

NAME: DATE OF BIRTH: EMQN1

REPORT NUMBER: DATE COLLECTED:

DATE RECEIVED:

DATE OF REPORT:

7011pgx\_lonCode\_0341

GENDER: SPECIMEN TYPE:

ORDERING PHYSICIAN:

COMPREHENSIVE PGX REPORT

#### **Current Patient Medication**

This patient is either not receiving any medication or may be receiving medications that are outside the scope of this report.

### SNPs of Importance for Venous Thromboembolism Risk

Gene	Protein change	Nucleotide change	Marker	Genotype	Results
F5	Arg534Gln	1601G>A	rs6025	C/C	Normal risk
F2		*97G>A	rs1799963	G/G	Normal risk
VKORC1		1173C>T	rs9923231	C/C	Low warfarin sensitivity; high warfarin dosage
VKORC1		-1639G>A	rs9934438	G/G	Low warfarin sensitivity; high warfarin dosage
MTHFR	Ala222Val	665C>T	rs1801133	G/G	Normal MTHFR enzyme function.
MTHFR	Glu429Ala	1286A>C	rs1801131	T/T	Normal MTHFR enzyme function.



#### GENOTYPE/HAPLOTYPE/PHENOTYPE DETAIL

_		
Gene	Genotype-Haplotype	Phenotype
CYP1A2	*1B/*1L	Extensive metaboliser
CYP2B6	*9/*9	Extensive metaboliser
CYP2C8	*1/*1	Extensive metaboliser
CYP2C9	*1/*9	Extensive metaboliser
CYP2C19	*1/*2	Intermediate metaboliser
CYP2D6	*1/*1	Extensive metaboliser
CYP3A4	*1A/*1B	Extensive metaboliser
CYP3A5	*3A/*6	Poor metaboliser
VKORC1	H3/H4	Sensitive to Warfarin
SLCO1B1	*1B/*1B	Extensive function
ABCB1	*1/*1	Extensive function
ABCG2	*1/*1	Extensive function
TPMT	*1/*3C	Intermediate metaboliser
UGT1A1	*1/*1	Extensive metaboliser
UGT2B7	*1a/*1a	Extensive metaboliser
UGT2B15	*1/*2	Extensive metaboliser
DPYD	*9A/*9A	Extensive metaboliser
OPRM1	*1/*1	Sensitive to Opioids

<u>Disclaimer:</u> No patient should evaluate or use the information contained herein without the advice, consultation and supervision of a licensed healthcare professional such as a pharmacist or physician.

Methodology: Genomic DNA was extracted from the sample under analysis. A pharmacogenomics Ion Ampliseq panel was used to detect 136 SNV/Indel targets in 40 genes as well as CYP2D6 copy number variation. This panel covers the commonly known targets in genes encoding drug metabolism enzymes and associated transport proteins. The following genes are included: ABCB1, ABCG2, ADRA2A, ANK1, APOE, COMT, CYP1A2, CYP2B6, CYP2C19, CYP2C8, CYP2C9, CYP2D6, CYP3A4, CYP3A5, DBH, DPYD, DRD1, DRD4, F2, F5, FLOT1, GABRA6, GABRP, GRIK4, HCP5, HLA-A, HTR2A, HTR2C, ITGB3, KIF6, MTHFR, OPRD1, OPRK1, OPRM1, SLCO1B1, TPMT, UGT1A1, UGT2B15, UGT2B7, VKORC1. Sequencing was carried out using the Next Generation Sequencing platform Ion Gene Studio S5 Prime System (Thermo Fisher Scientific).

<u>Limitations</u>: This test detects specific mutations with pharmacogenomics evidence in the genes described without analysing the full coding sequence of these genes. Every molecular test has an internal 0,5-1% chance of error. This is attributable to rare molecular events and factors relating to the preparation and analysis of the samples.

#### **PHARMACOGENOMICS**

Genetic Markers Tested for Pharmacogenomics:

Results are arranged by drug response. Each individual report contains six sections, including: Patient's current medication (if any), Medication history, genotype/haplotype/phenotype detail, PGx report, Genomic Test Results, and Patient Information Card. Inclusion of the PGx Report indicates that the tested individual: displays decreased efficacy to the drug (light green dots), should use the drug as directed (green dots), or exhibits increased toxicity to the drug (red dots). Inclusion of Genomic Test Results indicates genotype, haplotype, phenotype, or presence of mutation.

Organisation of Table:

- 1. Gene/Locus refers to gene or intergenic region of genetic marker location.
- 2. Marker refers to the tested marker's unique identifier.
- 3. Genotype/Haplotype refers to the particular marker's combination of nucleotides. The letter(s) on either side of the slash refer(s) to the two (2) copies of the patient DNA. Del and dashes denotes nucleotide indels in patient DNA. Empty cells indicate an absence of genotyping results.
- 4. Phenotype refers to the CYP specific drug metabolising capabilities of an individual.

See RISKS AND LIMITATIONS on the last pages of this Report.



## **PGx Report - Pain Management**

Type: Anti-inflammatory Agent, Analgesic, Antipyretic

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		The Nonsteroidal Antiinfla	ammatory Drugs (NSAIDs)			
	<u>Diclofenac</u>	UGT2B7	CYP2C9, UGT1A3, UGT1A9, CYP2E1, CYP3A4		<b>Ø</b>	
Acetic acid derivatives	<u>Nabumetone</u>	CYP1A2	CYP2C19, CYP3A4			
	Indomethacin	CYP2C9	CYP2C19			
	Meloxicam	CYP2C9	CYP1A2, CYP3A4, CYP3A5			
Enolic acid (Oxicam)	<u>Piroxicam</u>	CYP2C9	CYP3A4, CYP3A5			
derivatives	Tenoxicam	CYP2C9				
	Lornoxicam	CYP2C9				
	<u>Etoricoxib</u>	CYP3A4	CYP3A5, CYP2C9, CYP2D6, CYP1A2			
Selective COX-2 inhibitors (Coxibs)	<u>Parecoxib</u>	CYP2C9	CYP3A4, CYP3A5			
(GONIDO)	<u>Celecoxib</u>	CYP2C9	CYP2C19			
	<u>Ibuprofen</u>	CYP2C9	CYP2C19, CYP2C8, UGT1A3, UGT1A9, UGT2B7		<b>Ø</b>	
	<u>Flurbiprofen</u>	CYP2C9				
Propionic acid derivatives	Ketoprofen	CYP3A4	CYP2C9, CYP3A5, UGT1A6, UGT1A9, UGT2B7		<b>Ø</b>	
·	<u>Fenoprofen</u>	CYP2C9	UGT2B7		<b>Ø</b>	
	<u>Vicoprofen</u>	CYP2D6	CYP3A4			
	<u>Naproxen</u>	CYP2C9	CYP1A2, CYP2C8, UGT2B7, SULT1A1, UGT1A3, UGT1A6, UGT1A9		<b>Ø</b>	
Anthranilic acid derivatives (Fenamates)	Mefenamic acid	CYP2C9			<b>Ø</b>	
The Non-NSAIDs Analgesic	<u>Paracetamol</u>	UGT1A1, UGT1A6, UGT1A9, SULT1A1, GSHs	CYP2E1, CYP3A4, CYP3A5, CYP2D6, CYP1A2, ABCG2			<b>W</b>



## **PGx Report - Pain Management**

Type: Opioid

Drug Class	Generic Primary Mechanism Involved		Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Opioid A				,
Opium alkaloids	<u>Morphine</u>	UGT2B7	ABCB1, ABCC3, UGT1A1, UGT1A8, OPRM1, COMT		<b>Ø</b>	
Opium aikaioius	<u>Codeine</u>	CYP2D6	CYP3A4, UGT2B7, UGT2B4, FMO3, CYP3A5, OPRM1		<b>Ø</b>	
Esters of morphine	Diacetylmorphine (Heroin)	CES1	CES2, BCHE, OPRM1			
Ethan of acception	<u>Dihydrocodeine</u>	CYP3A4	CYP2D6, CYP3A5			
Ethers of morphine	<u>Ethylmorphine</u>	CYP2D6	CYP3A4, CYP3A5			
	<u>Hydrocodone</u>	CYP2D6	CYP3A4, CYP3A5, OPRM1			
Semi-synthetic alkaloid	<u>Hydromorphone</u>	UGT2B7				
derivatives	<u>Oxycodone</u>	CYP3A4	CYP3A5, CYP2D6, ABCB1, UGT2B7, COMT			<b>&gt;</b>
	<u>Oxymorphone</u>	UGT2B7				
		Syntheti	c opioids			
	<u>Alfentanyl</u>	CYP3A4	CYP3A5, ABCB1, OPRM1			
Anilidopiperidine derivatives	<u>Fentanyl</u>	CYP3A4	CYP3A5, ABCB1, OPRM1		<b>Ø</b>	
	<u>Sufentanil</u>	CYP3A4	CYP3A5, OPRM1			
Phenylpiperidine derivatives	<u>Meperidine</u>	CYP2B6	CYP3A4, CYP2C19, CYP3A5, UGT1A4		<b>Ø</b>	
	<u>Ketobemidone</u>	CYP2C9	CYP3A4, CYP3A5		<b>Ø</b>	
	<u>Dextropropoxyphene</u>	CYP3A4	CYP3A5, Renal Excretion			<b>W</b>
Diphenylpropylamine	<u>Levacetylmethadol</u>	CYP3A4	CYP3A5			
derivatives	<u>Loperamide</u>	CYP3A4	CYP2C8, CYP3A5			
	<u>Methadone</u>	CYP3A4	CYP2B6, CYP2D6, CYP3A5, ABCB1, UGT2B7, COMT		<b>Ø</b>	
Oripavine derivatives	<u>Buprenorphine</u>	CYP3A4	CYP3A5, CYP2C8, UGT1A1, UGT1A3, UGT2B7		<b>Ø</b>	
Morphinan derivatives	<u>Dextromethorphan</u>	CYP2D6	CYP3A4, CYP3A5			
	<u>Tramadol</u>	CYP2D6	CYP3A4, CYP2B6, CYP3A5, OPRM1, SLC22A1, COMT		0	
Others	<u>Tapentadol</u>	CYP2C9	CYP2C19, CYP2D6			
	<u>Tilidine</u>	CYP3A4	CYP2C19, CYP3A5			<b>&gt;</b>
	<u>Methylnaltrexone</u>	CYP2D6	CYP3A4, CYP3A5			
Anti-opioid	<u>Naloxone</u>	UGT2B7	UGT1A3			
	<u>Naltrexone</u>	UGT2B7	UGT1A1, UGT1A3, OPRM1		<u> </u>	

### **Additional SNPs of Importance for Pain Management**

Gene	Marker	Genotype	Drug	Level of Evidence	Results
OPRM1	rs1799971	A/A	Naloxone	2B	Patients may have lower cortisol response
OPRM1	rs1799971	A/A	Morphine	2B	Pain patients may experience increased efficacy of opioids and may be less susceptible to opioid addiction, and may require a decreased dose of opioids
OPRM1	rs1799971	A/A	Alfentanil	2B	Pain patients may experience increased efficacy of opioids and may be less susceptible to opioid addiction, and may require a decreased dose of opioids
OPRM1	rs1799971	A/A	Fentanyl	2B	Pain patients may experience increased efficacy of opioids and may be less susceptible to opioid addiction, and may require a decreased dose of opioids
OPRM1	rs1799971	A/A	Tramadol	2B	Pain patients may experience increased efficacy of opioids and may be less susceptible to opioid addiction, and may require a decreased dose of opioids
OPRM1	rs1799971	A/A	Hydrocodone	3	Patients may have a decreased risk for experiencing side effects, including constipation, dry mouth or respiratory depression
COMT	rs4680	G/G	Paroxetine	3	Patients may require a higher dose



### **PGx Report - Pain Management**

Type: Drugs Prescribed for the Treatment of Gout, Antirheumatic

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Drugs Prescr	ibed for Gout			
Uricosurics	<u>Sulfinpyrazone</u>	CYP2C9	CYP3A4, CYP3A5			
Mitotic inhibitors	Colchicine	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Febuxostat</u>	CYP1A2, CYP2C8	CYP2C9, UGT1A1, UGT1A3, UGT1A9, UGT2B7		<b>Ø</b>	
Xanthine oxidase inhibitors	<u>Allopurinol</u>	AOX1	Renal Excretion, HLA-B*5801		0	
	<u>Oxypurinol</u>	Renal Excretion				
Recombinant urate oxidase	Rasburicase		G6PD, CYB5R1, CYB5R2, CYB5R3, CYB5R4		<b>Ø</b>	_
DMARDs	<u>Leflunomide</u>	CYP1A2			<b>Ø</b>	
Anti-inflammatory	<u>Tofacitinib</u>	CYP3A4	CYP2C19, CYP3A5			<b>W</b>
	Abbreviatio	ns: DMARDs, Disease-modifying antirher	umatic drugs; RE, renal excretion (unchai	nged drug).		

## **PGx Report - Modulation of Cardiovascular Function**

Type: Antiarrhythmic

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
	<u>Quinidine</u>	CYP3A4, CYP2D6	CYP2E1, CYP3A5, CYP2C9, CYP2C8		<b>Ø</b>	
Antiarrhythmic class la	<u>Procainamide</u>	CYP2D6	NAT2		<b>Ø</b>	
Antiarmytinnic class ia	<u>Sparteine</u>	CYP2D6			<b>Ø</b>	
	<u>Disopyramide</u>	CYP3A4	CYP3A5, CYP1A2, CYP2C19			<b>W</b>
	Phenytoin	CYP2C19	CYP2C9, CYP3A4, CYP3A5, CYP2D6, ABCB1, EPHX1, HLA-B*1502			<b>W</b>
Antiarrhythmic class lb	<u>Tocainide</u>	UGTs			<b>Ø</b>	
, , , , , , , ,	<u>Lidocaine</u>	CYP1A2	CYP3A4, CYP3A5			
	<u>Mexiletine</u>	CYP2D6	CYP1A2			
	<u>Propafenone</u>	CYP2D6	CYP3A4, CYP1A2, CYP3A5			
Antiarrhythmic class Ic	<u>Flecainide</u>	CYP2D6				
	<u>Encainide</u>	CYP2D6				
	<u>Carvedilol</u>	CYP2D6	UGT1A1, UGT2B4, CYP2C9			
	<u>Bisoprolol</u>	CYP2D6	CYP3A4, CYP3A5			
Antiarrhythmic class II	<u>Metoprolol</u>	CYP2D6	CYP3A4, CYP3A5			
	<u>Propranolol</u>	CYP2D6	CYP1A2, CYP2C19, CYP3A4, CYP3A5, UGT1A9		<b>Ø</b>	
	<u>Amiodarone</u>	CYP3A4	CYP2C8, CYP3A5			
Antiarrhythmic class III	<u>Dronedarone</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Dofetilide</u>	Renal Excretion	CYP3A4, CYP3A5			
Antiarrhythmic class IV	<u>Diltiazem</u>	CYP3A4	CYP2C19, CYP3A5			<b>W</b>
Annamy minic class IV	<u>Verapamil</u>	CYP3A4	CYP2C8, CYP3A5, ABCB1			



### **PGx Report - Modulation of Cardiovascular Function**

Type: Antihypertensive I

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antihype	rtensives			
	<u>Losartan</u>	CYP2C9	CYP3A4, CYP3A5, UGT1A1, UGT1A3		<b>Ø</b>	
	<u>Azilsartan</u>	CYP2C9				
Angiotensin II receptor	<u>Irbesartan</u>	CYP2C9				
antagonist	<u>Telmisartan</u>	Biliary Excretion	UGT1A1			
	<u>Olmesartan</u>	Hydrolysis	Renal Excretion, SLCO1B1			<b>6</b>
	<u>Valsartan</u>	CYP2C9				
	Captopril	Renal Excretion	CYP2D6			
Angiotensin-Converting Enzyme Inhibitors	<u>Enalapril</u>	CES1, Renal Excretion	CYP3A4, CYP3A5			<b>W</b>
Enzymo minotoro	Trandolapril	CES1	CYP2D6, CYP2C9, Renal Excretion			
Renin inhibitors	Aliskiren	CYP3A4	CYP3A5, ABCB1			
Aldosterone Antagonists	<u>Eplerenone</u>	CYP3A4	CYP3A5			
Loop diuretic	<u>Torasemide</u>	CYP2C9	CYP2C8, Renal Excretion			
Loop diuretic	<u>Furosemide</u>	Renal Excretion	UGT1A9, UGT1A10			
Potassium-sparing diuretic	<u>Triamterene</u>	CYP1A2				
Vasopressin receptor antagonists	<u>Tolvaptan</u>	CYP3A4	CYP3A5		Ø	
Adrenergic release inhibitors	<u>Debrisoquine</u>	CYP2D6				
Peripheral Adrenergic Inhibitors	Reserpine	CYP2D6			0	
	<u>Metoprolol</u>	CYP2D6	CYP3A4, CYP3A5			
Beta-1 cardioselective beta- blockers	<u>Bisoprolol</u>	CYP2D6	CYP3A4, CYP3A5			
2.00.010	Nebivolol	CYP2D6				

### **PGx Report - Modulation of Cardiovascular Function**

Type: Antihypertensive II

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antihype	rtensives			
	<u>Timolol</u>	CYP2D6				
Nonselective beta-blockers	<u>Propranolol</u>	CYP2D6	CYP1A2, CYP2C19, CYP3A4, CYP3A5, UGT1A9		<b>Ø</b>	
Beta-blockers with alpha	Carvedilol	CYP2D6	UGT1A1, UGT2B4, CYP2C9			
activity	<u>Labetalol</u>	CYP2D6	CYP2C19, ABCB1, UGT1A1, UGT1A9, UGT2B7		<b>Ø</b>	
Alpha blockers	<u>Terazosin</u>	CYP3A4	CYP3A5			
Alpha blockers	<u>Doxazosin</u>	CYP2D6	CYP2C19, CYP3A4, CYP3A5			<b>W</b>
α-2 adrenergic agonist	<u>Clonidine</u>	CYP2D6	CYP1A2, CYP3A4, CYP3A5			
α-2 aurenergic agonist	<u>Tizanidine</u>	CYP1A2				
		Antihypertensives Cal	cium channel blockers			
	<u>Amlodipine</u>	CYP3A4	CYP3A5			
Dihydropyridine	<u>Nifedipine</u>	CYP3A4	CYP1A2, CYP2A6, CYP3A5		<b>Ø</b>	
Diffydropyfidiffe	Nimodipine	CYP3A4	CYP3A5			
	<u>Nicardipine</u>	CYP2C8	CYP2D6, CYP3A4, CYP3A5			
Benzothiazepine	<u>Diltiazem</u>	CYP3A4	CYP2C19, CYP3A5			<b>64</b>
Phenylalkylamine	<u>Verapamil</u>	CYP3A4	CYP2C8, CYP3A5, ABCB1			
Nonselective	<u>Bepridil</u>	CYP3A4	CYP3A5			
		Anti-pulmonary ar	terial hypertension			
ERA-Dual antagonists	<u>Bosentan</u>	CYP2C9	CYP3A4, CYP3A5, SLCO1B3			
Li ii i Duai amayomsis	<u>Macitentan</u>	CYP3A4	CYP2C19, CYP3A5			<b>W</b>
Phosphodiesterase inhibitors	<u>Sildenafil</u>	CYP3A4	CYP2C9, CYP3A5		<b>Ø</b>	
i nosphoulesterase initibitors	<u>Tadalafil</u>	CYP3A4	CYP3A5		<b>Ø</b>	
		Abbreviations: ERA, endo	thelin receptor antagonist.			



