

GRI Labs

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This report combines (i) an analysis of the patient's DNA by GRI, identifying relevant genetic variants that are informative for medication efficacy, safety, and dosing, with (ii) an interpretation of the identified DNA variants by GRI to bring you immediately actionable clinical guidance regarding safer, more effective medications and dosages for the patient. The Medication Report section lists the type of PGx guidance present on FDA-approved drug labels. Medications with no established FDA PGx guidance are provided solely for educational purposes.

Patient: Doe, Jane Date of Birth: Jan 01, 1990 Gender: Female	Physician: Dr. Exar Practice: Example		Specimen Sample ID:	type: Buccal swab ex225
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Medication Report Details (by class) References	Pg. 15 Pg. 39	ANKK1	G G	Normal function
Patient Information Card Genetic Summary Information	Pg. 40	АроЕ	ε3 ε3	See ApoE Genotype Info
† When multiple activities are listed, check in Medication Report Details (Pg. 15) for spo medication of interest.	ecific	ATM	C C	Increased likelihood of treatment success when taking metformin
Uncertain = No known diplotype/result (name for this combination of genetic variants; Unint	. ,	COMT(Val158Met)	G G	Normal function
Genotype.		CYP1A2	*1A *1A	Normal metabolizer
		CYP2B6	*1A *1A	Normal metabolizer

**CYP2C19** 

CYP2C8

CYP2C9

CYP2D6

CYP3A4

\*1|\*1

\*1|\*1

\*1|\*1

\*1Ax2|\*1A

\*1B|\*1B

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Normal metabolizer

Normal metabolizer

Normal metabolizer

Ultrarapid metabolizer

Ultrarapid metabolizer

Gene	Result	Activity †
CYP3A5	*1D *1D;	Normal metabolizer
	or *1A *1A;	
	or *1A *1D	
CYP4F2	*1 *1	Normal metabolizer
Factor V Leiden	Variant	See Thrombosis Profile
GRIK4	T T	Altered function
HLA-B*1502	WT WT	Negative
HTR2A	A A	Normal function
HTR2C(-759C>T)	C/C	Unknown Metabolizer
IFNL3	C C	Normal function
MTHFR	Uncertain Allele	See Thrombosis Profile
MTHFR (A1298C)	Uncertain Allele	See Thrombosis Profile
MTHFR (C677T)	Uncertain Allele	See Thrombosis Profile
OPRM1(A118G)	A A	Normal function
Prothrombin (F2)	Normal	See Thrombosis Profile
SLCO1B1	*1 *1	Normal liver uptake activity
TPMT	*1 *1	Normal metabolizer
VKORC1	*1 *1	Normal (with respect to Warfarin)

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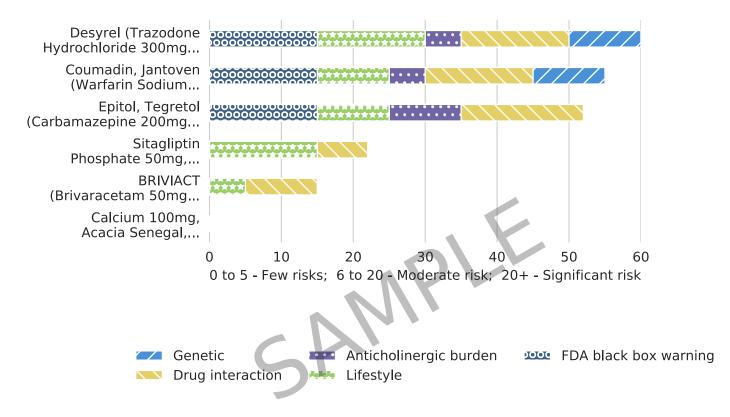
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### **Current Regimen Risk Chart**

This chart summarizes the various risk factors associated with each medication entered into GeneDose<sup>™</sup> Live for Jane Doe. The length of each colored segment represents the relative contribution of a risk category (detailed in the below legend) to the overall risk associated with the use of a medication. For further information, consult the *Current Regimen Risk Details* Pg. 4 section.



