ M	100 $\mu$ M	10 $\mu$ M	100 $\mu$ M
Ketorolac Tromethamine	87.9	84.7	1.5	1.5
Labetalol	108.7	111.4	1.2	1.2
Lamotrigine	129.4	109.7	1.4	1.6
Lansoprazole	113.0	93.7	1.2	1.3
Latanoprost	91.9	87.8	1.4	1.6
Letrozole	87.8	89.8	1.5	1.6
Levetiracetam	86.4	92.6	1.4	1.7
Levocetirizine Dihydrochloride	115.3	114.5	0.9	1.4
Levodopa	108.2	115.5	1.4	1.6
Levofloxacin	91.2	91.8	1.5	1.6
Levosulpiride	113.4	106.1	1.4	1.4
Levothyroxine	110.2	78.6	1.2	1.3
Lidocaine	106.9	102.8	1.4	1.5
Lisinopril	91.1	94.4	1.5	1.6
Lithium	97.5	93.6	1.5	1.7
Loratadine	116.1	99.2	1.2	1.0
Losartan Potassium	104.7	106.8	1.3	0.9
Lovastatin	92.3	78.6	1.2	0.9
Meclizine Hydrochloride	105.8	114.7	1.3	1.1
Medroxyprogesterone Acetate	61.1	81.3	1.2	1.0
meglitrol	108.7	111.3	1.3	1.6
Meloxicam	97.7	93.9	1.5	1.6
Mesalamine	113.8	109.0	1.3	1.4
Mesna	111.9	118.4	1.4	1.6
Meso-2,3,-Dimercaptosuccinic acid	107.9	112.3	1.2	1.1
Metaxalone	106.8	112.5	1.4	1.6
Metformin Hydrochloride	119.2	110.3	1.5	1.6
Methimazole	107.8	104.9	1.4	1.6
Methocarbamol	85.3	90.1	1.4	1.7
Methotrexate	88.5	90.4	1.5	1.5
Methylprednisolone	102.8	95.7	1.4	1.3
Metoclopramide Hydrochloride	113.3	113.3	1.5	1.7
Metoprolol	106.2	113.3	1.4	1.6
Metronidazole	117.0	119.5	1.4	1.6
Minocycline Hydrochloride	97.4	72.3	0.9	1.4
Mirtazapine	104.5	111.4	1.3	1.4
Mofezolac	118.0	113.8	1.5	1.6
Mometasone Furoate	98.6	105.2	0.8	1.5
Mometasone Furoate	68.9	64.2	1.1	1.4
Montelukast	110.1	97.0	0.9	1.1
Moxifloxacin Hydrochloride	113.7	96.9	1.4	1.4
Mycophenolate Mofetil	100.0	91.0	1.5	1.6
Mycophenolic acid	117.7	115.5	1.4	1.6
Nabumetone	97.7	93.6	1.4	1.5
Nadolol	114.0	106.4	1.6	1.7
Naphazoline Hydrochloride	111.9	106.7	1.4	1.5
Naproxen	93.9	88.7	1.4	1.6
Nebivolol Hydrochloride	103.8	89.0	1.2	1.2
Neomycin	112.2	107.8	1.3	1.5
Niacin	112.1	112.1	1.4	1.5
Nifedipine	93.1	97.1	1.3	1.2
Nitrofurantoin	95.1	93.4	1.3	1.5
Norethindrone	115.4	119.6	1.4	1.1
Nortriptyline Hydrochloride	101.2	99.6	1.4	1.5
Novobiocin	103.8	82.7	0.9	1.1
Ofloxacin	96.5	97.4	1.4	1.3
Olanzapine	111.5	105.1	1.3	1.0
Olmesartan Medoxomil	114.6	105.2	1.3	1.0
Olopatadine Hydrochloride	88.8	88.7	1.4	1.4
Olsalazine	84.8	85.2	1.4	1.5
Omega-3 acid ethyl ester Omtryg	121.6	104.2	1.4	1.3
Omeprazole	115.4	107.2	1.4	1.3
Oseltamivir Phosphate	117.0	107.3	1.4	1.6
Oxaprozin	116.8	118.3	1.4	1.1
Oxcarbazepine	109.9	104.1	1.3	1.3
Oxybutynin	87.2	97.0	1.4	1.2
P-3004Unergol	100.3	82.9	0.9	1.0
Pantoprazole Sodium	93.5	103.7	1.4	1.3
Parecoxib	113.5	117.0	1.3	1.3
Paroxetine	134.0	107.0	1.4	1.6
Pemetrexed	111.5	116.5	1.4	1.5
Penicillin V	69.6	72.4	1.5	1.7