### **PGx Report - Modulation of Cardiovascular Function**

Type: Cardiac stimulant, Vasodilator, Drugs Prescribed for the Treatment of Angina

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Cardiac s	timulants			
Digitalis glycosides	<u>Digoxin</u>	Renal Excretion	ABCB1, SLCO1B3, ABCB4			
	<u>Epinephrine</u>	MAO	COMT		0	
Adrenergic and dopaminergic	<u>Phenylephrine</u>	MAO	SULTs, UGTs		<b>Ø</b>	
agents	<u>Dopamine</u>	ALDH1A1, ALDH2	DBH, MAOA, MAOB, SULT1A3, SULT1A4, COMT		<b>Ø</b>	
	<u>Synephrine</u>	MAO			0	
		Vasodilators used i	n cardiac diseases			
		Other Drugs U	Ised in Angina			
Other cardiac preparations	<u>Ranolazine</u>	CYP3A4	CYP2D6, CYP3A5			
other durated proparations	<u>Ivabradine</u>	CYP3A4	CYP3A5		<b>Ø</b>	

### **PGx Report - Modulation of Cardiovascular Function**

Type: Dyslipidemia

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Drug Therapy for Hypercholeste	rolemia and Dyslipidemia (Liver)			
	Atorvastatin	CYP3A4, HMGCR	HMGCR, ABCG2, CYP3A5, ABCB1, ABCG8, UGT1A1, UGT1A3, UGT2B7, KIF6		<b>Ø</b>	
	Fluvastatin	CYP2C9, SLCO1B1	HMGCR, ABCG2, CYP3A4, CYP2C8, UGT1A1, UGT1A3, UGT2B7		<b>Ø</b>	
	<u>Lovastatin</u>	CYP3A4, SLCO1B1	CYP3A5, HMGCR, UGT1A1, UGT1A3		<b>Ø</b>	
HMG CoA reductase	<u>Cerivastatin</u>	CYP3A4, SLCO1B1	HMGCR, CYP2C8, CYP3A5			
inhibitors Statins	<u>Pitavastatin</u>	UGT1A3, UGT2B7	CYP2C9, CYP2C8, ABCB1, HMGCR			
	<u>Pravastatin</u>	SLCO1B1, HMGCR	KIF6, APOE, ABCA1			
	Simvastatin	CYP3A4, SLCO1B1	ABCG2, HMGCR, CYP3A5, ABCB1, SLCO2B1, UGT1A1, UGT1A3, UGT2B7, KIF6		0	
	Rosuvastatin	UGT1A1	UGT1A3, ABCG2, HMGCR			
MTTP inhibitors	<u>Lomitapide</u>	CYP3A4	CYP3A5, LDLR			
		Drug Therapy for Hypercholest	erolemia and Dyslipidemia (GI)	'		
Cholesterol absorption inhibitors	<u>Ezetimibe</u>	UGT1A1	UGT1A3, UGT2B15			
		Drug Therapy for Hypercholesteroler	mia and Dyslipidemia (Blood vessels)			
Fibrates	<u>Gemfibrozil</u>	CYP3A4	CYP3A5, UGT2B7, UGT1A1, UGT1A3, UGT1A9, UGT2B15			
	<u>Clofibrate</u>	UGT2B7			<b>Ø</b>	
		Drug Therapy for familia	al hypercholesterolemia	'		
Cholesterol-reducing drug (antisense oligonucleotide)	Mipomersen	Nuclease, Renal Excretion	LDLR		<b>Ø</b>	
Abbreviations: MTTP, microso	omal triglyceride transfer p	orotein; GI, gastrointestinal tract. Rosuvasta CY		ative Statins since a	re not extensively n	netabolised by the

### **Additional SNPs of Importance for Treatment Using Statins**

Gene	Marker	Genotype	Drug	Level of Evidence	Results
KIF6	rs20455	A/G	Atorvastatin	3	Intermediate risk for adverse cardiovascular events
KIF6	rs20455	A/G	Pravastatin	3	Intermediate risk for adverse cardiovascular events
APOE	rs7412	C/C	Atorvastatin	2A	Less responsive to Statin treatment
APOE	rs7412	C/C	Pravastatin	3	Less responsive to Statin treatment
APOE	rs7412	C/C	Simvastatin	3	Less responsive to Statin treatment
ITGB3	rs5918	T/T	Clopidogrel	3	Patients may have an increased antiplatelet effect to a 300 or 600 mg loading dose of Clopiodgrel



### **PGx Report - Modulation of Cardiovascular Function**

Type: Anticoagulant, Antiplatelet

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Blood Coagulation and Anticoa	agulant, and Antiplatelet Drugs			
	<u>Warfarin</u>	CYP2C9, VKORC1	CYP4F2, CYP2C19, CYP1A2, CYP3A4, EPHX1, PROC, PROS1		<b>Ø</b>	
Vitamin K antagonist	<u>Acenocoumarol</u>	CYP2C9, VKORC1	CYP4F2, CYP2C19, CYP1A2			
	<u>Phenprocoumon</u>	CYP2C9, VKORC1	CYP4F2, CYP3A4, CYP2C8, EPHX1			
Direct factor Xa inhibitors	Rivaroxaban	CYP3A4	CYP2J2, CYP3A5			
Direct factor Ad Illinoitors	<u>Apixaban</u>	CYP3A4	CYP3A5			
		Antiplate	et Drugs			
ADP receptor (P2Y12) inhibitors Nucleotide/nucleo side analogues	Ticagrelor	CYP3A4	CYP3A5		<b>Ø</b>	
ADP receptor (P2Y12)	Clopidogrel	CYP2C19	ABCB1, ABCC3			
inhibitors Thienopyridines	<u>Prasugrel</u>	BCHE, CYP3A4	CYP2B6, CYP2C9, CYP2C19, CYP3A5, CYP2D6			<b>\</b>
Irreversible cyclooxygenase inhibitors	<u>Aspirin</u>	GLYAT, UGTs, Renal Excretion	CYP2C9, CYP3A4, CYP3A5		<b>Ø</b>	
Phosphodiesterase inhibitors	Cilostazol	CYP3A4	CYP2C19, CYP3A5			<b>W</b>
Protease-activated receptor-1 (PAR-1) antagonists	<u>Vorapaxar</u>	CYP3A4	CYP2J2, CYP3A5		<b>Ø</b>	
		Abbreviations: P2Y12, pu	rinergic receptor P2Y12.			

### **PGx Report - Internal Medicine**

Type: Drugs Prescribed for the Modulation of Respiratory Function

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Respi	ratory			
Anticholinergic	<u>Umeclidinium</u>	CYP2D6				
Anticholinergic	<u>Aclidinium</u>	CYP2D6	CYP3A4, CYP3A5		<b>Ø</b>	
	<u>Arformoterol</u>	CYP2D6, UGT1A1	CYP2C19			
	<u>Indacaterol</u>	UGT1A1, CYP3A4	CYP3A5, CYP1A2, CYP2D6			
Beta2-adrenergic agonist	<u>Formoterol</u>	CYP2D6	CYP2C19, CYP2C9, CYP2A6			
	Salmeterol	CYP3A4	CYP3A5			
	<u>Vilanterol</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Budesonide</u>	CYP3A4	CYP3A5		<b>Ø</b>	
Corticosteroid	<u>Fluticasone</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	Mometasone	CYP3A4	CYP3A5			
Discoule of the second section of	Roflumilast	CYP3A4	CYP1A2, CYP3A5			
Phosphodiesterase inhibitor	Theophylline	CYP1A2	CYP2E1			
5-lipoxygenase inhibitor	<u>Zileuton</u>	CYP1A2	CYP2C9, CYP3A4, CYP3A5		0	
	<u>Montelukast</u>	CYP3A4	CYP2C9, CYP3A5, SLCO2B1, ABCC1		0	
Leukotriene receptor-1 antagonist	<u>Pranlukast</u>	CYP3A4	CYP3A5			
anayonsi	<u>Zafirlukast</u>	CYP2C9	CYP3A4, CYP3A5		0	
Treatment of cystic fibrosis (specifics mutations in the CFTR gene)	Ivacaftor	CYP3A4	CYP3A5, CFTR		0	
		Abbreviations: CFTR, Cystic fibrosis tr	ansmembrane conductance regulator.			



### **PGx Report - Internal Medicine**

Type: Antiemetic

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antie	metic			
Antiemetic, 5-HT3 receptor	Dolasetron	CYP3A4	CYP2D6, CYP3A5		<b>Ø</b>	
antagonist Indole derivative	Tropisetron	CYP3A4	CYP2D6, CYP3A5			
Antiemetic, 5-HT3 receptor antagonist Isoquinoline derivative	Palonosetron	CYP1A2	CYP2D6, CYP3A4, CYP3A5		<b>Ø</b>	
Antiemetic, 5-HT3 receptor antagonist Indazole derivative	Granisetron	CYP3A4	CYP3A5			
Antiemetic, 5-HT3 receptor antagonist	Ondansetron	CYP2B6	CYP1A2, CYP2D6, CYP3A4, ABCB1			
	<u>Domperidone</u>	CYP3A4	CYP3A5		<b>Ø</b>	
Antiemetic, dopamine-	<u>Prochlorperazine</u>	CYP2D6	CYP3A4, CYP3A5			
receptor antagonist	Metoclopramide	CYP2D6	CYP1A2, CYB5R1, CYB5R2, CYB5R3, CYB5R4		<b>Ø</b>	
Antiemetic, NK1 receptor antagonist	<u>Aprepitant</u>	CYP3A4	CYP3A5, CYP1A2, CYP2C19			<b>W</b>
	<u>Diphenhydramine</u>	CYP2D6	CYP3A4, CYP3A5, UGT1A3, UGT1A4		<b>Ø</b>	
Antiemetic, H1 histamine receptor antagonist	<u>Hydroxyzine</u>	ADHs	CYP3A4, CYP3A5		<b>Ø</b>	
- Coopie amagemen	Promethazine	CYP2D6	UGT1A3, UGT1A4, SULTs		<b>Ø</b>	
Cannabinoids	<u>Dronabinol</u>	CYP2C9	CYP2C19, CYP3A4, CYP3A5			
Benzodiazepines	<u>Lorazepam</u>	UGT2B15	UGT2B7			
Derizoulazepines	Midazolam	CYP3A4	CYP3A5		<b>2</b>	
Anticholinergics	Scopolamine	CYP3A4	CYP3A5			
Steroids	<u>Dexamethasone</u>	CYP3A4	CYP17A1, CYP3A5		<b>2</b>	
		Abbreviations: 5-HT, Ser	otonin; NK1, neurokinin 1.			

## **PGx Report - Internal Medicine**

Type: Drugs Prescribed for the Treatment of Peptic Ulcers and/or Gastro-Oesophageal Reflux Disease

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
Histamine H2-receptor antagonists	<u>Ranitidine</u>	Renal Excretion	CYP1A2, CYP2C19, FMO3, CYP3A4, CYP3A5			<b>\undersity</b>
	<u>Omeprazole</u>	CYP2C19	CYP3A4, CYP2C9, CYP3A5			<b>W</b>
	<u>Dexlansoprazole</u>	CYP2C19	CYP3A4, CYP3A5			<b>W</b>
	<u>Esomeprazole</u>	CYP2C19	CYP3A4, CYP3A5			<b>W</b>
Proton-pump inhibitor	Lansoprazole	CYP3A4	CYP2C19, CYP3A5			<b>W</b>
	Rabeprazole	Non Enz	CYP2C19, CYP3A4, CYP3A5			<b>W</b>
	<u>llaprazole</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Pantoprazole</u>	CYP2C19	CYP3A4, CYP2D6, CYP2C9, CYP3A5			<b>W</b>
		Abbreviations: Non Enz, n	on-enzymatic metabolism.			



### **PGx Report - Internal Medicine**

Type: Drugs Prescribed for the Treatment of Functional Gastrointestinal Disorders, Obesity

Drug Class Generic		Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Drugs for functional ga	strointestinal disorders			
Acting on serotonin receptors	Alosetron	CYP2C9	CYP3A4, CYP1A2			
5-HT3 antagonists	Cilansetron	CYP3A4	CYP2D6, CYP1A2, CYP2C19, CYP3A5		<b>Ø</b>	
Acting on serotonin receptors	<u>Mosapride</u>	CYP3A4	CYP3A5			
5-HT4 agonists	<u>Prucalopride</u>	Renal Excretion	CYP3A4, CYP3A5			
		Gastrop	rokinetic			
Serotonin 5-HT4 receptor	<u>Cisapride</u>	CYP3A4	CYP3A5			
agonist	<u>Cinitapride</u>	CYP3A4	CYP2C8, CYP3A5		<b>Ø</b>	
Parasympatho mimetic	<u>Itropride</u>	FMO3				
_	Metoclopramide	CYP2D6	CYP1A2, CYB5R1, CYB5R2, CYB5R3, CYB5R4		0	
Dopamine antagonists	<u>Clebopride</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Domperidone</u>	CYP3A4	CYP3A5		<b>Ø</b>	
		Antiprop	oulsives			
	<u>Loperamide</u>	CYP3A4	CYP2C8, CYP3A5			
Opioids	<u>Morphine</u>	UGT2B7	ABCB1, ABCC3, UGT1A1, UGT1A8, OPRM1, COMT		<b>Ø</b>	
·		Centrally acting a	inti-obesity drugs			
Stimulant/ Amphetamine/	Sibutramine	CYP3A4	CYP3A5			
Appetite suppressant agent	<u>Phentermine</u>	Renal Excretion	CYP3A4, CYP3A5			
Anorectic	<u>Lorcaserin</u>	CYP2D6	CYP3A4, CYP3A5			

### **PGx Report - Internal Medicine**

Type: Diabetes

Drug Class			Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antidiabetic S	ecretagogues			
Meglitinides	<u>Repaglinide</u>	CYP2C8	SLCO1B1, CYP3A4, CYP3A5, ABCC8			
Wegittilides	<u>Nateglinide</u>	CYP2C9	CYP3A4, CYP3A5		<b>Ø</b>	
	<u>Chlorpropamide</u>	Renal Excretion	CYP2D6, G6PD			
Sulfonylurea 1st generation	<u>Tolazamide</u>	CYP2C9				
	<u>Tolbutamide</u>	CYP2C9	CYP2C19, CYP2C8			
	<u>Glipizide</u>	CYP2C9	G6PD			
	<u>Glyburide</u>	CYP3A4	CYP2C9, CYP2C19, CYP3A5, G6PD			
Sulfonylurea 2nd generation	<u>Gliquidone</u>	CYP2C9				
	Gliclazide	CYP2C9	CYP2C19			
	<u>Glimepiride</u>	CYP2C9	G6PD			
	Saxagliptin	CYP3A4	CYP3A5			
DPP-IV inhibitor	Alogliptin	Renal Excretion	CYP2D6, CYP3A4, CYP3A5			
DPP-IV IIIIIIDILOI	Linagliptin	Renal Excretion	CYP3A4, CYP3A5			
	Sitagliptin	CYP3A4	CYP2C8, CYP3A5			
		Antidiabetic	Sensitisers	·		
Thiazolidinediones	<u>Pioglitazone</u>	CYP2C8	CYP3A4, CYP3A5		<b>Ø</b>	
THIAZUIUITIEUIUTIES	Rosiglitazone	CYP2C8	CYP2C9		<b>Ø</b>	
		Antidiabe	etic Other			
SGLT2 inhibitors	Canagliflozin	UGT1A9, UGT2B4	CYP3A4, CYP3A5		<b>Ø</b>	
	Abbrevia	tions: DPP-IV, Dipeptidyl peptidase-4; SC	GLT2, sodium/glucose cotransporter 2 or	gliflozins.		