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### **Current Regimen Risk Detail**



### **Severe Risks**

Black box warning for Coumadin, Jantoven (Warfarin Sodium 10mg Oral tablet) and bleeding

Black box warning for Desyrel (Trazodone Hydrochloride 300mg Oral tablet) and suicidal ideation

Black box warning for Epitol, Tegretol (Carbamazepine 200mg Oral tablet) and rash

Black box warning for Epitol, Tegretol (Carbamazepine 200mg Oral tablet) and aplastic anemia

Black box warning for Epitol, Tegretol (Carbamazepine 200mg Oral tablet) and agranulocytosis

### Strong regimen anticholinergic burden

The cumulative effect of taking multiple medicines with anticholinergic properties termed as anticholinergic burden can adversely impact cognition, physical function and increase the risk of mortality.

### Major Risks

### Sitagliptin Phosphate 50mg, Simvastatin 40mg Oral tablet with Grapefruit Juice

• avoid combination unless benefit outweighs potential risk

Coadministration with grapefruit juice increases the peak serum concentrations and the AUC of lovastatin and may have a similar effect on simvastatin. Grapefruit juice should be avoided or minimized in patients taking simvastatin.

### Desyrel (Trazodone Hydrochloride 300mg Oral tablet) with Grapefruit Juice

• avoid combination unless benefit outweighs potential risk

Advise patients to avoid grapefruit juice while taking trazodone due to increased trazodone exposure and associated adverse effects including QT prolongation.

Genetic warning for Coumadin, Jantoven (Warfarin Sodium 10mg Oral tablet)

Individuals with this combination of alleles may benefit from an increased dose of Warfarin. The FDA

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table recommends a therapeutic dose of 5-7 mg/day.

### Genetic warning for Desyrel (Trazodone Hydrochloride 300mg Oral tablet)

Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose, or using an alternative medication.

### Moderate Risks

### Coumadin, Jantoven (Warfarin Sodium 10mg Oral tablet) with Grapefruit Juice

- monitor for evidence of bleeding
- monitor international normalized ratio (INR) or prothrombin time

Closely monitor the INR if coadministration of warfarin with grapefruit/grapefruit juice is necessary as concurrent use may increase the exposure of warfarin leading to increased bleeding risk.

### Coumadin, Jantoven (Warfarin Sodium 10mg Oral tablet) with Alcohol

- monitor for evidence of bleeding
- monitor international normalized ratio (INR) or prothrombin time

In studies of healthy subjects anticoagulated with warfarin, small to moderate amounts of wine do not alter the INR. Acute intoxication from ethanol may enhance the hypoprothrombinemic response to warfarin due to inhibition of warfarin's metabolism.

### Sitagliptin Phosphate 50mg, Simvastatin 40mg Oral tablet with Alcohol

- monitor blood glucose concentration and for changes in glycemic control
- warn against driving or operating machinery or performing other hazardous tasks until drug effects are known

Patients should be advised to limit alcohol (ethanol) ingestion when treated with an antidiabetic agent. Ethanol inhibits gluconeogenesis, which can increase the risk for hypoglycemia. In some patients, hypoglycemia can be prolonged.

### Desyrel (Trazodone Hydrochloride 300mg Oral tablet) with Alcohol

- avoid combination unless benefit outweighs potential risk
- warn against driving or operating machinery or performing other hazardous tasks until drug effects are known
- monitor for an increase in CNS depression

Trazodone may enhance the response to alcohol and other CNS depressants. Trazodone can have

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significant sedative effects. The patient should be cautioned accordingly.

### Epitol, Tegretol (Carbamazepine 200mg Oral tablet) with Grapefruit Juice

• consult literature

Grapefruit juice has been shown to increase carbamazepine peak and trough serum concentrations and the AUC by up to 40 percent. Grapefruit juice contains a compound that inhibits CYP3A4 in enterocytes. Increased sedation or other side effects may occur.