Pheniramine Maleate	112.4	102.4	1.4	1.6
Phenytoin	112.6	109.7	1.4	1.6
Pioglitazone Hydrochloride	117.6	123.8	1.3	1.2
pipotiazine	107.0	112.3	1.0	1.4
Piretanide	109.4	108.8	1.4	1.4
Piroxicam	109.8	110.2	1.2	1.5
Piroxicam	97.6	91.0	1.3	1.6
Potassium Citrate	107.6	114.3	1.4	1.6
Pramipexole Dihydrochloride	104.3	107.0	1.3	1.4
Pranoprofen	110.9	105.2	1.4	1.6
Prasugrel Hydrochloride	95.9	83.6	1.3	1.6
Pravastatin Sodium	69.7	88.0	1.3	1.4
Prednisolone	107.2	97.9	1.3	1.4
Prednisone	102.8	98.1	1.4	1.2
Probenecid	120.7	116.4	1.3	1.6
Progesterone	117.5	114.1	1.5	1.5
Promethazine Hydrochloride	112.0	110.9	1.4	1.5
Propafenone	116.2	111.7	1.4	1.5
Propagermanium	119.6	115.5	1.6	1.7
Propranolol Hydrochloride	107.3	102.6	1.4	1.5
Quetiapine Fumarate	114.4	101.0	1.3	0.9
Quinapril	117.3	106.8	1.3	1.5
Raloxifene Hydrochloride	110.5	86.8	1.2	1.2
Ramipril	83.8	105.7	1.3	1.5
Ranitidine	91.8	96.5	1.3	1.6
Reserpine	96.2	84.7	0.8	1.1
Risperidone	110.5	113.2	1.4	1.3
Rivaroxaban	112.9	104.3	1.1	1.2
Ropinirole Hydrochloride	119.6	104.9	1.3	1.6
Sertraline Hydrochloride	98.7	88.0	1.4	1.2
Sildenafil	95.9	89.4	1.3	1.1
Silymarin	98.5	97.0	1.2	1.1
Simvastatin	96.1	93.8	0.7	1.0
Sitagliptin Phosphate	109.1	108.2	1.3	1.1
Sivelestat	79.2	82.1	1.5	1.6
Sodium phenylbutyrate	109.2	107.0	1.4	1.6
Sodium salicate	117.7	115.9	1.6	1.7
Spironolactone	105.1	104.0	1.2	1.0
Sulfamethoxazole	110.2	109.3	1.4	1.6
Sulfasalazine	112.3	104.7	1.3	1.6
Sulindac	109.0	100.0	1.3	1.5
Sulpiride	109.6	98.1	1.4	1.5
Sulprostone	105.0	116.3	1.3	1.5
Sumatriptan	109.7	102.6	1.4	1.5
Tacrolimus	71.7	73.0	1.0	1.1
Tamoxifen Citrate	101.6	106.9	1.2	0.8
Tamsulosin Hydrochloride	92.0	87.9	1.3	1.3
Terazosin	106.0	83.7	1.2	1.1
Tetracycline	109.2	106.6	1.2	1.0
Tianeptine	114.7	111.2	1.2	1.1
Tiaprofen	101.2	107.0	1.4	1.6
Ticagrelor	106.7	108.1	1.7	1.5
Timolol	89.0	98.9	1.4	1.7
Tipranavir	94.1	97.7	0.9	1.0
Tizanidine	108.8	109.8	1.4	1.6
Tolterodine Tartrate	117.0	105.2	1.5	1.6
Topiramate	111.6	112.6	1.4	1.7
Torsemide	108.0	114.5	1.4	1.6
Travoprost	98.5	105.1	1.3	1.6
Trazodone Hydrochloride	86.6	101.5	1.3	1.3
Trepibutone	91.2	92.2	1.5	1.7
Triamcinolone	108.8	110.6	1.4	1.6
Triamterene	87.6	93.3	1.4	1.4
Triflusal	112.1	112.4	1.3	1.5
Trimethoprim	110.6	102.5	1.4	1.5
Tropine indometacinat	97.1	83.3	1.0	0.9
Valacyclovir	101.8	94.4	1.4	1.6
Valsartan	113.3	114.8	1.3	1.5
Venlafaxine Hydrochloride	103.1	95.3	1.4	1.6
Verapamil	111.5	110.3	1.3	1.0
Verapamil Hydrochloride	114.7	103.5	1.3	1.2
Vilazodone Hydrochloride	99.3	104.2	0.7	1.5
Warfarin	111.2	112.5	1.4	1.5
Xipamide	100.1	92.0	1.4	1.5
Zafirlukast	102.3	84.6	0.7	1.1