## **PGx Report - Internal Medicine**

Type: Migraine, Antihistamine, Abortifacient, Drugs Prescribed for the Treatment of Hyperparathyroidism, Dermatology

Drug Class Generic		Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Anti-m	igraine			
	<u>Almotriptan</u>	CYP3A4	CYP2D6, CYP3A5			
	<u>Eletriptan</u>	CYP3A4	CYP3A5			
elective serotonin (5-HT1)	<u>Frovatriptan</u>	CYP1A2				
agonists	<u>Naratriptan</u>	CYP1A2	CYP2C8, CYP2C9, CYP2D6			
	<u>Sumatriptan</u>	MAO	UGTs, HTR2A			
	<u>Zolmitriptan</u>	CYP1A2			<b>Ø</b>	
Ergot alkaloids	<u>Dihydroergotamine</u>	CYP3A4	CYP3A5		<b>Ø</b>	
Ligot aikaioius	<u>Ergotamine</u>	CYP3A4	CYP3A5		<b>Ø</b>	
			tamines	'		
Aminoalkyl ethers	<u>Diphenhydramine</u>	CYP2D6	CYP3A4, CYP3A5, UGT1A3, UGT1A4			
Substituted alkylamines	Chlorpheniramine	CYP3A4	CYP3A5		<b>Ø</b>	
Phenothiazine derivatives	<u>Promethazine</u>	CYP2D6	UGT1A3, UGT1A4, SULTs			
	<u>Hydroxyzine</u>	ADHs	CYP3A4, CYP3A5		<b>Ø</b>	
Piperazine derivatives	<u>Cyclizine</u>	CYP2D6			<b>Ø</b>	
	<u>Cetirizine</u>	Renal Excretion				
	<u>Terfenadine</u>	CYP3A4	CYP3A5			
	<u>Loratadine</u>	CYP3A4, CYP2D6	CYP3A5, CYP2C8, CYP2C9			
Other antihistamines	<u>Fexofenadine</u>	Biliary Excretion	Renal Excretion, CYP3A4, CYP3A5, SLCO2B1		<b>Ø</b>	
	<u>Desloratadine</u>	CYP2C8	UGT2B10		<b>Ø</b>	
	<u>Astemizole</u>	CYP3A4	CYP3A5		<b>Ø</b>	
		Treatment of secondar	ry hyperparathyroidism			
Calcimimetic	<u>Cinacalcet</u>	CYP3A4	CYP2D6, CYP3A5, CYP1A2			
			facient			
Progestin Antagonist	<u>Mifepristone</u>	CYP3A4	CYP3A5			
		•	Antipsoriatics			
Retinoids	Etretinate	CYP26A1				
	<u>Acitretin</u>	CYP26A1				
D			gy Anti-acne			
Retinoid	<u>Isotretinoin</u>	CYP2C8 Abbreviations: BE	CYP2C9, CYP3A4, CYP2B6, CYP3A5			



## **PGx Report - Psychiatry**

Type: Antidepressant I

Drug Class Generic		Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antidep	ressants			
	<u>Citalopram</u>	CYP2C19, CYP2D6	CYP3A4, CYP3A5, SLC6A4, HTR2A			<b>W</b>
	<u>Escitalopram</u>	CYP3A4, CYP2C19	CYP2D6, CYP3A5, SLC6A4, HTR2C			<b>W</b>
	<u>Dapoxetine</u>	CYP2D6	CYP3A4, CYP3A5, FMO1			
SSRIs	Fluoxetine	CYP2D6	CYP3A4, CYP2C9, CYP3A5, CYP2C19, SLC6A4, HTR2A		0	
	<u>Paroxetine</u>	CYP2D6	CYP3A4, CYP1A2, CYP3A5, CYP2C9, SLC6A4, HTR2A, DRD3		<b>Ø</b>	
	<u>Sertraline</u>	CYP2B6	CYP2C19, CYP2C9, CYP3A4, CYP2D6, SLC6A4		<b>Ø</b>	
	<u>Fluvoxamine</u>	CYP2D6	CYP1A2, SLC6A4, HTR2A		<b>Ø</b>	
SMSs	<u>Vilazodone</u>	CYP3A4	CYP3A5, CYP2C19, CYP2D6			<b>6</b>
	<u>Levomilnacipran</u>	CYP3A4	CYP2C8, CYP3A5, CYP2C19, CYP2D6		<b>Ø</b>	
SNRIs	<u>Milnacipran</u>	UGTs	Renal Excretion			<b>W</b>
Sivnis	<u>Venlafaxine</u>	CYP2D6	CYP2C19, CYP3A4, CYP2C9, CYP3A5, SLC6A3, SLC6A4, HTR2A		<b>Ø</b>	
	<u>Duloxetine</u>	CYP2D6	CYP1A2, HTR2A			
	<u>Atomoxetine</u>	CYP2D6	CYP2C19, CYP3A4, CYP3A5, SLC6A2			
NRIs	Reboxetine	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Maprotiline</u>	CYP2D6	CYP1A2		<b>Ø</b>	
TCAs that preferentially inhibit the reuptake of	Clomipramine	CYP2D6	CYP3A4, CYP2C19, CYP1A2, CYP2C9, SLC6A4, HTR2A		0	
serotonin	<u>Imipramine</u>	CYP1A2, CYP2D6	CYP2C19, CYP3A4, CYP3A5, UGT1A3, UGT1A4			
TCAs that preferentially	<u>Desipramine</u>	CYP2D6	CYP1A2, CYP2C19		<b>Ø</b>	
inhibit the reuptake of	<u>Nortriptyline</u>	CYP2D6	CYP1A2, CYP2C19, ABCB1, SLC6A4		<b>Ø</b>	
norepinephrine	Protriptyline	CYP2D6				



### **PGx Report - Psychiatry**

Type: Antidepressant II

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antidep	ressants			
TCAs that fairly balanced	<u>Amitriptyline</u>	CYP2D6	CYP3A4, CYP2C19, CYP2C9, CYP1A2, CYP2B6, UGT1A3, UGT1A4		<b>Ø</b>	
serotonin-norepinephrine reuptake inhibitors	<u>Doxepin</u>	CYP2D6, CYP2C19	CYP1A2, CYP3A4, CYP3A5, UGT1A3, UGT1A4			<b>\</b>
reaptane ministers	Dosulepin	CYP2D6, CYP2C9	CYP3A4, CYP1A2, CYP3A5, CYP2C19		<b>Ø</b>	
TeCAs	Mianserin	CYP2D6	CYP3A4, CYP1A2, CYP2B6, CYP3A5			
Teoas	<u>Amoxapine</u>	CYP2D6	CYP3A4, CYP3A5			
TCA with antipsychotic and sedative properties	Trimipramine	CYP2D6	CYP2C19, CYP2C9		0	
MAOI	Tranylcypromine	MAO	CYP3A4, CYP2A6, CYP3A5, CYP2C19, CYP2D6		<b>Ø</b>	
	<u>Moclobemide</u>	CYP2C19	CYP2D6, CYP1A2, HTR2A			<b>W</b>
·		Atypical anti	depressants			
SMSs	<u>Vortioxetine</u>	CYP2D6	CYP2C9, CYP3A4, CYP3A5, UGTs, CYP2A6, CYP2C8, CYP2C19, CYP2B6		<b>Ø</b>	
NaSSAs	<u>Mirtazapine</u>	CYP1A2	CYP2D6, CYP3A4, CYP3A5, SLC6A4, HTR2A		<b>Ø</b>	
CADI	<u>Trazodone</u>	CYP3A4	CYP2D6, CYP3A5			
SARIs	Nefazodone	CYP2D6, CYP3A4	CYP3A5, UGT1A6		<u> </u>	
Antidepressant and smoking cessation aid	<u>Bupropion</u>	CYP2B6	CYP2E1, CYP3A4, CYP2C9, CYP2D6, CYP1A2, CYP3A5		0	
Antidepressant and anti- anxiety	sant and anti-		CYP3A5			

Abbreviations: SSRI, serotonin selective reuptake inhibitor; SMS, Serotonin modulator and stimulator; SNRI, serotonin-norepinephrine reuptake inhibitor; NRI, norepinephrine reuptake inhibitor; TCA, tricyclic antidepressant; TeCA, tetracyclic antidepressant; MAOI, monoamine oxidase inhibitor; NaSSA, noradrenergic and specific serotonergic antidepressant; SARI, serotonin antagonist and reuptake inhibitor.

### **Additional SNPs of Importance for Treatment Using Antidepressants**

Gene	Marker	Genotype	Drug	Level of Evidence	Results
GRIK4	rs1954787	T/T	Citalopram	1B	Patients may have a decreased chance of response to Citalopram treatment
GRIK4	rs1954787	T/T	Antidepressants	2B	Patients with Depressive Disorder or Depression may be less likely to respond to antidepressant treatment
ADRA2A	rs1800544	G/C	SSRIs	3	Patients with major depressive disorder may have an increased response to selective serotonin reuptake inhibitors
GABRP	rs10036156	C/C	SSRIs	3	Patients with Depressive Disorder or Depression may be more likely to respond to antidepressant treatment
GABRP	rs10036156	C/C	Venlafaxine	3	Patients with Depressive Disorder or Depression may be more likely to respond to antidepressant treatment
HTR2A	rs6313	G/G	Paroxetine	3	Patients with depression may have an increased risk of adverse medication reactions

# Additional SNPs of Importance for the Treatment of Depression and Psychosis, and the Treatment of Alcohol and Tobacco Use Disorders

Gene	Marker	Genotype	Drug	Level of Evidence	Results
HTR2A	rs6311	C/C	Escitalopram	3	Patients with anxiety disorder may have a decreased risk of adverse cognitive effects
HTR2A	rs6311	C/C	Fluvoxamine	3	Depressive patients may have an increased risk of gastrointestinal side effects and increased response
COMT	rs4680	G/G	Fluvoxamine	3	Schizophrenia patients may have a decreased risk for developing extrapyramidal symptoms
COMT	rs4680	G/G	Venlafaxine	3	Patients with Depressive Disorder may have increased response but patients with Anxiety Disorders may have a decreased response
COMT	rs4680	G/G	Paroxetine	3	Depressive patients may have a decreased response or decreased improvement
ANKK1/DRD2	rs1800497	G/A	Bupropion	1B	Patients may be less likely to quit smoking
ANKK1/DRD2	rs1800497	G/A	Antipsychotics	2A	Schizophrenia patients may have an increased risk for tardive dyskinesia
ANKK1/DRD2	rs1800497	G/A	Ethanol	2B	Patients may have an increased risk for Alcoholism
ANKK1/DRD2	rs1800497	G/A	Clozapine Olanzapine Risperidone	2B	Patients may have increased risk of side effects including hyperprolactinemia and weight gain
ANKK1/DRD2	rs1800497	G/A	Nicotine	3	Patients may have an increased likelihood of smoking cessation when treated with nicotine replacement therapy
ANKK1/DRD2	rs1800497	G/A	Risperidone	3	Schizophrenia patients may have more improvement in symptoms



### **PGx Report - Psychiatry**

Type: Typical Antipsychotic

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Typical an	tipsychotic			
	<u>Bromperidol</u>	CYP3A4	CYP3A5			
Butyrophenones	<u>Droperidol</u>	CYP3A4	CYP3A5			
	<u>Haloperidol</u>	UGTs, CYP3A4	CYP1A2, CYP2D6, CYP3A5, SLC6A4, HTR2C		<b>Ø</b>	
	Chlorpromazine	CYP2D6	CYP1A2, UGT1A3, UGT1A4, CYP3A4, CYP3A5		<b>Ø</b>	
Phenothiazines with aliphatic	Levomepromazine	CYP3A4	CYP1A2, CYP3A5			
side-chain	<u>Promazine</u>	CYP1A2	CYP3A4, CYP2C19, CYP2C9, CYP3A5			
	Cyamemazine	CYP1A2	CYP3A4, CYP2C9, CYP2C8, CYP3A5			
	Fluphenazine	CYP2D6				
Phenothiazines with	Perphenazine	CYP2D6				
piperazine structure	Prochlorperazine	CYP2D6	CYP3A4, CYP3A5			
	<u>Trifluoperazine</u>	CYP1A2	UGT1A4			
Phenothiazines with piperidine structure	Thioridazine	CYP2D6	CYP1A2, CYP3A4, CYP2C19, CYP3A5		<b>Ø</b>	
Phenothiazines used as an anti-histamine, sedative, and antiemetic	Promethazine	CYP2D6	UGT1A3, UGT1A4, SULTs		0	
Diphenyl-butylpiperidine	<u>Pimozide</u>	CYP3A4, CYP2D6	CYP1A2, CYP3A5			
Thioxanthene derivative	<u>Thiothixene</u>	CYP1A2	CYP3A4, CYP3A5			
Thioxanthene derivative	Zuclopenthixol	CYP2D6	CYP3A4, CYP3A5			
Tricyclics	<u>Loxapine</u>	CYP1A2	CYP3A4, CYP2D6, UGT1A3, UGT1A4, CYP3A5		0	

### **PGx Report - Psychiatry**

Type: Atypical antipsychotic

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Atypical ar	ntipsychotic			
	<u>Quetiapine</u>	CYP3A4, CYP2D6	CYP3A5, CYP1A2, CYP2C9, CYP2C19, SLC6A4		<b>Ø</b>	
Diazepines, Oxazepines, Thiazepines and Oxepines	<u>Asenapine</u>	CYP1A2, UGT1A4	CYP2D6, CYP3A4, CYP3A5			
	<u>Clozapine</u>	CYP1A2, CYP2D6	CYP3A4, FMO3, CYP2C9, CYP2C19, CYP3A5, CYP2A6, UGT1A3, UGT1A4, SLC6A3, SLC6A4, SLC1A1, HTR2C, DRD3			
	<u>Sertindole</u>	CYP2D6	CYP3A4, CYP3A5			
Indole derivatives	<u>Ziprasidone</u>	CYP3A4	AOX1, CYP3A5			
	<u>Lurasidone</u>	CYP3A4	CYP3A5			
Benzamides	<u>Sulpiride</u>	Renal Excretion				
Denzamilies	<u>Amisulpride</u>	Renal Excretion				
	<u>Aripiprazole</u>	CYP2D6	CYP3A4, CYP3A5, DRD3			
-	Risperidone	CYP2D6	CYP3A4, CYP3A5, ABCB1, SLC6A4, SLC1A1, HTR2A, HTR2C, DRD3		<b>Ø</b>	
Other antipsychotics	<u>lloperidone</u>	CYP2D6	CYP3A4, CYP3A5			
	<u>Paliperidone</u>	CYP2D6	CYP3A4, CYP3A5			
	<u>Zotepine</u>	CYP3A4	CYP1A2, CYP3A5, CYP2D6			



# Additional SNPs of Importance in Treatment that Includes the Use of Antipsychotics and for the Treatment of Autism

Gene	Marker	Genotype	Drug	Level of Evidence	Results
HTR2A	rs6311	C/C	Risperidone	3	Children with Autism may have poorer response to treatment
HTR2C	rs3813929	C/C	Olanzapine	3	Patients with psychiatric disorders or schizophrenia may have an increased risk of weight gain
COMT	rs4680	G/G	Haloperidol	3	Schizophrenia patients may have a decreased risk for developing extrapyramidal symptoms
DRD1	rs4532	C/T	Methylphenidate or Dextroamphetamine	3	Patients with attention deficit hyperactivity disorder (ADHD) may have a decreased severity of social withdrawal or nausea

Other genetic and clinical factors may also influence a patient's response to medications.



### **PGx Report - Neurology**

### Type: Drugs Prescribed for the Treatment of ADHD, Related Drugs

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Anti ADHD	Stimulants			
Amphetamine	<u>Dextroamphetamine</u>	Renal Excretion, CYP2D6	DBH, FMO3, GLYAT			<b>W</b>
Amphetamine	<u>Levoamphetamine</u>	Renal Excretion, CYP2D6	FMO3			<b>W</b>
NDRI	<u>Dexmethylphenidate</u>	CYP2D6	Renal Excretion		<b>Ø</b>	
	<u>Lisdexamfetamine</u>	Hydrolysis	CYP2D6, Renal Excretion			<b>W</b>
Psychostimulant	<u>Methylphenidate</u>	CYP2D6	Renal Excretion, SLC6A2, SLC6A3, SLC6A4, DRD3		<b>Ø</b>	
		Anti ADHD N	on-stimulants			
NERI	<u>Atomoxetine</u>	CYP2D6	CYP2C19, CYP3A4, CYP3A5, SLC6A2		<b>Ø</b>	
Central alpha-2 Adrenergic Agonist	Clonidine	CYP2D6	CYP1A2, CYP3A4, CYP3A5		<b>Ø</b>	
	Bupropion	CYP2B6	CYP2E1, CYP3A4, CYP2C9, CYP2D6, CYP1A2, CYP3A5		<b>Ø</b>	
Antidonus	<u>Imipramine</u>	CYP1A2, CYP2D6	CYP2C19, CYP3A4, CYP3A5, UGT1A3, UGT1A4		<b>Ø</b>	
Antidepressants	<u>Desipramine</u>	CYP2D6	CYP1A2, CYP2C19			
	<u>Milnacipran</u>	UGTs	Renal Excretion			<b>6</b>
	Reboxetine	CYP3A4	CYP3A5			
Wakefulness-promoting	Modafinil	Hydrolysis, CYP2D6	CYP1A2, CYP3A4, CYP2B6, CYP3A5			
agent	<u>Armodafinil</u>	CYP3A4	CYP3A5			
		Anti-ins	somnia			
Melatonin Receptor Agonist	Ramelteon	CYP1A2	CYP2C19, CYP3A4, CYP3A5			<b>W</b>
Abbrev	iations: ADHD, Attention defi	cit hyperactivity disorder; NERI; norepine	phrine reuptake inhibitor, NDRI, norepine	phrine-dopamine re	uptake inhibitor.	

### **PGx Report - Neurology**

### Type: Drugs Prescribed for the Treatment of Epilepsy

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antiep	oileptic			
Barbiturates	<u>Phenobarbital</u>	CYP2C19	ABCB1			<b>&gt;</b>
Carbamates	<u>Felbamate</u>	CYP3A4	CYP2E1, CYP3A5			<b>W</b>
Carboxamides	Carbamazepine	CYP3A4, EPHX1	CYP2C8, CYP2B6, UGT2B7, CYP1A2, CYP3A5, ABCB1, HLA-B*1502, HLA- A*3101, ABCC2		<b>Ø</b>	
Fatty acids	<u>Tiagabine</u>	CYP3A4	CYP3A5, CYP1A2, CYP2D6, CYP2C19			<b>W</b>
Fructose derivatives	<u>Topiramate</u>	Renal Excretion	CYPs, UGTs			
GABA analogues	<u>Gabapentin</u>	Renal Excretion				
	<u>Pregabalin</u>	Renal Excretion				
Hydantoin	<u>Phenytoin</u>	CYP2C19	CYP2C9, CYP3A4, CYP3A5, CYP2D6, ABCB1, EPHX1, HLA-B*1502			•
nyuantoin	<u>Mephenytoin</u>	CYP2C19	CYP2C8, CYP2C9, CYP2B6, CYP1A2, CYP2D6		<b>Ø</b>	
Oxazolidinediones	<u>Trimethadione</u>	CYP2C9	CYP2E1, CYP3A4, CYP3A5			
Oxazolidirlediorles	<u>Paramethadione</u>	CYP2C9				
Pyrimidinedione	<u>Primidone</u>	CYP2C9	CYP2C19			
	<u>Brivaracetam</u>	CYP2C19, CYP2C9	CYP3A4, CYP3A5, CYP2C8, CYP2B6			<b>6</b>
Pyrrolidines	Levetiracetam	Renal Excretion				
	Seletracetam	Renal Excretion				
Succinimides	<u>Ethosuximide</u>	CYP3A4	CYP3A5, CYP2E1			•
Sulfonamides	Zonisamide	CYP3A4	CYP2C19, CYP3A5			•
Other	Lacosamide	CYP2C9	CY2C19, CYP3A4			
Other	<u>Perampanel</u>	CYP3A4	CYP3A5		0	
		Abbreviations: GABA, ga	amma-aminobutyric acid.			