### Epitol, Tegretol (Carbamazepine 200mg Oral tablet) with Alcohol

- warn against driving or operating machinery or performing other hazardous tasks until drug effects are known
- monitor for an increase in CNS depression

Because of its primary CNS effect, caution should be used when carbamazepine is taken with other centrally acting drugs such as ethanol.

### BRIVIACT (Brivaracetam 50mg Oral tablet) with Alcohol

• monitor for an increase in CNS depression

The effects of ethanol on psychomotor function, alertness, attention span, body sway, saccadic reaction time, and memory may be increased during co-administration with brivaracetam.

### Desyrel (Trazodone Hydrochloride 300mg Oral tablet) may reduce effect of BRIVIACT (Brivaracetam 50mg Oral tablet)

- monitor for altered clinical response to drug therapy
- dosage increase may be required

Trazodone can lower the seizure threshold of anticonvulsants. Patients may require increased concentrations of anticonvulsants to achieve equivalent effects if trazodone is added. Drowsiness may be additive.

BRIVIACT (Brivaracetam 50mg Oral tablet) may have its effect decreased by, and may increase effect of Epitol, Tegretol (Carbamazepine 200mg Oral tablet)

- monitor for signs of drug toxicity
- monitor for altered clinical response to drug therapy
- warn against driving or operating machinery or performing other hazardous tasks until drug effects are known
- dosage reduction may be required

Coadministration with carbamazepine may increase exposure to the active metabolite of carbamazepine, carbamazepine-epoxide. A 26% decrease in the plasma concentration of brivaracetam

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has also been observed during co-administration.

Coumadin, Jantoven (Warfarin Sodium 10mg Oral tablet) may have their effect increased by Desyrel (Trazodone Hydrochloride 300mg Oral tablet)

- use combination with caution
- monitor patient clinically
- monitor for evidence of bleeding
- monitor international normalized ratio (INR) or prothrombin time

Patients should be instructed to monitor for signs and symptoms of bleeding while taking trazodone concurrently with anticoagulants and to promptly report any bleeding events to the practitioner. There is an increased risk of bleeding.

# Epitol, Tegretol (Carbamazepine 200mg Oral tablet) may decrease concentration of Desyrel (Trazodone Hydrochloride 300mg Oral tablet)

- monitor for altered clinical response to drug therapy
- dosage increase may be required

Consider increasing the trazodone dose based on therapeutic response when coadministered with carbamazepine. Concurrent use may decrease trazodone exposure.

Sitagliptin Phosphate 50mg, Simvastatin 40mg Oral tablet may increase effect of Coumadin, Jantoven (Warfarin Sodium 10mg Oral tablet)

- use combination with caution
- monitor for evidence of bleeding
- monitor international normalized ratio (INR) or prothrombin time

Simvastatin can potentiate the anticoagulant effects of warfarin. Simvastatin should be prescribed cautiously in patients receiving warfarin, with appropriate INR monitoring.

Epitol, Tegretol (Carbamazepine 200mg Oral tablet) may reduce effect of Coumadin, Jantoven (Warfarin Sodium 10mg Oral tablet)

- monitor for altered clinical response to drug therapy
- monitor international normalized ratio (INR) or prothrombin time

Closely monitor the INR if coadministration of warfarin with carbamazepine is necessary as concurrent use may decrease the exposure of warfarin leading to reduced efficacy.

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### **Minor Risks**

Sitagliptin Phosphate 50mg, Simvastatin 40mg Oral tablet may have its concentration decreased by Epitol, Tegretol (Carbamazepine 200mg Oral tablet)

• monitor for altered clinical response to drug therapy

Carbamazepine, which is a CYP3A4 inducer, may decrease the efficacy of HMG-Co-A reductase inhibitors which are CYP3A4 substrates, including simvastatin.