## **PGx Report - Neurology**

Type: Anxiolytic, Hypnotic, Sedative, Anticonvulsant, Muscle Relaxants

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Anxiolytic, Hypnotic, Sedative, An	ticonvulsant, and Muscle Relaxant			
	Midazolam	CYP3A4	CYP3A5		<b>Ø</b>	
Benzodiazepine Short-acting	<u>Triazolam</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Brotizolam</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Alprazolam</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Bromazepam</u>	CYP1A2	CYP2D6		<b>Ø</b>	
-	Clobazam	CYP2C19	CYP3A4, CYP3A5, CYP2B6			<b>W</b>
	Flunitrazepam	CYP2C19	CYP2C9, CYP3A4, CYP3A5, NAT2			<b>W</b>
	<u>Estazolam</u>	CYP3A4	CYP3A5			
	Clonazepam	CYP3A4	CYP2C19, CYP3A5, NAT2			<b>W</b>
Benzodiazepine	Oxazepam-r	UGT2B7	UGT1A9			
Intermediate-acting	Oxazepam-s	UGT2B15				
	<u>Quazepam</u>	CYP3A4	CYP2C19, CYP3A5			<b>W</b>
	<u>Lormetazepam</u>	CYP3A4	CYP3A5			
	Lorazepam-r	UGT2B7				
	Lorazepam-s	UGT2B15				
	<u>Nitrazepam</u>	CYP3A4	CYP3A5, NAT2			
	<u>Temazepam</u>	CYP2C19	CYP3A4, CYP3A5, UGT2B7			<b>64</b>
	<u>Diazepam</u>	CYP2C19, CYP3A4	CYP3A5, CYP2B6, CYP1A2			<b>W</b>
	Clorazepate	CYP3A4	CYP3A5			
Benzodiazepine Long-acting	Chlordiazepoxide	CYP3A4	CYP3A5			
	<u>Flurazepam</u>	CYP3A4	CYP3A5		0	
	Nordazepam	CYP3A4	CYP3A5		0	
	Zolpidem	CYP3A4	CYP3A5, CYP1A2, CYP2D6		0	
anhanzadiazanina hun	Zaleplon	AOX1, CYP3A4	CYP3A5		<u> </u>	
onbenzodiazepine hypnotic	Zopiclone	CYP3A4	CYP2C8, CYP2C9, CYP3A5		0	
	Eszopiclone	CYP3A4	CYP2E1, CYP3A5			0.4



### **PGx Report - Neurology**

### Type: Drugs Prescribed for the Treatment of Alzheimer's and Parkinson's, Related Drugs

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Anti-Alzheir	ner disease			
	<u>Tacrine</u>	CYP1A2	CYP2D6			
Acetylcholinesterase inhibitor	<u>Donepezil</u>	CYP2D6	CYP3A4, CYP3A5		<b>Ø</b>	
Acetylcrioiniesterase illilibitor	Rivastigmine	ACHE	BCHE, CHAT			
	<u>Galantamine</u>	CYP2D6	CYP3A4, CYP3A5			
NMDA receptor antagonist	<u>Memantine</u>	Renal Excretion	UGTs			
·		Anti-Parkin:	son disease			
Inhibitor of MAO-B	<u>Selegiline</u>	CYP2B6	CYP2C9, CYP3A4, CYP3A5, CYP2A6, FMO3		<b>Ø</b>	
	<u>Rasagiline</u>	CYP1A2				
COMT inhibitors	Entacapone	UGT1A9, CYP3A4	CYP2A6, CYP3A5, UGT1A6, UGT2B7, UGT2B15		<b>Ø</b>	
	<u>Bromocriptine</u>	CYP3A4	CYP3A5		<b>Ø</b>	
Dopamine receptor agonists	<u>Pramipexole</u>	Renal Excretion	DRD3			
	Ropinirole	CYP1A2	UGTs, Renal Excretion			
Anticholinergics - Antimuscarinics	<u>Diphenhydramine</u>	CYP2D6	CYP3A4, CYP3A5, UGT1A3, UGT1A4			
Anti-hyperkinetic movement	<u>Tetrabenazine</u>	CYP2D6	CYP1A2			
Anti-amyotrophic lateral sclerosis drug	Riluzole	CYP1A2			<b>Ø</b>	
		Anti-multip	le sclerosis			
Dihydroorotate dehydrogenase inhibitor	<u>Teriflunomide</u>	Hydrolysis	NATs , SULTs			
		Improvement of walking in pa	atients with multiple sclerosis			
Selective blocker of members of voltage-activated K+ channels	<u>Dalfampridine</u>	Renal Excretion	CYP2E1			
		Abbreviations: NMDA, N-methyl-D-asparta	ate; COMT, Catechol-O-methyltransferase	е.		

### Additional SNP of Importance for different Medical Condition and personality

Gene	Marker	Genotype	Results
APOE	rs429358	T/T	Normal risk for Alzheimer's disease
GABRA6	rs3219151	T/T	Decreased risk for schizophrenia
DRD4	rs1800955	T/C	Increased susceptibility to novelty seeking personality
ABCG2	rs2231142	G/G	Increased risk for Gout

### **Additional SNP of Importance for hypersensitivity**

Gene	Marker	Genotype	HLA	Drug	Results
HLA-A	rs1061235	A/T	HLA-A*3101	Carbamazepine	3.2-fold increased risk for Carbamazepine hypersensitivity syndrome
FLOT1	rs3909184	G/C	HLA-B*1502	Carbamazepine Lamotrigine Phenytoin	Increased risk for medication-induced hypersensitivity
HCP5	rs2395029	T/T	HLA-B*5701	Abacavir	Normal risk for medication-induced hypersensitivity

The variant allele for rs1061235(T) serves as a proxy for the HLA-A\*3101 allele, the variant allele for rs2990184(C) serves as a proxy for the HLA-B\*1502 allele, the variant allele for rs2995029(G) serves as a proxy for the HLA-B\*5701 allele.



## **PGx Report -Infectious Disease**

Type: Antibiotics

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antibacterials: protein s	ynthesis inhibitors 50S			
Amphenicols	Chloramphenicol	CYP2C9	UGT2B7		<b>Ø</b>	
Lincosamides	Clindamycin	CYP3A4	CYP3A5		<b>Ø</b>	
		Antib	iotic			
Macrolides	Clarithromycin	CYP3A4	CYP3A5			
	<u>Erythromycin</u>	CYP3A4				
	Telithromycin	CYP3A4	CYP3A5		<b>Ø</b>	
		Antibacterials: nuc	leic acid inhibitors			
DHPS inhibitor Intermediate- acting sulfonamides	Sulphamethoxazole	Renal Excretion	NAT2, CYP2C9			
Anaerobic DNA inhibitors/	<u>Tinidazole</u>	CYP3A4	CYP3A5		<b>Ø</b>	
Nitroimidazole	<u>Ornidazole</u>	CYP3A4	CYP3A5			
DNA-dependent RNA	Rifampicin	CYP3A4	CYP2C8, CYP3A5, CYP2C19, CYP2A6, RE			<b>\</b>
polymerase inhibitors	<u>Rifabutin</u>	CYP3A4	CYP1A2, CYP3A5		<b>Ø</b>	
Other drugs against	<u>Bedaquiline</u>	CYP3A4	CYP2C8, CYP2C19, CYP3A5			
mycobacteria	<u>Pyrazinamide</u>	AOX1, XDH	CYP1A2, CYP3A4, CYP3A5, RE			
		Abbreviations: DHPS, Di	hydropteroate synthase.			

## **PGx Report -Infectious Disease**

Type: Antimalarial, Anthelmintic, Antifungal

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antim	alarial			
	<u>Chloroquine</u>	CYP2C8	CYP3A4, CYP3A5, G6PD			
Aminoquinolines	<u>Hydroxychloroquine</u>	CYP2D6	CYP2C8, CYP3A4, CYP3A5		<b>Ø</b>	
Aminoquinonnes	<u>Amodiaquine</u>	CYP2C8				
	<u>Primaquine</u>	CYP2D6	G6PD			
Methanolquinolines	<u>Quinine</u>	CYP3A4, CYP2D6	CYP2C19, CYP3A5, G6PD			
	<u>Mefloquine</u>	CYP3A4	CYP3A5			
	<u>Artemisinin</u>	CYP3A4	CYP2B6, CYP3A5			
Artemisinin and derivatives	<u>Artemether</u>	CYP3A4	CYP3A5			
-	<u>Arteether</u>	CYP3A4	CYP2B6, CYP3A5			
Biguanides	<u>Proguanil</u>	CYP2C19				
Other antimalarials	<u>Halofantrine</u>	CYP3A4	CYP3A5			
Other antimatanais	<u>Pentamidine</u>	CYP2C19	CYP1A2, CYP2D6			<b>W</b>
'		Anthe	mintic			
Benzimidazoles	<u>Albendazole</u>	CYP3A4	CYP1A2, CYP3A5			
		Antifu	ngals			
Imidazoles	<u>Ketoconazole</u>	CYP3A4	UGT1A1, FMO3, CYP26A1		<b>Ø</b>	
	<u>Itraconazole</u>	CYP3A4			<b>Ø</b>	
Triazoles	<u>Voriconazole</u>	CYP2C19	CYP2C9, CYP3A4, CYP3A5			<b>W</b>
	<u>Fluconazole</u>	Renal Excretion				
Allylamines	<u>Terbinafine</u>	CYP2C9	CYP1A2, CYP3A4, CYP2C8, CYP2C19		<b>Ø</b>	_



### **PGx Report -Infectious Disease**

Type: Antiretroviral, Antiviral

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
	<u>Lopinavir</u>	CYP3A4	SLCO1B1, CYP3A5, ABCC1, ABCC2		<b>Ø</b>	
	Ritonavir	CYP3A4	CYP2D6, CYP3A5, ABCC1		<b>Ø</b>	
Protease inhibitor 1st	<u>Saquinavir</u>	CYP3A4	CYP3A5		<b>Ø</b>	
generation	<u>Indinavir</u>	CYP3A4	CYP2D6, CYP3A5, ABCC4		<b>Ø</b>	
	<u>Nelfinavir</u>	CYP2C19	CYP3A4, CYP3A5			<b>W</b>
	<u>Fosamprenavir</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Atazanavir</u>	CYP3A4	CYP3A5, ABCB1			
Protease inhibitor 2nd generation	<u>Darunavir</u>	CYP3A4	CYP3A5, SLCO3A1			
	<u>Tipranavir</u>	CYP3A4	CYP3A5			
NNRTI 1st generation	<u>Delavirdine</u>	CYP3A4	CYP2D6, CYP3A5			
NINTI 1St generation	<u>Efavirenz</u>	CYP2B6	CYP2A6, ABCB1, SLCO3A1, ABCG2			
NNRTI 2nd generation	<u>Nevirapine</u>	CYP3A4	CYP2B6, CYP3A5, ABCB1, SLCO3A1			
	<u>Etravirine</u>	CYP3A4	CYP2C9, CYP2C19, CYP3A5			
	Rilpivirine	CYP3A4	CYP3A5			
Nucleoside reverse	Zidovudine	UGT2B7	Renal Excretion, UGT1A9, SLCO3A1, ABCC1, ABCC4		<b>Ø</b>	
ranscriptase inhibitor (NRTI)	<u>Abacavir</u>	ADH6	UGT1A1, ADK, HLA-B*5701		<b>Ø</b>	
	<u>Zanamivir</u>	Renal Excretion				
Neuraminidase inhibitors/release phase	<u>Peramivir</u>	Renal Excretion				
minore of ordered prides	<u>Oseltamivir</u>	BCHE, ACHE	Renal Excretion			<b>W</b>
CCR5 Co-receptor Antagonist	Maraviroc	CYP3A4	CYP3A5			
	<u>Boceprevir</u>	CYP3A4	IFNL3, CYP3A5		<b>Ø</b>	
Hepatitis C Virus NS3/4A	<u>Telaprevir</u>	CYP3A4	CYP3A5, IFNL3		<b>Ø</b>	
Protease Inhibitor	<u>Paritaprevir</u>	CYP3A4	CYP3A5			
	Simeprevir	CYP3A4	CYP2C8, CYP2C19, CYP3A5, IFNL3			<b>W</b>
	<u>Enfuvirtide</u>	CYP2C19	CYP2E1, CYP1A2			
Other antivirals	Raltegravir	UGT1A1	SLCO1A2		<b>Ø</b>	
Other antivirais	<u>Elvitegravir</u>	CYP3A4	CYP3A5		0	
	Dolutegravir	UGT1A1, CYP3A4	CYP3A5		<b>Ø</b>	

## **PGx Report - Oncology, Hematology**

Type: Antineoplastic I

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Alkylatin	g agents			
Nitrogen mustard analogues	Cyclophosphamide	CYP2B6	CYP2C19, CYP3A4, CYP2C9, CYP3A5, EPHX1, ALDH1A1, ABCC3		<b>Ø</b>	
	<u>Iphosphamide</u>	CYP2B6	CYP3A4, CYP3A5			
Nitrosoureas	<u>Carmustine</u>	CYP1A2	Renal Excretion			<b>W</b>
·		Antimet	abolites			
Folic acid analogues	<u>Methotrexate</u>	Renal Excretion	AOX1, SLCO1B1, SLC19A1, ABCC1, ABCC2, ABCC3, ABCG2			
	<u>Pemetrexed</u>	Renal Excretion	SLC19A1			
	<u>Mercaptopurine</u>	XO	TPMT, NUDT15, AOX1, SLC19A1			<b>W</b>
	<u>Tioguanine</u>	HPRT1	TPMT, NUDT15			<b>W</b>
Purine analogues	<u>Cladribine</u>	DCK	Renal Excretion			
	<u>Clofarabine</u>	DCK	Renal Excretion			
	Nelarabine	ADA	DCK, Renal Excretion, XO			
Pyrimidine analogues	Fluorouracil	DPYD, TYMS, MTHFR	NQO1, GSTP1, UMPS, TYMP, SLC19A1, ABCG2			<b>\(\rightarrow\)</b>
Pyrimidine analogues	Cytarabine	CES1, CES2, CDA	TYMP, DPYD, TYMS, SLCO1B1, SLC29A1		<b>Ø</b>	



## **PGx Report - Oncology, Hematology**

Type: Antineoplastic II

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Plant alkaloids and o	ther natural products			
Vinca alkaloids and	<u>Vincristine</u>	CYP3A4	CYP3A5, ABCC3			
analogues	<u>Vinblastine</u>	CYP3A4	CYP3A5			
Podophyllotoxin derivatives	Etoposide	CYP3A4	CYP3A5, CYP1A2, CYP2E1, ABCB1, UGT1A1			•
	<u>Teniposide</u>	CYP2C19	CYP3A4, CYP3A5, ABCB1			<b>W</b>
Taxanes	<u>Paclitaxel</u>	CYP2C8	CYP3A4, CYP3A5, ABCB1, SLC29A1			
Taxanes	<u>Docetaxel</u>	CYP3A4	CYP3A5, EPHX1, SLCO1B3, ABCC6			
		Cytotoxic antibiotics a	ind related substances			
Anthracyclines and related substances	Doxorubicin	ALDH1A1, ABCB1, GSTP1, NQO1	CYP3A4, CYP2B6, CYP3A5, CYP2C8, CYP2D6, ABCC2, ABCC3		<b>Ø</b>	
		Other antined	plastic agents			
Platinum compounds <u>Cisplatin</u>		Renal Excretion, NQO1, GSTP1	EPHX1, GSTM1, ABCB1, XPC, LRP2, SLC19A1, ABCC2, ABCC3			
Derivative of camptothecin	<u>Irinotecan</u>	UGT1A1, CYP3A4, CES1, CES2	CYP3A5, CYP2B6, UGT1A4, SLC01B1, BCHE, UGT1A9, UGT1A10, SLC19A1, SLC01B3, ABCG2		<b>Ø</b>	

### **PGx Report - Oncology, Hematology**

Type: Antineoplastic Targeted Therapy I

Drug Class Generic Pri		Primary Mechanism Involved		May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
<u>'</u>		Protein kinase in	hibitor (receptor)			
Established to the feet of	<u>Erlotinib</u>	CYP3A4	CYP1A2, CYP3A5			
Epidermal growth factor receptor (EGFR)	<u>Gefitinib</u>	CYP3A4	CYP2D6, CYP3A5, ABCG2			
	<u>Vandetanib</u>	CYP3A4	FMO3, FMO1, CYP3A5			
GFR and epidermal growth factor receptor (HER2)	<u>Lapatinib</u>	CYP3A4, CYP2C19	CYP2C8, CYP3A5, HLA-DQA1*0201, HLA-DRB1*0701		<b>Ø</b>	
iacioi receptor (HEHZ)	<u>Neratinib</u>	CYP3A4	CYP3A5			
C-KIT and PDGFR	<u>Masitinib</u>	CYP3A4	CYP3A5		<b>Ø</b>	
FLT3	Lestaurtinib	CYP3A4	CYP3A5			
RET, VEGFR and EGFR	<u>Vandetanib</u>	CYP3A4	FMO3, FMO1, CYP3A5			
c-MET and VEGFR2	<u>Cabozantinib</u>	CYP3A4	CYP2C8, CYP3A5			
	<u>Axitinib</u>	CYP3A4	CYP1A2, CYP2C19, CYP3A5, UGT1A1		0	
	Nintedanib	CYP1A2	CYP2C9, CYP2C19, CYP2D6, CYP2E1		<b>Ø</b>	
	<u>Pazopanib</u>	CYP3A4, UGT1A1	CYP1A2, CYP2C8, CYP3A5		<b>Ø</b>	
Multiple targets (c-KIT, GFR, PDGFR and VEGFR)	<u>Ponatinib</u>	CYP3A4	CYP2C8, CYP2D6, CYP3A5		<b>Ø</b>	
, , , , , , , , , , , , , , , , , , , ,	<u>Regorafenib</u>	CYP3A4	UGT1A9, CYP3A5		<b>Ø</b>	
	<u>Sorafenib</u>	CYP3A4	UGT1A9, CYP3A5		<b>Ø</b>	
	<u>Sunitinib</u>	CYP3A4	CYP3A5, ABCG2			
	<u>Toceranib</u>	CYP3A4	CYP3A5			
<u> </u>		Protein kinase inhib	` ' '	'		'
	<u>Imatinib</u>	CYP3A4	CYP3A5, ABCB1, SLCO1A2, SLC22A4, ABCG2			
BCR-ABL	<u>Nilotinib</u>	CYP3A4, UGT1A1	CYP1A2, CYP2C9, CYP2C19, CYP2D6, CYP3A5, ABCG2			
	<u>Dasatinib</u>	CYP3A4	CYP3A5, ABCG2		<b>Ø</b>	
	<u>Ponatinib</u>	CYP3A4	CYP2C8, CYP2D6, CYP3A5		<b>Ø</b>	
Src	<u>Bosutinib</u>	CYP3A4	CYP3A5		<b>Ø</b>	
	Lestaurtinib	CYP3A4	CYP3A5		<b>Ø</b>	
Janus kinase	<u>Ruxolitinib</u>	CYP3A4	CYP3A5		<b>Ø</b>	
Janus Kinase	<u>Pacritinib</u>	CYP3A4	CYP3A5		0	
	Tofacitinib	CYP3A4	CYP2C19, CYP3A5			•



### **PGx Report - Oncology, Hematology**

Type: Antineoplastic Targeted Therapy II

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Protein kinase inhil	oitor (non-receptor)			
EML4-ALK	<u>Ceritinib</u>	CYP3A4	CYP2C9, CYP3A5		<b>Ø</b>	
LIVIL4-ALIX	<u>Crizotinib</u>	CYP3A4	CYP3A5			
Bruton tyrosine kinase	<u>Ibrutinib</u>	CYP3A4	CYP2D6, CYP3A5			
BRAF inhibitor (V600E mutation-positive)	<u>Dabrafenib</u>	CYP2C8	CYP3A4, CYP3A5, G6PD		<b>Ø</b>	
<u> </u>		Other Targe	eted therapy	'		
mTOR Inhibitors	Sirolimus	CYP3A4	CYP3A5		<b>Ø</b>	
IIITOIT IIIIIIDIOIS	Everolimus	CYP3A4	CYP2C8, CYP3A5		<b>Ø</b>	
Hedgehog pathway inhibitor	<u>Vismodegib</u>	CYP2C9	CYP3A4, CYP3A5			
		Hormone antagonist	s and related agents	'		'
	<u>Toremifene</u>	CYP3A4	CYP2D6, CYP3A5		<b>Ø</b>	
Selective oestrogen receptor modulators (SERM)	<u>Tamoxifen</u>	CYP3A4, CYP2D6, CYP2C9	CYP3A5, CYP2B6, FMO1, FMO3, CYP2C19, CYP1A2, UGT1A3, UGT1A4, SULT1A1, F2, F5, ABCC2		<b>Ø</b>	
SERD	<u>Fulvestrant</u>	CYP3A4	CYP3A5			
	<u>Flutamide</u>	CYP1A2	CYP3A4, CYP3A5			
Anti andrana	Nilutamide	CYP2C19	FMO3			<b>6.0</b>
Anti-androgens	<u>Bicalutamide</u>	CYP3A4	CYP3A5			
	Enzalutamide	CYP2C8	CYP3A4, CYP3A5		<b>6</b>	
	Anastrozole	CYP3A4	CYP3A5, UGT1A4		0	
Aromatase inhibitors	<u>Letrozole</u>	CYP3A4	CYP2A6, CYP3A5		<u> </u>	
	Exemestane	CYP3A4	CYP3A5		<u> </u>	
Other hormone antagonists and related agents	<u>Abiraterone</u>	CYP3A4	CYP3A5, SULT2A1		0	
		Hema	tologic	'		·
Thrombopoiesis Stimulating Agent	Eltrombopag	CYP1A2	CYP2C8, F5, SERPINC1		<b>Ø</b>	

Abbreviations: C-KIT, tyrosine-protein kinase Kit; PDGFR, Platelet-derived growth factor receptor; FLT3, FMS-like tyrosine kinase-3; RET, RET proto-oncogene; VEGFR, Vascular endothelial growth factor receptor; Src, Proto-oncogene tyrosine-protein kinase Src; EML4-ALK, echinoderm microtubule associated protein like 4 – anaplastic lymphoma kinase; BRAF, proto-oncogene B-Raf; mTOR, mammalian target of rapamycin; SERD, selective oestrogen receptor down-regulator.

### **PGx Report - Organ Transplantation**

Type: Immunosuppressive, Immunomodulation

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Immunosu	uppressive			
Antimetabolite	Mycophenolate mofetil	CYP3A4	CYP3A5, CYP2C8, UGT2B7, UGT1A8, UGT1A9, SLCO1B1, SLCO1B3, ABCC2, HPRT1			
	<u>Azathioprine</u>	XO	TPMT, NUDT15, AOX1			<b>W</b>
	<u>Pimecrolimus</u>	CYP3A4	CYP3A5			
Calcineurin Inhibitors	<u>Tacrolimus</u>	CYP3A4	CYP3A5, ABCB1, UGT2B7		<b>Ø</b>	
	<u>Cyclosporine</u>	CYP3A4	CYP3A5, ABCB1, UGT2B7, ABCC2		<b>Ø</b>	
mTOR Inhibitors	<u>Temsirolimus</u>	CYP3A4	CYP3A5		<b>Ø</b>	
IIITOR IIIIIbilois	<u>Everolimus</u>	CYP3A4	CYP2C8, CYP3A5		<b>Ø</b>	
	Immuno		nodulation			
Immunomodulator and anti- angiogenic	<u>Pomalidomide</u>	CYP1A2	CYP3A4, CYP2C19, CYP2D6, CYP3A5		<b>Ø</b>	



### **PGx Report - Anaesthesiology**

Type: Anaesthetic, Muscle Relaxant

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Inhaled Ar	naesthetics			
		Intravenous age	ents (non-opioid)			
Barbiturates	<u>Hexobarbital</u>	CYP2C19	CYP2C9, CYP2E1, CYP1A2			<b>W</b>
Daibiturates	<u>Thiamylal</u>	CYP2C9			<b>Ø</b>	
	<u>Diazepam</u>	CYP2C19, CYP3A4	CYP3A5, CYP2B6, CYP1A2			<b>W</b>
Benzodiazepines	<u>Lorazepam</u>	UGT2B15	UGT2B7		0	
	<u>Midazolam</u>	CYP3A4	CYP3A5		0	
Other Anaesthetics	<u>Ketamine</u>	CYP3A4	CYP2B6, CYP2C9, CYP3A5			
		Skeletal mus	cle relaxants			
	Carisoprodol	CYP2C19				<b>W</b>
Muscle Relaxants	Cyclobenzaprine	CYP1A2	CYP2D6, CYP3A4, CYP3A5, UGT1A4		<b>Ø</b>	
	<u>Tizanidine</u>	CYP1A2			<b>Ø</b>	

## **PGx Report - Urology**

Type: Drugs Prescribed for the Treatment of Incontinence, Erectile Dysfunction, Benign Prostatic Hypertrophy

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity		
Drugs for urinary frequency and incontinence								
	<u>Oxybutynin</u>	CYP3A4	CYP3A5		<b>Ø</b>			
Anticholinergic	<u>Tolterodine</u>	CYP2D6, CYP3A4	CYP2C9, CYP3A5, CYP2C19					
Anticholinergic	<u>Solifenacin</u>	CYP3A4	CYP3A5					
	<u>Darifenacin</u>	CYP2D6	CYP3A4, CYP3A5					
		Drugs used in ere	ectile dysfunction			1		
	<u>Sildenafil</u>	CYP3A4	CYP2C9, CYP3A5					
	<u>Tadalafil</u>	CYP3A4	CYP3A5					
Phosphodiesterase inhibitors	<u>Vardenafil</u>	CYP3A4	CYP2C9, CYP3A5					
	<u>Avanafil</u>	CYP3A4	CYP3A5					
	<u>Udenafil</u>	CYP3A4	CYP3A5					
'		Drugs used in benign	prostatic hypertrophy					
	<u>Alfuzosin</u>	CYP3A4	CYP3A5, Renal Excretion			<b>W</b>		
Alpha-adrenoreceptor antagonists	<u>Tamsulosin</u>	CYP3A4	CYP2D6, CYP3A5, Renal Excretion			<b>W</b>		
anagomoto	Silodosin	CYP3A4	UGT2B7, CYP3A5					
Testosterone-5-alpha	<u>Finasteride</u>	CYP3A4	CYP3A5					
reductase inhibitors	<u>Dutasteride</u>	CYP3A4	CYP3A5					



### **PGx Report - Endocrinology**

Type: Contraceptives, Androgens, Antiandrogens, Glucocorticoid, Thyroid

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Hormonal co	ontraceptives			
Oestrogens	Ethinylestradiol	CYP3A4, CYP2C9	CYP3A5, CYP2C19, CYP1A2, UGT1A1		<b>Ø</b>	
Oestrogens	<u>Estradiol</u>	CYP1A2	CYP3A4, CYP3A5, CYP2C8, UGT1A1, UGT1A9		<b>Ø</b>	
	<u>Desogestrel</u>	CYP3A4, HSD3B1	CYP3A5, CYP2C9, CYP2C19, UGT1A1		<b>Ø</b>	
Progestogens	<u>Dienogest</u>	CYP3A4	CYP3A5			
	<u>Mestranol</u>	CYP2C9				
F	Levonorgestrel	CYP3A4	CYP3A5		<b>2</b>	
Emergency contraceptives	<u>Ulipristal</u>	CYP3A4	CYP1A2, CYP2D6, CYP3A5		0	
		Andro	ogens			
3-oxoandrosten-(4) derivatives	Testosterone	CYP3A4, CYP19A1	HSD3B2, CYP3A5, UGT2B15, SULTs		<b>Ø</b>	
		Antiano				
Antiandrogens	Cyproterone	CYP3A4	CYP3A5			
		Other sex hormones and mod	• •			
	<u>Raloxifene</u>	UGT1A1	UGT1A8, UGT1A10			
Selective oestrogen receptor modulators (SERMs)	<u>Bazedoxifene</u>	UGT1A1	UGT1A8, UGT1A10			
modulators (SERMS)	<u>Ospemifene</u>	CYP3A4	CYP2C9, CYP3A5, CYP2C19, CYP2B6		<b>Ø</b>	
		Steroid h	normone			
	<u>Dexamethasone</u>	CYP3A4	CYP17A1, CYP3A5			
Glucocorticoids	Cortisol (hydrocortisone)	CYP3A4	CYP3A5		<b>Ø</b>	
	<u>Prednisone</u>	HSD11B2	CYP3A4, CYP3A5, SLC19A1, SULTs, UGTs		<b>Ø</b>	
		Thyroid I	hormone	'		
Thyroid hormones	<u>Levothyroxine</u>	DIO2	UGT1A1, SULTs		<b>Ø</b>	
Thyrola normones	Liothyronine	DIO2	UGT1A1, UGT1A9, SULTs			
	The	re are additional SERMs (Tamoxifen and	Toremifene) described under antineoplas	tics)		

### **PGx Report - Recreational Drugs**

Type: Alcohol, Barbiturates, Benzodiazepines, Cannabinoids, Synthetic Cannabis, Dissociative Drugs, Tobacco

Drug Class	Generic	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
Amphetamines	3,4-methylenedioxy- methamphetamine (MDMA)	Renal Excretion, CYP2D6	CYP1A2, CYP3A4, CYP3A5, FMO3			
	<u>Methamphetamine</u>	CYP2D6, Renal Excretion	DBH, FMO3, ACSM1, GLYAT, DRD3			<b>W</b>
Barbiturates	<u>Amobarbital</u>	CYP3A4	CYP3A5, CYP2B6, CYP2C9, CYP2A6		<b>Ø</b>	
Daibiturates	<u>Phenobarbital</u>	CYP2C19	ABCB1			<b>W</b>
	<u>Alprazolam</u>	CYP3A4	CYP3A5			
Benzodiazepines	Clonazepam	CYP3A4	CYP2C19, CYP3A5, NAT2			<b>W</b>
Delizodiazepines	<u>Lorazepam</u>	UGT2B15	UGT2B7			
	<u>Diazepam</u>	CYP2C19, CYP3A4	CYP3A5, CYP2B6, CYP1A2			<b>W</b>
	Cannabidiol (CBD)	CYP3A4	CYP2C19, CYP3A5			<b>W</b>
Cannabinoids & Related Drugs	Delta 9-tetra hydrocannabinol (△9 THC)	CYP2C9	CYP2C19, CYP3A4, CYP3A5		<b>Ø</b>	
	Cannabinol (CBN)	CYP2C9	CYP2C19, CYP3A4, CYP3A5			
Synthetic Cannabis	<u>JWH-018</u>	CYP1A2	CYP2C9			
Synthetic Garinabis	AM2201	CYP1A2	CYP2C9			
Dissociative Drugs	<u>Ketamine</u>	CYP3A4	CYP2B6, CYP2C9, CYP3A5			
Dissociative Drugs	Phencyclidine (PCP)	CYP3A4	CYP3A5, CYP2A6, CYP1A2		_	<b>&gt;</b>
Ecgonine derivative	<u>Cocaine</u>	BCHE, CES2	CYP3A4, CYP3A5, SLC6A3			<b>W</b>
Ergoline derivatives	Lysergic acid diethylamide (LSD)	CYP3A4	CYP3A5		0	



### **Additional SNPs of Importance for Recreational Drugs**

Gene	Marker	Genotype	Drug	Level of Evidence	Results
OPRD1	rs2236857	T/T	Heroin		Patients may have a lower tendency for heroin addiction
OPRK1	rs702764	T/T	Opioids		Neonates may display reduced abstinence syndrome due to in-utero opioid exposure
OPRK1	rs1051660	C/C	Opioids		Patients may have a lower tendency for Opioids addiction
DBH	rs1611115	C/C	Analgesics	3	Patients with substance withdrawal syndrome may have an increased likelihood of headache when discontinuing the use of analgesics (such as opioids, NSAIDs, triptans, ergot)



#### Clinical Pharmacogenetics Implementation Consortium (CPIC)

#### Medications Affected by Patient Genetic Results

#### Clinical Annotation for CYP2D6\*1, \*1XN, \*2, \*2XN, \*3, \*4, \*5, \*6, \*10, \*17, \*40, \*41

Codeine and Pain Haplotype: \*1/\*1 Evidence Level 1A Efficacy,

Toxicity/ADR

Patients treated with codeine may have 1) increased metabolism/clearance of codeine 2) increased likelihood of response to codeine and 3) decreased but not absent risk for side effects as compared to patients with non-functional or reduced function

-- https://www.pharmgkb.org/clinicalAnnotation/1183616718

#### Clinical Annotation for CYP2D6\*1, \*1XN, \*2, \*2XN, \*3, \*4, \*5, \*6, \*10

**Tramadol and Pain** Haplotype: \*1/\*1 Evidence Level 1B Dosage, Efficacy, Toxicity/ADR

Patients treated with tramadol may have 1) increased metabolism of tramadol 2) increased likelihood of response to tramadol and 3) decreased but not absent risk for side effects as compared to patients with non-functional or reduced function alleles.

-- https://www.pharmgkb.org/clinicalAnnotation/1183618159

#### Clinical Annotation for rs1045642 (ABCB1)

Fentanyl, methadone, morphine, opioids, oxycodone, tramadol and Pain

Evidence Level 2B Dosage

Patients may experience decreased efficacy of fentanyl methadone morphine tramadol oxycodone or other opioids and thus may require an increased dose those drugs as compared to patients with the AA or AG genotype athough this is contradicted in some studies.

-- https://www.pharmgkb.org/clinicalAnnotation/1444704833

#### Clinical Annotation for rs3745274 (CYP2B6)

#### Methadone and Heroin Dependence

Patients treated with methadone for heroin addiction may require a decreased dose of the drug as compared to patients with the GG or GT genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1183702975

#### Clinical Annotation for rs1045642 (ABCB1)

Fentanyl, methadone, morphine, opioids, oxycodone, tramadol and Pain

Genotype: G/G

Genotype: T/T

Haplotype: \*1/\*1

Evidence Level 2B Dosage

Evidence Level 2A Dosage

Patients may experience decreased efficacy of fentanyl, methadone, morphine, tramadol, oxycodone or other opioids and thus may require an increased dose those drugs as compared to patients with the AA or AG genotype, although this is contradicted in some studies.

-- https://www.pharmgkb.org/clinicalAnnotation/1444704833

#### Clinical Annotation for rs1799971 (OPRM1)

Alfentanil, Drugs used in opioid dependence, fentanyl, heroin, morphine, naltrexone, opioids, tramadol, Heroin Dependence, Opioid-Related Disorders, Pain and Postoperative Pain

Genotype: A/A Evidence Level 2B Metabolism/PK

Individuals may experience increased efficacy of opioids for pain and opioid related drugs to treat addiction and may require a decreased dose of opioids as compared to individuals with the AG and GG genotypes. However this has been contradicted in some studies. In some studies the AA and AG genotypes were found to have a increased efficacy and to require a decreased dose as compared to the GG genotype.