SANRIE

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### **Thrombosis Profile**

Tested Genes (Alleles)	Genotype	Predicted Phenotype	Clinical Guidance
Prothrombin (F2)	Normal	Variant alleles detected. It is important for	Individuals homozygous for the Factor V Leiden mutation have an approximately
Factor V Leiden	Homozygous variant	individuals possessing this allelic variant to	80-fold increased risk of venous thrombosis as compared to individuals without the
MTHFR (A1298C)	Homozygous variant	understand the clinical risks and the genetic implications of their	mutation. Patients who are homozygous for either MTHFR variant may have a further increased risk for venous thrombosis if they
MTHFR (C677T)	Homozygous variant	result. Patients should be counseled by their physician or genetic counselor	also possess the Factor V Leiden 1691A allele.

### **General Description**

Genetic analyses of three genes (four alleles) considered to increase the risk for venous thrombosis were performed using molecular genetic techniques. The presence of the Prothrombin (Factor 2) gene allele 20210A and Factor V Leiden allele 1691A are risk factors for venous thrombosis. This risk may be further increased by the use of estrogen therapy, oral contraceptives, pregnancy, and surgery.

Patients who are homozygous for MTHFR 677T or MTHFR 1298C may have a further increased risk for venous thrombosis if they also possess the Factor V Leiden 1691A allele. However the MTHFR alleles alone do not predict a significant risk for venous thrombosis.

#### **References and Useful Information:**

- Factor V Leiden Working Group; ACMG Laboratory Quality Assurance Molecular Subcommittee of the ACMG Laboratory Quality Assurance Committee AMERICAN COLLEGE OF MEDICAL GENETICS; Standards and Guidelines for Clinical Genetics Laboratories; 2006 Edition
  - Middeldorp S, Henkens CM, Koopman MM, van Pampus ECM, Hamulyák K, van der Meer J, Prins MH, Büller HR. The incidence of venous thromboembolism in family members of patients with factor V Leiden mutation and venous thrombosis. Ann Intern Med 1998;128:15-20.
  - Vandenbroucke JP, Koster T, Briet E, Reitsma PH, Bertina RM, Rosendaal FR. Increased risk of venous thrombosis in oral contraceptive users who are carriers of factor V Leiden mutation. Lancet 1994;344:1453-1457.
  - Rosendaal FR, Koster T, Vandenbroucke JP, Reitsma PH. High risk of thrombosis in patients homozygous for factor V Leiden (activated protein C resistance). Blood 1995;85(6):1504-1508.
  - Reich LM, Bower M, Key NS. Role of the geneticist in testing and counseling for inherited thrombophilia. Genet Med 2003;5:133-143.
  - Tosetto A, Rodeghiero F, Martinelli I, De Stefano V, Missiaglia E, Chiusolo P, Mannucci PM. Additional genetic risk factors for venous thromboembolism in carriers of the factor V Leiden mutation. Br J Haematol 1998;103:871-876.
  - De Stefano V, Martinelli I, Mannucci PM, Paciaroni K, Chiusolo P, Casorelli I, Rossi E, Leone G. The risk of recurrent deep venous thrombosis among heterozygous carriers of both factor V Leiden and the G20210A prothrombin mutation. N Engl J Med 1999;341:801-806.
- M. Adams, P.D. Smith, D. Martin, J.R. Thompson, D. Lodwick, N.J. Samani. Genetic analysis of thermolabile methylenetetrahydrofolate reductase as a risk factor for myocardial infarction. QJM. 1996 Jun;89(6):437-44.

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### **ApoE Genotype Information**<sup>†</sup>

Tested Genes (Alleles)	Genotype	Predicted Phenotype	Clinical Guidance
ΑροΕ (ε2, ε3, ε4)	ε3 ε3	Often associated with normal lipid metabolism.	Typical cardiovascular disease risk expected.

### **General Description**

Genetic analysis in the ApoE gene was performed using molecular genetic techniques. The genotype is based on genotyping results for this patient at SNPs rs429358 and rs7412.

ApoE  $\epsilon$ 3 is the most common allele—found in about 60% of people. The presence of  $\epsilon$ 2 or  $\epsilon$ 4 alleles may be a risk factor for multiple conditions including cardiovascular disease. ApoE  $\epsilon$ 2 carriers may be more likely to develop familial dysbetalipoproteinemia or type III hyperlipoproteinemia.