#### Clinical Annotation for rs1799971 (OPRM1)

**Naloxone** Genotype: A/A Evidence Level 2B Efficacy

Patients treated with naloxone may have lower cortisol response as compared to patients with the AG or GG genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/655385241

#### Clinical Annotation for rs1799971 (OPRM1)

**Ethanol** Evidence Level 2B Toxicity/ADR Genotype: A/A

Patients may have a decreased but not absent severity of intoxication and a decreased response when exposed to ethanol as compared to patients with the AG and GG genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/981204641

#### Clinical Annotation for rs2231142 (ABCG2)

**Allopurinol and Gout** Genotype: G/G Evidence Level 2B Efficacy

Patients with gout may have improved response when treated with allopurinol as compared to patients with the GT or TT genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1447982582

#### Clinical Annotation for rs762551 (CYP1A2)

#### Leflunomide and Rheumatoid Arthritis

Genotype: C/A

Evidence Level 3 Toxicity/ADR

Patients with rheumatoid arthritis who are treated with leflunomide may have a decreased but not absent risk of toxicity as compared to patients with the CC genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/655384902

#### Clinical Annotation for CYP2C19\*1, \*2, \*3, \*4, \*5, \*6, \*8

Clopidogrel Haplotype: \*1/\*2 Evidence Level 1A Efficacy, Toxicity/ADR

Patients with one functional allele may have 1) poor metabolism of clopidogrel and decreased formation of active drug metabolite resulting in decreased response 2) may have an increased risk for secondary cardiovascular events when treated with clopidogrel as compared to patients with two functional alleles (\*1/\*1).

-- https://www.pharmgkb.org/clinicalAnnotation/1043858794

#### Clinical Annotation for rs4149056 (SLCO1B1)

#### Simvastatin, Muscular Diseases and Central **Core Myopathy**

Genotype: T/T

Evidence Level 1A Toxicity/ADR

Patients may have a lower risk of simvastatin-related myopathy as compared to patients with the CT or CC genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/655384011

#### Clinical Annotation for CYP2C9\*1, \*2, \*3

Warfarin, Cardiovascular Diseases and Heart Haplotype: \*1/\*9 Evidence Level 1A Dosage **Diseases** 

-- https://www.pharmgkb.org/clinicalAnnotation/981238341

#### Clinical Annotation for rs9923231 (VKORC1)

Warfarin Genotype: C/C Evidence Level 1A Dosage

Patients may require an increased dose of warfarin as compared to patients with the CT or TT genotype.

-- https://www.pharmgkb.org/variant/rs9923231?previousQuery=rs9923231



#### Clinical Annotation for rs7294 (VKORC1)

Warfarin Genotype: C/C Evidence Level 1B Dosage

Patients treated with warfarin may require a lower dose as compared to patients with the TC or TT genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/655384733

#### Clinical Annotation for rs1045642 (ABCB1)

Digoxin Genotype: G/G Evidence Level 2A

Patients may have increased metabolism and decreased serum concentration of digoxin as compared to patients with the AA and AG genotypes.

-- https://www.pharmgkb.org/clinicalAnnotation/981204372

#### Clinical Annotation for rs7412 (APOE)

Genotype: C/C

Genotype: T/T

Genotype: T/T

Genotype: C/C

Genotype: G/G

### Atorvastatin, Coronary Disease and Hyperlipidemias

Patients treated with atorvastatin may have a reduced response (less reduction in LDL-cholesterol) as compared to patients with the CT or TT genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1183492249

#### Clinical Annotation for rs4149056 (SLCO1B1)

#### Cerivastatin and Rhabdomyolysis

Patients may have a lower risk of cerivastatin-related rhabdomyolysis as compared to patients with the CC or CT genotype. Cerivastatin was withdrawn from the market because of 52 deaths attributed to drug-related rhabdomyolysis that lead to kidney failure.

-- https://www.pharmgkb.org/clinicalAnnotation/981344897

#### Clinical Annotation for rs4149056 (SLCO1B1)

Pravastatin Genotype: T/T Evidence Level 2A Metabolism/PK

Patients may have decreased plasma concentrations of pravastatin as compared to patients with the CC or CT genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/981345293

#### Clinical Annotation for rs4149056 (SLCO1B1)

#### Rosuvastatin and Hypercholesterolemia

Patients may have lower plasma concentrations of rosuvastatin as compared to patients with the CC genotype. No association is seen between genotypes of this variant and change in LDL-cholesterol levels in response to rosuvastatin treatment.

-- https://www.pharmgkb.org/clinicalAnnotation/981345350

#### Clinical Annotation for rs7294 (VKORC1)

#### Acenocoumarol and phenprocoumon

Patients may require a decreased dose of phenprocoumon or acenocoumarol as compared to patients with the CT or TT genotypes although this has been contradicted in some studies.

-- https://www.pharmgkb.org/clinicalAnnotation/1445585748

#### Clinical Annotation for rs2231142 (ABCG2)

### Rosuvastatin, Hypercholesterolemia and Myocardial Infarction

Patients treated with rosuvastatin 1) may have lower plasma concentrations of rosuvastatin 2) may have a reduced response to treatment as determined by a lower reduction in LDL-C as compared to patients with the TT genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1154221922



Evidence Level 2A Efficacy

Evidence Level 2A Toxicity/ADR

Evidence Level 2A

Evidence Level 2A Dosage

Evidence Level 2B Efficacy

#### Clinical Annotation for rs762551 (CYP1A2)

Clopidogrel Genotype: C/A Evidence Level 3 Efficacy

Patients may have decreased on-treatment platelet reactivity when treated with clopidogrel as compared to patients with the CC genotype. However another study found no association with risk of major adverse cardiac events.

-- https://www.pharmgkb.org/clinicalAnnotation/982030732

#### Clinical Annotation for rs1045642 (ABCB1)

Ondansetron Genotype: G/G Evidence Level 2A Efficacy

Patients may have increased likelihood of nausea and vomiting shortly after being treated with treated with ondansetron as compared to patients with genotype AA.

-- https://www.pharmgkb.org/clinicalAnnotation/1183632195

#### Clinical Annotation for rs10509681 (CYP2C8)

Rosiglitazone Genotype: T/T Evidence Level 2A Dosage

Patients may have decreased metabolism of rosiglitazone a larger change in HbA1c and an increased risk of edema as compared to patients with the CC (CYP2C8\*3/\*3) or CT (CYP2C8\*3/\*1) genotype. One study found no association with blood glucose levels.

-- https://www.pharmgkb.org/clinicalAnnotation/655384653

#### Clinical Annotation for rs762551 (CYP1A2)

#### Deferasirox and beta-Thalassemia

Genotype: C/A Evidence Level 3 Metabolism/PK

Patients with beta-thalassemia may have increased concentrations of deferasirox as compared to patients with the AA genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1444666564

#### Clinical Annotation for rs2470890 (CYP1A2)

#### Deferasirox and beta-Thalassemia

Genotype: C/C

Evidence Level 3 Metabolism/PK

Patients with beta-thalassemia may have increased concentrations of deferasirox as compared to patients with the TT genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1444666571

#### Clinical Annotation for CYP2C19\*1, \*2, \*3

#### Sertraline and Major Depressive Disorder

Haplotype: \*1/\*2

Evidence Level 1A Metabolism/PK

Patients treated with sertraline may have increased clearance as compared to patients with the \*2/\*2 or \*2/\*3 diplotype and a decreased clearance as compared to patients with the \*1/\*1 diplotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1183619004

#### Clinical Annotation for CYP2C19\*1, \*17, \*2, \*3, \*4

Citalopram, escitalopram and Major Depressive Disorder

Haplotype: \*1/\*2

Haplotype: \*1/\*1

Evidence Level 1A Efficacy, Toxicity/ADR

Patients treated with citalopram or escitalopram may have a decreased drug clearance/metabolism and decreased tolerance as compared to patients with the CYP2C19\*1/\*1 genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1183620386

#### Clinical Annotation for CYP2D6\*1, \*1XN, \*2, \*2XN, \*3, \*4, \*5, \*6, \*10

Paroxetine, Major Depressive Disorder, Mental Disorders and Obsessive-Compulsive Disorder

Evidence Level 1A Efficacy, Toxicity/ADR, Metabolism/PK

Patients treated with paroxetine may have 1) a decreased clearance of paroxetine as compared to patients with more than 2 functional CYP2D6 alleles (\*1XN \*2XN) and 2) an increased clearance of paroxetine as compared to patients with two non-functional CYP2D6 alleles (\*3 \*4 \*5 \*6) or \*10/\*10 genotype.



#### Clinical Annotation for CYP2D6\*1, \*1XN, \*2, \*2XN, \*3, \*4, \*5, \*6, \*10

#### Nortriptyline and Major Depressive Disorder

Haplotype: \*1/\*1

Haplotype: \*1/\*1

Evidence Level 1A Efficacy, Toxicity/ADR, Metabolism/PK

Patients treated with nortriptyline may have decreased metabolism of nortriptyline as compared to patients with a duplication of a functional CYP2D6 gene and 1) increased metabolism of nortriptyline (decreased nortriptyline plasma levels) 2) a decreased but not absent risk for side effects as compared to patients with CYP2D6 non-functional alleles (\*3 \*4 \*5 \*6) or reduced function alleles (\*10).

-- https://www.pharmgkb.org/clinicalAnnotation/1183618725

#### Clinical Annotation for CYP2D6\*1, \*3, \*4, \*5, \*6, \*10

Fluvoxamine, Depressive Disorder, Major Depressive Disorder, Mental Disorders and Obsessive-Compulsive Disorder Evidence Level 1A Efficacy, Toxicity/ADR, Metabolism/PK

Patients treated with fluvoxamine may have 1) decreased steady-state plasma concentration-to-dose (C/D) ratio as compared to patients with the \*1/\*5 \*1/\*10 \*5/\*10 \*10/\*10 genotype 2) decreased plasma concentrations 3) decreased risk of developing gastrointestinal side effects as compared to patients with the \*5/\*10 \*10/\*10 genotype and 4) decreased AUC Cmax and half-life time of fluvoxamine as compared to patients with two non-functional CYP2D6 alleles (poor metaboliser phenotypes). However contradictory findings are reported.

-- https://www.pharmgkb.org/clinicalAnnotation/1183700410

#### Clinical Annotation for CYP2D6\*1, \*1XN, \*2, \*2XN, \*3, \*4, \*5, \*6, \*10, \*41

Amitriptyline, Major Depressive Disorder, Mental Disorders and Mood Disorders Evidence Level 1A Efficacy, Toxicity/ADR

Patients treated with amitriptyline may have 1) decreased nortriptyline plasma levels and increased clearance of amitryptyline and 2) a decreased but not absent risk for side effects as compared to patients with CYP2D6 non-functional alleles (\*3 \*4 \*5 \*6) or reduced function alleles and a decreased metabolism of amitriptyline as compared to patients with duplication of a functional CYP2D6 gene.

-- https://www.pharmgkb.org/clinicalAnnotation/1183617651

#### Clinical Annotation for rs1800497 (ANKK1)

#### **Bupropion and Tobacco Use Disorder**

Genotype: G/A

Haplotype: \*1/\*1

Evidence Level 1B Efficacy

Patients treated with bupropion may be less likely to quit smoking as compared to patients with the GG genotype although this has been contradicted in one study.

-- https://www.pharmgkb.org/clinicalAnnotation/655385422

#### Clinical Annotation for rs1800497 (ANKK1, DRD2)

Antipsychotics, clozapine, olanzapine, risperidone, Hyperprolactinemia, tardive dyskinesia and Weight gain

**Genotype:** *G/A* Evidence Level 2B Toxicity/ADR

Patients may have increased risk of side effects including hyperprolactinaemia and weight gain but decreased risk of tardive dyskinesia during treatment with antipsychotic drugs as compared to patients with the GG genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/655385495

#### Clinical Annotation for rs3813929 (HTR2C)

#### **Antipsychotics and Mental Disorders**

Genotype: C/C

Evidence Level 2B Toxicity/ADR

Patients with psychiatric disorders who are treated with antipsychotics may have an increased risk of weight gain as compared to patients with the TT genotype. However some studies find no association with weight gain.



#### Clinical Annotation for rs1954787 (GRIK4)

#### Antidepressants, Depression, Depressive **Disorder and Major Depressive Disorder**

Genotype: T/T Evidence Level 2B Efficacy

Evidence Level 3 Toxicity/ADR

Evidence Level 3 Toxicity/ADR

Evidence Level 3 Dosage, Toxicity/ADR

Patients with Depressive Disorder or Depression may be less likely to respond to antidepressant treatment as compared to patients with the CC genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/982030381

#### Clinical Annotation for rs762551 (CYP1A2)

**Olanzapine** Genotype: C/A Evidence Level 3 Efficacy

Patients with psychiatric disorders who are treated with olanzapine may have an increased response to olanzapine based on not decreased mean dose-/body weight-normalised olanzapine serum concentrations as compared to patients with the AA genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/655385604

#### Clinical Annotation for rs762551 (CYP1A2)

Genotype: C/A

Genotype: G/A

Genotype: C/A

Genotype: C/C

Antipsychotics, chlorpromazine, fluphenazine, thioridazine, trifluoperazine and Schizophrenia

Patients may have increased QT interval when treated with antipsychotics chlorpromazine fluphenazine thioridazine and trifluoperazine in people with Schizophrenia as compared to patients with genotype AA.

-- https://www.pharmgkb.org/clinicalAnnotation/1183679775

#### Clinical Annotation for CYP1A2\*1A, \*1F

Clozapine and Schizophrenia Haplotype: \*1B/\*1L Evidence Level 3 Toxicity/ADR

-- https://www.pharmgkb.org/clinicalAnnotation/1444608250

#### Clinical Annotation for rs2069514 (CYP1A2)

#### **Antipsychotics and Schizophrenia**

Schizophrenia patients may have increased severity of tardive dyskinesia when treated with antipsychotics in people who were smokers as compared to patients with the GG genotype. Genotype AG is not associated with increased QT interval in Schizophrenia patients treated with antipsychotics as compared to genotype GG.

-- https://www.pharmgkb.org/clinicalAnnotation/981201888

#### Clinical Annotation for rs762551 (CYP1A2)

#### **Paroxetine and Major Depressive Disorder**

Patients may require an increased dose of paroxetine and may have an increased risk of fatigue when treated with paroxetine as compared to patients with the CC genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/982031767

#### Clinical Annotation for rs2470890 (CYP1A2)

#### **Paroxetine and Major Depressive Disorder**

Patients with major depressive disorder who are treated with paroxetine may be less likely to experience remission as compared to patients with the TT genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1183617623

#### Clinical Annotation for rs2069526 (CYP1A2)

#### **Escitalopram and Major Depressive Disorder**

Genotype: T/T Evidence Level 3 Toxicity/ADR

Patients with major depressive disorder may have reduced metabolism of escitalopram as compared to patients with the GG genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1183622557



Evidence Level 3 Efficacy

#### Clinical Annotation for rs1902023 (UGT2B15)

Lorazepam and oxazepam Genotype: A/C Evidence Level 2B

Subjects with the AC genotype may have decreased clearance of oxazepam or lorazepam as compared to subjects with the CC genotype or increased clearance as compared to subjects with the AA genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/655387798

#### Clinical Annotation for rs762551 (CYP1A2)

#### Carbamazepine and Epilepsy

Genotype: C/A

Evidence Level 3 Metabolism/PK

Paediatric patients with epilepsy may have decreased clearance of carbamazepine as compared to paediatric patients with epilepsy and the AA genotypes.

-- https://www.pharmgkb.org/clinicalAnnotation/1447983940

#### Clinical Annotation for rs28365063 (UGT2B7)

#### Carbamazepine and Epilepsy

Genotype: A/A

Evidence Level 3 Metabolism/PK

Patients with epilepsy may have decreased clearance of carbamazepine as compared to patients with the AG or GG genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1447987066

#### Clinical Annotation for CYP2C19\*1, \*17, \*2, \*3

#### **Voriconazole and Mycoses**

Haplotype: \*1/\*9

Evidence Level 1B Metabolism/PK

-- https://www.pharmgkb.org/clinicalAnnotation/1183689217

#### Clinical Annotation for rs1045642 (ABCB1)

#### **Nevirapine and HIV Infections**

Haplotype: \*1/\*1

Evidence Level 2A Toxicity/ADR

Patients with HIV-1 infection who are treated with nevirapine may have an increased risk for nevirapine hepatotoxicity as compared to patients with the AA genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/655386244

#### Clinical Annotation for rs3745274 (CYP2B6)

#### **Nevirapine and HIV Infections**

Genotype: T/T

Evidence Level 2A

Patients with HIV infection may have decreased clearance of and increased exposure to nevirapine as compared to patients with the GG genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/981202294

#### Clinical Annotation for rs28399499 (CYP2B6)

Nevirapine and HIV Genotype: T/T

Evidence Level 2A

Patients may have decreased plasma drug exposure when treated with nevirapine as compared to patients with the CT or CC genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/981201854

#### Clinical Annotation for rs2279343 (CYP2B6)

**Efavirenz and HIV** Genotype: A/A Evidence Level 2A Metabolism/PK

Patients with HIV may have increased clearance and decreased plasma concentration of efavirenz as compared to patients with the GG genotype.



#### Clinical Annotation for rs28399499 (CYP2B6)

Efavirenz and HIV Genotype: T/T Evidence Level 2A Metabolism/PK

Patients may have decreased plasma drug exposure when treated with efavirenz as compared to patients with the CT or CC genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/981201844

#### Clinical Annotation for TPMT\*1, \*2, \*3A, \*3B, \*3C, \*4

### Azathioprine, mercaptopurine, purine analogues and thioguanine

**Haplotype:** \*1/\*3C Evidence Level 1A Dosage, Toxicity/ADR

Patients treated with thiopurine drugs and purine analogues: 1) may have decreased inactivation of thiopurines due to deficient TPMT activity and 2) may have an increased risk for toxicity when receiving thiopurine drugs and purine analogues as compared to patients with the \*1/\*1 genotype. These effects may be more pronounced in those who are homozygous for two non-functional variants (e.g. \*2/\*3A) than in those who are heterozygous for the non-functional variant (\*1/\*2).

-- https://www.pharmgkb.org/clinicalAnnotation/1184648909

#### Clinical Annotation for rs1801133 (MTHFR)

Carboplatin and Non-Small-Cell Lung Carcinoma

Genotype: G/G

Haplotype: \*1/\*1

Evidence Level 2A Efficacy

Patients may have 1) decreased likelihood of response 2) decreased progression free survival to carboplatin in people with Non-Small-Cell Lung Carcinoma as compared to patients with genotype AA.

-- https://www.pharmgkb.org/clinicalAnnotation/981220481

#### Clinical Annotation for rs1045642 (ABCB1)

Methotrexate, Burkitt Lymphoma, Drug Toxicity, T-Cel Lymphomal, Precursor Cell Lymphoblastic Leukemia-Lymphoma and Toxic liver disease Evidence Level 2A Toxicity/ADR

Patients with lymphoma or leukemia who are treated with methotrexate may have lower concentrations of the drug and may have a reduced but not absent risk of toxicity as compared to patients with the AA genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/1296599132

#### Clinical Annotation for rs1801133 (MTHFR)

#### **Methotrexate and Neoplasms**

Genotype: G/G

Evidence Level 2A Dosage, Efficacy,

Toxicity/ADR

Patients with leukemia or lymphoma who are treated with methotrexate: 1) may have better response to treatment 2) may be at decreased risk of toxicity 3) may require a higher dose of methotrexate and 4) may be at lower risk of folate deficiency as compared to patients with the AA or AG genotype. This association has been contradicted or not found in multiple studies.

 $-- \underline{https://www.pharmgkb.org/clinicalAnnotation/827848365}$ 

#### Clinical Annotation for rs1801133 (MTHFR)

#### Cyclophosphamide

Genotype: G/G

Evidence Level 2A Toxicity/ADR

Patients may have decreased likelihood of Drug Toxicity when treated with cyclophosphamide as compared to patients with the AA genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/981204929

#### Clinical Annotation for rs4148323 (UGT1A1)

SN-38 and Neoplasms Genotype: G/G Evidence Level 2A

Cancer patients may have increased metabolism of SN-38 when treated with irinotecan as compared to patients with the AA genotype. SN-38 is the active metabolite of irinotecan and is glucuronidated by UGT1A1. One in vitro study found increased enzyme activity for the G allele compared to the A allele.



#### Clinical Annotation for rs4148323 (UGT1A1)

Irinotecan and Neoplasms Genotype: G/G Evidence Level 2A

Cancer patients treated with irinotecan-based regimens may have a decreased risk of neutropenia as compared to patients with the AA genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/981201713

#### Clinical Annotation for rs776746 (CYP3A5)

Tacrolimus, heart transplantation, hemopoietic stem cell transplant, Kidney Transplantation and lung transplantation

Evidence Level 1A Dosage, Metabolism/PK

Patients who are recipients of a kidney heart lung or hematopoeitic stem cell transplant or have other diseases who are treated with tacrolimus may have increased metabolism of tacrolimus resulting in decreased exposure and may require a higher dose as compared to patients with the CC genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/981203719

#### Clinical Annotation for rs2740574 (CYP3A4)

#### **Tacrolimus and Organ Transplantation**

Genotype: C/T

Genotype: C/T

Genotype: C/T

Evidence Level 2A Dosage

Transplant recipients may require an increased dose of tacrolimus as compared to patients with the TT (\*1/\*1) genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/655387058

#### Clinical Annotation for rs776746 (CYP3A5)

Tacrolimus and liver transplantation

Evidence Level 2A Dosage,

Metabolism/PK

Patients who are recipients of a liver transplantation from a donor with the CT (CYP3A5 \*1/\*3) genotype may have increased metabolism of tacrolimus resulting in decreased exposure and may require a higher dose as compared to patients who receive a liver transplantation from a donor with the CC (\*3/\*3) genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/982046323

#### Clinical Annotation for rs776746 (CYP3A5)

#### Tacrolimus and transplant rejection

Genotype: C/T

Evidence Level 2A Efficacy

Patients with the CT genotype (\*1/\*3) and recipients of kidney or hematopoietic stem cell transplant who are treated with tacrolimus may have an increased risk of transplant rejection as compared to patients with the CC genotype (\*3/\*3) or a decreased risk of transplant rejection as compared to patients with the TT genotype (\*1/\*1).

-- https://www.pharmgkb.org/clinicalAnnotation/981203808

#### Clinical Annotation for rs776746 (CYP3A5)

#### Sirolimus and Transplantation

Genotype: C/T

Evidence Level 2A Dosage

Patients who are recipients of transplants may have increased metabolism of sirolimus and require a higher dose as compared to patients with the CC genotype (\*3/\*3).

-- https://www.pharmgkb.org/clinicalAnnotation/981203936

#### Clinical Annotation for rs6025 (F5)

### Hormonal contraceptives for systemic use and Thrombosis

Genotype: C/C

Evidence Level 2A Toxicity/ADR

Patients may have a decreased risk of experiencing thrombosis when receiving oral contraceptives as compared to patients with the CT or TT genotype (carriers of Factor V Leiden). Both Factor V Leiden and oral contraceptives have been found to independently increase the risk for thrombosis but together they may have a cumulative effect on thrombosis risk.



#### Clinical Annotation for rs1799963 (F2)

Hormonal contraceptives for systemic use, Stroke and Venous Thrombosis

**Genotype:** *G/G* Evidence Level 3 Toxicity/ADR

Patients not taking oral contraceptives (OCs) may have a decreased risk for deep vein thrombosis (DVT) as compared to patients with the AG genotype who are taking oral contraceptives. Current evidence suggests that patients with the AG mutation who are taking oral contraceptives experience an increase risk for DVT due to the cumulative effect of both the contraceptives and the AG genotype. At the time of writing there are no studies that show a significant increase in risk for DVT when considering only the AG genotype. Additionally some contradictory evidence exists for this association.

-- https://www.pharmgkb.org/clinicalAnnotation/1444672766

#### Clinical Annotation for rs4680 (COMT)

**Nicotine and Tobacco Use Disorder** 

Genotype: G/G Evidence Level 2A Efficacy

Patients treated with nicotine replacement therapy may have a decreased likelihood of smoking cessation and increased risk of relapse as compared to patients with the AA genotype. However some contradictory evidence exists.

-- https://www.pharmgkb.org/clinicalAnnotation/981202618

#### Clinical Annotation for rs1800497 (ANKK1)

Ethanol and Alcoholism Genotype: G/A Evidence Level 2B Toxicity/ADR

Patients may have an increased risk for Alcoholism when exposed to ethanol as compared to patients with the GG genotype.

-- https://www.pharmgkb.org/clinicalAnnotation/981204890

#### Clinical Annotation for rs762551 (CYP1A2)

**Caffeine and Myocardial Infarction** 

Genotype: C/A Evidence Level 3 Toxicity/ADR

Patients may have an increased risk of nonfatal myocardial infarction with increased coffee consumption as compared to patients with the AA genotype.



#### **Genomic Test Results**

#### Genotype/Haplotype Details

CYP1A2

Allele Tested: \*1A, \*1B, \*1C, \*1D, \*1E, \*1F, \*1G, \*1J, \*1K, \*1L, \*3, \*4, \*5, \*6, \*7, \*8, \*11, \*15, \*16.

Genetic results: CYP1A2 \*1B/\*1L
Phenotype: Extensive metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
CYP1A2	Asn516Asn	1548T>C	*1B	rs2470890	C/C
CYP1A2		-3860G>A	*1C	rs2069514	G/A
CYP1A2		-2467delT	*1D	rs35694136	T/-
CYP1A2		-739T>G	*1E	rs2069526	T/T
CYP1A2		-729C>T	*1K	rs12720461	C/C
CYP1A2		-163C>A	*1F	rs762551	C/A
CYP1A2	Asp348Asn	1042G>A	*3	rs56276455	G/G
CYP1A2	lle386Phe	1156A>T	*4	rs72547516	A/A
CYP1A2	Cys406Tyr	1217G>A	*5	rs55889066	G/G
CYP1A2	Arg431Trp	1291C>T	*6	rs28399424	C/C
CYP1A2	Splicing defect	1253+1G>A	*7	rs56107638	G/G
CYP1A2	Arg456His	1367G>A	*8	rs72547517	G/G
CYP1A2	Phe186Leu	558C>A	*11	rs72547513	C/C
CYP1A2	Pro42Arg	125C>G	*15	rs72547511	C/C
CYP1A2	Arg377Gln	1130G>A	*16	rs72547515	G/G

CYP1A2 is the most important gene in the metabolism of: Asenapine, Bromazepam, Carmustine, Clozapine, Cyamemazine, Cyclobenzaprine, Eltrombopag, Estradiol, Febuxostat, Flutamide, Frovatriptan, Imipramine, Leflunomide, Lidocaine, Loxapine, Mirtazapine, Nabumetone, Naratriptan, Nintedanib, Palonosetron, Pomalidomide, Promazine, Pyrazinamide, Ramelteon, Rasagiline, Riluzole, Ropinirole, Tacrine, Theophylline, Thiothixene, Tizanidine, Triamterene, Trifluoperazine, Zileuton, Zolmitriptan.

Drugs and substances known to induce CYP1A2 activity include: beta-naphthoflavone, char-grilled meat, Marijuana, Modafinil, Omeprazole, Tobacco.

Drugs and substances known to inhibit CYP1A2 activity include: Amiodarone, Efavirenz, Fluoroquinolones, Fluvoxamine, Ticlopidine, Verapamil.

CYP1A2 activity is dependent upon hepatic and renal function status as well as age.

#### Genotype/Haplotype Details

CYP2B6

Allele Tested: \*1, \*4, \*5, \*6, \*7, \*9, \*10, \*11, \*18, \*16, \*22, \*28.

Genetic results: CYP2B6 \*9/\*9
Phenotype: Extensive metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
CYP2B6	Lys262Arg	785A>G	*4	rs2279343	A/A
CYP2B6	Arg487Cys	1459C>T	*5/*7	rs3211371	C/C
CYP2B6	Gln172His	516G>T	*6/*9	rs3745274	T/T
CYP2B6	Arg22Cys	64C>T	*10	rs8192709	C/C
CYP2B6	Met46Leu	136A>G	*11	rs35303484	A/A
CYP2B6	lle328Thr	983T>C	*16	rs28399499	T/T
CYP2B6		-82T>C	*22	rs34223104	T/T
CYP2B6	Arg378Ter	1132C>T	*28	rs34097093	C/C

CYP2B6 is the most important gene in the metabolism of: Bupropion, Cyclophosphamide, Efavirenz, Iphosphamide, Meperidine, Ondansetron, Selegiline, Sertraline.

Drugs and substances known to induce CYP2B6 activity include: Artemisinin, Carbamazepine, Efavirenz, Nevirapine, Phenobarbital, Phenytoin, Rifampicin.

Drugs and substances known to inhibit CYP2B6 activity include: Clopidogrel, Orphenadrine, Thiotepa, Ticlopidine, Voriconazole.

#### Genotype/Haplotype Details

CYP2C8

Allele Tested: \*1, \*2, \*3, .

Genetic results: CYP2C8 \*1/\*1

Phenotype: Extensive metaboliser



Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
CYP2C8	lle269Phe	805A>T	*2	rs11572103	A/A
CYP2C8	Lvs399Ara	1196A>G	*3	rs10509681	T/T

CYP2C8 is the most important gene in the metabolism of: Amodiaquine, Chloroquine, Dabrafenib, Desloratadine, Enzalutamide, Isotretinoin, Nicardipine, Paclitaxel, Pioglitazone, Repaglinide, Rosiglitazone.

Drugs and substances known to induce CYP2C8 activity include: Rifampicin.

Drugs and substances known to inhibit CYP2C8 activity include: Gemfibrozil, Montelukast, Trimethoprim.

#### Genotype/Haplotype Details

CYP2C9

Allele Tested: \*1, \*2, \*3, \*4, \*5, \*6, \*7, \*8, \*9, \*10, \*11, \*13, \*15, \*16, \*27.

Genetic results: CYP2C9 \*1/\*9
Phenotype: Extensive metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
CYP2C9	Arg144Cys	430C>T	*2	rs1799853	C/C
CYP2C9	Ile359Leu	1075A>C	*3	rs1057910	A/A
CYP2C9	lle359Asn	1076T>C	*4	rs56165452	T/T
CYP2C9	Asp360Glu	1080C>G	*5	rs28371686	C/C
CYP2C9	Lys273Argfs	817delA	*6	rs9332131	A/A
CYP2C9	Leu19lle	55C>A	*7	rs67807361	C/C
CYP2C9	Arg150His/Leu	449G>A/T	*8/*27	rs7900194	G/G
CYP2C9	His251Arg	752A>G	*9	rs2256871	A/G
CYP2C9	Glu272Gly	815A>G	*10	rs9332130	A/A
CYP2C9	Arg335Trp	1003C>T	*11	rs28371685	C/C
CYP2C9	Leu90Pro	269T>C	*13	rs72558187	T/T
CYP2C9	Ser162Ter	485C>A	*15	rs72558190	C/C
CYP2C9	Thr299Ala	895A>G	*16	rs72558192	A/A

CYP2C9 is the most important gene in the metabolism of: Acenocoumarol, Alosetron, Azilsartan, Bosentan, Cannabinol (CBN), Celecoxib, Chloramphenicol, Delta 9-tetra hydrocannabinol (\$\Delta\$9\_THC), Dronabinol, Fenoprofen, Flurbiprofen, Fluvastatin, Gliclazide, Glimepiride, Gliquidone, Ibuprofen, Indomethacin, Irbesartan, Ketobemidone, Lacosamide, Lornoxicam, Losartan, Mefenamic acid, Meloxicam, Mestranol, Naproxen, Nateglinide, Paramethadione, Parecoxib, Phenprocoumon, Piroxicam, Primidone, Sulfinpyrazone, Tapentadol, Tenoxicam, Terbinafine, Thiamylal, Tolazamide, Tolazamide, Torasemide, Trimethadione, Valsartan, Vismodegib, Warfarin, Zafirlukast.

Drugs and substances known to induce CYP2C9 activity include: Carbamazepine, Nevirapine, Phenobarbital, Rifampicin, Secobarbital.

Drugs and substances known to inhibit CYP2C9 activity include: Amentoflavone, Amiodarone, Apigenin, Isoniazid, Fluconazole, Miconazole, Sulfaphenazole, Valproic acid.

#### Genotype/Haplotype Details

CYP2C19

Allele Tested: \*1, \*2, \*2B, \*4, \*5, \*6, \*7, \*8, \*9, \*10, \*17.

Genetic results: CYP2C19 \*1/\*2 Phenotype: Intermediate metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
CYP2C19	Splicing defect	681G>A	*2	rs4244285	G/A
CYP2C19	Glu92Asp	276G>C	*2B	rs17878459	G/G
CYP2C19	Met1Val	1A>G	*4	rs28399504	A/A
CYP2C19	Arg433Trp	1297C>T	*5	rs56337013	C/C
CYP2C19	Arg132Gln	395G>A	*6	rs72552267	G/G
CYP2C19	Splicing defect	819+2T>A	*7	rs72558186	T/T
CYP2C19	Trp120Arg	358T>C	*8	rs41291556	T/T
CYP2C19	Arg144His	431G>A	*9	rs17884712	G/G
CYP2C19	Pro227Leu	680C>T	*10	rs6413438	C/C
CYP2C19		-806C>T	*17	rs12248560	C/C

CYP2C19 is the most important gene in the metabolism of: Brivaracetam, Carisoprodol, Citalopram, Clobazam, Clopidogrel, Dexlansoprazole, Diazepam, Enfuvirtide, Esomeprazole, Flunitrazepam, Hexobarbital, Mephenytoin, Moclobemide, Nelfinavir, Nilutamide, Omeprazole, Pantoprazole, Pentamidine, Phenobarbital, Phenytoin, Proguanil, Rabeprazole, Temazepam, Teniposide, Voriconazole.

Drugs and substances known to induce CYP2C19 activity include: Artemisinin, Carbamazepine, Efavirenz, Norethisterone, Rifampicin, Ritonavir, St. John's Wort.

Drugs and substances known to inhibit CYP2C19 activity include: Chloramphenicol, Esomeprazole, Felbamate, Fluvoxamine, Isoniazid, Lansoprazole, Moclobemide, Omeprazole.



#### Genotype/Haplotype Details

#### CYP2D6

Allele Tested: \*1, \*2A, \*2D, \*3, \*4A, \*4B, \*4K, \*4M, \*4N, \*5, \*6A, \*6C, \*7, \*8, \*9, \*10, \*11, \*12, \*14A, \*14B, \*15, \*17, \*20, \*29, \*34, \*35, \*36, \*39, \*41, \*69, and CNVs.

Genetic results: CYP2D6 \*1/\*1
Phenotype: Extensive metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
CYP2D6	Arg296Cys	886C>T	*2	rs16947	G/G
CYP2D6	Ser486Thr	1457G>C	*2	rs1135840	G/G
CYP2D6	Val136Val	408G>C	*2A	rs1058164	G/G
CYP2D6	Arg259Glyfs	775delA	*3	rs35742686	T/T
CYP2D6	Splicing defect	506-1G>A	*4	rs3892097	C/C
CYP2D6	CNV assay		*5/XN	CYP2D6_CNVs	**
CYP2D6	Trp152Glyfs	454delT	*6	rs5030655	A/A
CYP2D6	His324Pro	971A>C	*7	rs5030867	T/T
CYP2D6	Gly169Ter/Arg	505G>T/A	*8/*14	rs5030865	G/G
CYP2D6	Lys281del	841_843delAAG	*9	rs5030656	T/T
CYP2D6	Pro34Ser	100C>T	*10	rs1065852	G/G
CYP2D6	Splicing defect	181-1G>C	*11	rs201377835	G/G
CYP2D6	Gly42Arg	124G>A	*12	rs5030862	C/C
CYP2D6	46fs	137-138insT	*15	rs774671100	-/-
CYP2D6	Thr107lle	320C>T	*17	rs28371706	G/G
CYP2D6	211fs	1973_1974insG	*20	rs72549354	-/-
CYP2D6	Val136Met	406G>A	*29/*70	rs61736512	C/C
CYP2D6	Val11Met	31G>A	*35	rs769258	C/C
CYP2D6	(sing-dup)		*36	CYP2D7/2D6 hybrid *36	WT/WT
CYP2D6	Splicing defect	985+39G>A	*41	rs28371725	C/C

CYP2D6 is the most important gene in the metabolism of: Aclidinium, Amitriptyline, Amoxapine, Arformoterol, Aripiprazole, Atomoxetine, Bisoprolol, Carvedilol, Chlorpromazine, Clomipramine, Clonidine, Codeine, Cyclizine, Dapoxetine, Darifenacin, Debrisoquine, Desipramine, Dexmethylphenidate, Dextromethorphan, Diphenhydramine, Donepezil, Dosulepin, Doxazosin, Doxepin, Duloxetine, Encainide, Ethylmorphine, Flecainide, Fluoxetine, Fluoxetine, Fluoxetine, Fluoxetine, Fluoxetine, Fluoxetine, Fluoxetine, Methylphenidate, Metoprolol, Mexiletine, Mianserin, Modafinil, Nebivolol, Nefazodone, Nortriptyline, Paliperidone, Paroxetine, Perphenazine, Primaquine, Procainamide, Prochlorperazine, Promethazine, Propafenone, Propranolol, Protriptyline, Reserpine, Risperidone, Sertindole, Sparteine, Tetrabenazine, Thioridazine, Timolol, Tolterodine, Tramadol, Trimipramine, Umeclidinium, Venlafaxine, Vicoprofen, Vortioxetine, Zuclopenthixol.

In Caucasians, approximately 6 -10% are CYP2D6 poor metabolisers and up to 7% are ultrarapid drug metabolisers.

Drugs and substances known to induce CYP2D6 activity include: Dexamethasone, Glutethimide, Rifampicin.

Drugs and substances known to inhibit CYP2D6 activity include: Bupropion, Fluoxetine, Paroxetine, Quinidine, Ritonavir.

#### Genotype/Haplotype Details

CYP3A4

Allele Tested: \*1A, \*1B, \*2, \*3, \*6, \*12, \*13, \*15, \*17, \*20, \*22.

Genetic results: CYP3A4 \*1A/\*1B

Phenotype: Extensive metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
CYP3A4		-392A>G	*1B	rs2740574	C/T
CYP3A4	Ser222Pro	664T>C	*2	rs55785340	A/A
CYP3A4	Met445Thr	1334T>C	*3	rs4986910	A/A
CYP3A4	Asp277Glufs	830_831insA	*6	rs4646438	-/-
CYP3A4	Leu373Phe	1117C>T	*12	rs12721629	G/G
CYP3A4	Pro416Leu	1247C>T	*13	rs4986909	G/G
CYP3A4	Arg162Gln	485G>A	*15	rs4986907	C/C
CYP3A4	Phe189Ser	566T>C	*17	rs4987161	A/A
CYP3A4	Lys487_Pro488delinsLysThrArgfs	1461_1462insA	*20	rs67666821	-/-
CYP3A4		522-191C>T	*22	rs35599367	G/G

#### Genotype/Haplotype Details

CYP3A5

Allele Tested: \*1A, \*1D, \*2, \*3A, \*3B, \*3K, \*3L, \*6, \*7, \*8, \*9.

Genetic results: CYP3A5 \*3A/\*6
Phenotype: Poor metaboliser



Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
CYP3A5	Thr398Asn	1193C>A	*2	rs28365083	G/G
CYP3A5	Splicing defect	689-1A>G	*3	rs776746	C/T
CYP3A5	His30Tyr	58C>T	*3B	rs28383468	G/G
CYP3A5	Leu32Thrfs	92_93insG	*3B	rs200579169	-/-
CYP3A5		*14C>T	*1D/*3	rs15524	A/G
CYP3A5	Splicing defect	624G>A	*6	rs10264272	C/T
CYP3A5	Thr346Tyrfs	1035_1036insT	*7	rs41303343	-/-
CYP3A5	Arg28Cys	82C>T	*8	rs55817950	G/G
CYP3A5	Ala337Thr	1009G>A	*9	rs28383479	C/C
CYP3A5	Phe446Ser	1337T>C	*3K	rs41279854	A/A

CYP3A4/5 are the most important genes in the metabolism of: Abiraterone, Albendazole, Alfentanyl, Alfuzosin, Aliskiren, Almotriptan, Alprazolam, Amiodarone, Amlodipine, Amobarbital, Anastrozole, Apixaban, Aprepitant, Armodafinil, Arteether, Artemether, Artemisinin, Astemizole, Atazanavir, Atorvastatin, Avanafil, Axitinib, Bedaquiline, Bepridil, Bicalutamide, Boceprevir, Bosutinib, Bromocriptine, Bromperidol, Brotizolam, Budesonide, Buprenorphine, Buspirone, Cabozantinib, Cannabidiol (CBD), Carbamazepine, Certinib, Cerivastatin, Chlordiazepoxide, Chlorpheniramine, Cilansetron, Cilostazol, Cinacalcet, Cinitapride, Cisapride, Clarithromycin, Clebopride, Clindamycin, Clonazepam, Clorazepame, Cortisol (hydrocortisone), Crizotinib, Cyclosporine, Cyproterone, Darunavir, Dasatinib, Delavirdine, Desogestrel, Dexamethasone, Dextropropoxyphene, Dienogest, Dihydrocodeine, Dihydrocorgotamine, Diltiazem, Disopyramide, Docetaxel, Dolasetron, Domperidone, Dronedarone, Droperidol, Dutasteride, Eletriptan, Elvitegravir, Eplerenone, Ergotamine, Erlotinib, Erythromycin, Escitalopram, Estazolam, Eszopiclone, Ethinylestradiol, Ethosuximide, Etoposide, Etoricoxib, Etravirine, Everolimus, Exemestane, Felbamate, Fentanyl, Finasteride, Flurazepam, Fluticasone, Fosamprenavir, Fulvestrant, Gefitinib, Gemfibrozil, Glyburide, Granisetron, Halofantrine, Halofantrine, Halofantrine, Halofantrine, Levacetylmethadol, Levomepromazine, Levomilnacipran, Levonorgestrel, Loperamide, Lopinavir, Loratadine, Lormetazepam, Lovastatin, Lurasidone, Lysergic acid diethylamide (LSD), Macitentan, Maraviroc, Masitinib, Mefloquine, Methadone, Midazolam, Mifepristone, Mometasone, Montelukast, Mosapride, Mycophenolate mofetil, Neratinib, Nevirapine, Nifedipine, Nifedipine, Nifedipine, Nifedipine, Panalukast, Prednisone, Quazepam, Quetapam, Quetapame, Quinidine, Quinine, Ranolazine, Reboxetine, Regorafenib, Rifampicin, Rifampicin, Rifonavir, Rivaroxaban, Roflumilast, Ruxolitinib, Salmeterol, Savagliptin, Scopolamine, Sibutramine, Sildenafil, Silodosin, Simeprevir,

Drugs and substances known to induce CYP3A4/5 activity include: Carbamazepine, Efavirenz, Nevirapine, Phenobarbital, Phenytoin, Pioglitazone, Rifabutin, Rifampicin, St. John's Wort, Troglitazone.

Drugs and substances known to inhibit CYP3A4/5 activity include: Chloramphenicol, Clarithromycin, Grapefruit juice flavonoids, Indinavir, Itraconazole, Ketoconazole, Nefazodone, Nelfinavir, Ritonavir.

#### Genotype/Haplotype Details

VKORC1

Allele Tested: H1, H2, H3, H4, H6, H7.

Genetic results: VKORC1 H3/H4

Phenotype: Sensitive to Warfarin

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
VKORC1		497T>G	H2	rs2884737	A/A
VKORC1		2255T>C	H3	rs2359612	A/G
VKORC1		1542C>G	H3	rs8050894	C/G
VKORC1		1173T>C	H4	rs9934438	G/G
VKORC1		-1639A>G	H4	rs9923231	C/C
VKORC1		3730G>A	H7	rs7294	C/C

The VKORC1 gene encodes the vitamin K epoxide reductase enzyme, the drug target of Warfarin.

#### Genotype/Haplotype Details

ABCB1

Allele Tested: \*1, \*2.

Genetic results: ABCB1 \*1/\*1
Phenotype: Extensive function

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
ABCB1	lle1145lle	3435C>T	*2	rs1045642	G/G

ABCB1 is an important pharmacokinetic gene modifying drug disposition. Pharmaceutical agents affected include: Alfentanyl, Aliskiren, Atazanavir, Atorvastatin, Carbamazepine, Cisplatin, Clopidogrel, Cyclosporine, Digoxin, Doxorubicin, Efavirenz, Etoposide, Fentanyl, Imatinib, Labetalol, Methadone, Morphine, Nevirapine, Nortriptyline, Ondansetron, Oxycodone, Paclitaxel, Phenobarbital, Phenytoin, Pitavastatin, Risperidone, Simvastatin, Tacrolimus, Verapamil.



#### Genotype/Haplotype Details

ABCG2

Allele Tested: \*1, \*141K.

Genetic results: ABCG2 \*1/\*1

Phenotype: Extensive function

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
ABCG2	Gln141Lys	421C>A	*141K	rs2231142	G/G

ABCG2 is an important pharmacokinetic gene affecting drug disposition. Pharmaceutical agents affected include: Paracetamol, Atorvastatin, Docetaxel, Doxorubicin, Erlotinib, Fluoropyrimidines, Gefitinib, Imatinib, Irinotecan, Lovastatin, Lamivudine, Methotrexate, Pazopanib, Paclitaxel, Pravastatin, Uricosurics, Zidovudine.

#### Genotype/Haplotype Details

**TPMT** 

Allele Tested: \*1, \*2, \*3A, \*3B, \*3C, \*4.

Genetic results: TPMT \*1/\*3C

Phenotype:Intermediate metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
TPMT	Ala80Pro	238G>C	*2	rs1800462	G/G
TPMT	Ala154Thr	460G>A	*3A or *3B	rs1800460	C/C
TPMT	Tyr240Cys	719A>G	*3A or *3C	rs1142345	T/C
TPMT	Splicing defect	626-1G>A	*4	rs1800584	C/C

TPMT contribute in the metabolism of several drugs including: Azathioprine, Mercaptopurine, Thioguanine.

#### Genotype/Haplotype Details

UGT1A1

Allele Tested: \*1, \*6, \*28, \*36, \*37.

Genetic results: UGT1A1 \*1/\*1

Phenotype: Extensive metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
UGT1A1	Gly71Arg	211G>A	*6	rs4148323	G/G

A(TA)(7,5,8)TAA

UGT1A1 is the most important gene in the metabolism of: Bazedoxifene, Ezetimibe, Irinotecan, Raloxifene, Raltegravir, Rosuvastatin.

UGT1A1 contribute in the metabolism of several drugs including: Abacavir, Paracetamol, Arformoterol, Atorvastatin, Axitinib, Buprenorphine, Carvedilol, Desogestrel, Dolutegravir, Ethinylestradiol, Estradiol, Etoposide, Febuxostat, Fluvastatin, Gemfibrozil, Indacaterol, Ketoconazole, Labetalol, Levothyroxine, Liothyronine, Losartan, Lovastatin, Morphine, Naltrexone, Nilotinib, Pazopanib, Simvastatin, Telmisartan.

#### Genotype/Haplotype Details

UGT2B7

Allele Tested: \*1a, \*1d.

UGT1A1

Genetic results: UGT2B7 \*1a/\*1a

Phenotype: Extensive metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
UGT2B7	Arg124Arg	372A>G	*1d	rs28365063	A/A

UGT2B7 is the most important gene in the metabolism of: Clofibrate, Diclofenac, Hydromorphone, Morphine, Lorazepam-r, Naloxone, Naltrexone, Oxazepam-r, Oxymorphone, Zidovudine.



6/6

rs8175347

\*28, \*36, \*37

#### Genotype/Haplotype Details

UGT2B15

Allele Tested: \*1, \*2.

Genetic results: UGT2B15 \*1/\*2
Phenotype: Extensive metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
UGT2B15	Tyr85Asp	253G>T	*2	rs1902023	A/C

UGT2B15 is the most important gene in the metabolism of: Lorazepam-s, Oxazepam-s.

#### Genotype/Haplotype Details

DPYD

Allele Tested: \*1, \*2A, \*9A, \*9B, \*10. Genetic results: DPYD \*9A/\*9A Phenotype: Extensive metaboliser

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
DPYD		1905+1G>A	*2A	rs3918290	G/G
DPYD	Cys29Arg	85T>C	*9A/*9B	rs1801265	G/G
DPYD	Arg886His	2612C>T	*9B	rs1801267	C/C
DPYD	Val995Phe	2983G>T	*10	rs1801268	C/C

DPYD is the most important gene in the metabolism of: Cytarabine, Fluorouracil, Tegafur.

#### Genotype/Haplotype Details

OPRM1

Allele Tested: \*1, \*2.

Genetic results: OPRM1 \*1/\*1
Phenotype: Sensitive to Opioids

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
OPRM1	Asn40Asp	118A>G	*2	rs1799971	A/A

#### Genotype/Haplotype Details

APOE

Allele Tested: \*3, \*2, \*4, \*1.

Genetic results: APOE

Gene	Protein change	Nucleotide change	Allele	Marker	Genotype
APOE	Arg176Cys	526C>T	*2	rs7412	C/C
APOE	Cys130Arg	388T>C	*4	rs429358	T/T



Testing purpose(s): 1) To provide information on how tested individual's genetic profile may affect carrier status for: a) certain inherited disease, b) reaction to certain drugs, c) risk of certain common health conditions, and/or d) response to selected diet, exercise, and/or nutrition recommendations. 2) To obtain information on tested individual's ancient ancestry. Testing purposes are dependent upon specific genetic testing ordered by patient's healthcare professional. Based on testing results, patients should make no changes to medical care [including, but not limited to, changes in dosage or frequency of medication, diet and/or exercise regimens, or pregnancy planning] without the advice of and consultation with a healthcare professional.

Genetic testing is an evolving science. Current testing protocols and results are based on the current/existing developments, information and testing techniques known at this time.

In the future, new variants may be identified and/or more research may be developed on the significance of currently identified variants that will drive changes in the interpretation of previously obtained genetic testing results. Current testing may not include identification of certain variants associated with: diet, exercise or nutrition; disease; and/or, drug response due to these issues.

Factors such as age, diet, ethnicity, family health history, and/or personal health, not related to genetics can also impact the likelihood of developing certain conditions or exhibiting certain drug reactions. Therefore, patients may not always exhibit and/or require the specific diet, nutrition and/or exercise, disease, or drug response expected or consistent with his/her genetic test results.

The genetic associations of certain conditions, particularly those related to diet and exercise, have only been observed/studied in Caucasian populations only. This limitation means that interpretations and recommendations are made in the context of Caucasian-only studies and results may or may not be relevant to those tested who are non-Caucasian or mixed ethnicity individuals.

Healthcare professionals may recommend additional testing to be performed by an independent laboratory or consult with an outside, independent genetic counselor or healthcare professional.

### **Examples of different levels of evidence for PGx SNPs**

Gene	Marker	Level of Evidence	Drugs
TPMT	rs1142345	1A	Azathioprine, Mercaptopurine, Thioguanine
CYP2D6	rs16947	1A	Amitriptyline, Codeine, Nortriptyline, Paroxetine
VKORC1	rs9923231	1A	Warfarin
SLCO1B1	rs4149056	1A	Simvastatin
CYP2D6	rs16947	1B	Tramadol
VKORC1	rs9923231	1B	Acenocoumarol
CYP2D6	rs16947	2A	Flecainide, Doxepin, Desipramine, Atomoxetine, Risperidone, Clomipramine, Imipramine, Venlafaxine
SLCO1B1	rs4149056	2A	Cerivastatin, Pravastatin, Rosuvastatin
ABCB1	rs1045642	2A	Digoxin, Nevirapine, Methotrexate
CYP2D6	rs16947	3	Timolol, Carvedilol, Haloperidol, Aripiprazole, Metoprolol, Citalopram, Escitalopram, Tamoxifen
VKORC1	rs9923231	3	Phenprocoumon
SLCO1B1	rs4149056	3	Repaglinide, Irinotecan, Mycophenolate mofetil, Atorvastatin, Methotrexate, Olmesartan
ABCB1	rs1045642	3	Paclitaxel, Phenytoin, Fluorouracil, Dicloxacillin, Capecitabine, Nortriptyline, Oxaliplatin, Verapamil, Fexofenadine, Atorvastatin, Simvastatin, Sirolimus, Talinolol, Tamoxifen, Morphine, Efavirenz, Vincristine, Imatinib, Olanzapine, Risperidone, Cyclosporine, Tacrolimus, Atazanavir, Phenobarbital, Codeine, Clopidogrel, Etoposide, Oxaliplatin
CYP2D6	rs16947	4	Methylphenidate, Bufuralol
SLCO1B1	rs4149056	4	Lopinavir, Atrasentan
ABCB1	rs1045642	4	Carbamazepine

Level 1A Annotation for a variant-drug combination in a CPIC or medical society-endorsed PGx guideline, or implemented at a PGRN site or in another major health system.

Level 1B Annotation for a variant-drug combination where the preponderance of evidence shows an association. The association must be replicated in more than one cohort with significant p-values, and preferably will have a strong effect size.

Level 2A Annotation for a variant-drug combination that qualifies for level 2A where the variant is within a VIP (Very Important Pharmacogene) as defined by PharmGKB. The variants in level 2A are in known pharmacogenes, so functional significance is more likely.

Level 2B Annotation for a variant-drug combination with moderate evidence of an association. The association must be replicated but there may be some studies that do not show statistical significance, and/or the effect size may be small.

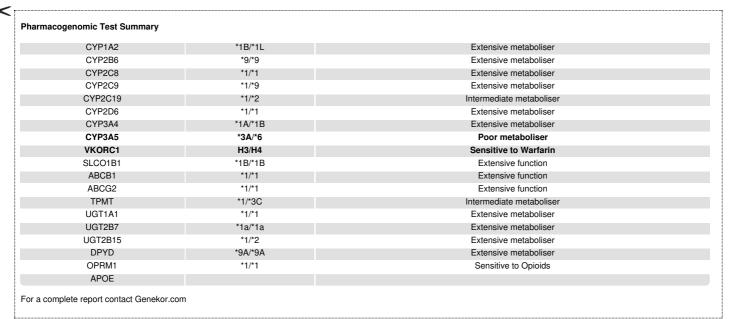
Level 3 Annotation for a variant-drug combination based on a single significant (not yet replicated) or annotation for a variant-drug combination evaluated in multiple studies but lacking clear evidence of an association.

Level 4 Annotation based on a case report, non-significant study or in vitro, molecular or functional assay evidence only.



#### **Patient Information Card**

An easily portable summary of the report patients can share with their medical professionals. (Please cut along dotted line.)



#### AbG8MwDiN9V8Gcp2R-oJ