† Predicted phenotype, clinical significance, relative risk, and interpretations reported for each genotype are associated with cardiovascular risk only. The interpretations should not be used to determine the relative risk of other diseases. Other factors important to understanding total risk should be considered.

Medication Summary					
Cardiac	5				
Therapeutic Class	Standard Precautions	A Caution / Info	Change recommended		
Antiarrhythmics		Flecainide Propafenone			
Anticoagulants	Acenocoumarol Warfarin (CYP4F2)	Warfarin			
Anticonvulsants	Phenytoin				
Antiplatelet Agents	Clopidogrel Prasugrel	Ticagrelor			
Beta Blockers	Nebivolol Propranolol	Carvedilol Metoprolol Timolol			
Statins	Simvastatin	Atorvastatin			

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Gastroenterology			
Therapeutic Class	Standard Precautions	A Caution / Info	Change recommended
Antidepressants	Mirtazapine	Trazodone	Amitriptyline Clomipramine Desipramine Doxepin Nortriptyline
Antiemetics			Ondansetron Tropisetron
Endocrine-Metabolic Agents			Eliglustat
Immunosuppressants	Azathioprine Mercaptopurine Thioguanine	Cyclosporine	
Nonsteroidal Anti- Inflamatory Drugs (NSAIDs)	Celecoxib		
Proton Pump Inhibitors (PPIs)	Esomeprazole Rabeprazole	Dexlansoprazole Lansoprazole Omeprazole Pantoprazole	
Selective Serotonin Reuptake Inhibitors (SSRIs)	Citalopram Escitalopram		Paroxetine
Infectious Disease			
Therapeutic Class	Standard Precautions	Caution / Info	Change recommended
Antifungals	Voriconazole		Ketoconazole
Pain	-		_
Therapeutic Class	Standard Precautions	A Caution / Info	Change recommended
Analgesics, Opioid	Methadone (CYP2B6)		
Anticonvulsants	Brivaracetam Carbamazepine Clobazam Oxcarbazepine Phenytoin		
Antidepressants	Mirtazapine	Duloxetine	Amitriptyline

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Pain			
Therapeutic Class	Standard Precautions	A Caution / Info	Change recommended
	Moclobemide	Trazodone Venlafaxine Vortioxetine	Clomipramine Desipramine Doxepin Nortriptyline Protriptyline
Antipsychotics	Olanzapine		
Beta Blockers	Nebivolol Propranolol	Timolol	
Endocrine-Metabolic Agents			Eliglustat
Immunosuppressants	Azathioprine Mercaptopurine	Cyclosporine Tacrolimus	
Muscle Relaxants	Carisoprodol		
Nonsteroidal Anti- Inflamatory Drugs (NSAIDs)	Celecoxib Diclofenac Flurbiprofen Ibuprofen Lornoxicam Meloxicam Piroxicam	RLE	
Opioids	Alfentanil Fentanyl (OPRM1) Hydromorphone Morphine Oxycodone (CYP3A5) Sufentanil	Buprenorphine Fentanyl Hydrocodone	Codeine Oxycodone Tramadol
Selective Serotonin Reuptake Inhibitors (SSRIs)	Citalopram Escitalopram Fluoxetine Sertraline	Fluvoxamine	Paroxetine
Psychotropic			
Therapeutic Class	Standard Precautions	A Caution / Info	Change recommended
Anti-ADHD Agents	Amphetamine Dexmethylphenidate Dextroamphetamine Lisdexamfetamine Methylphenidate (COMT)	Atomoxetine Clonidine Methylphenidate (ADRA2A)	

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Psychotropic			
Therapeutic Class	Standard Precautions	A Caution / Info	Change recommended
	Carbamazepine Clobazam Oxcarbazepine Phenytoin		
Antidementia Agents		Donepezil	
Antidepressants	Bupropion Mirtazapine Moclobemide	Duloxetine Trazodone Venlafaxine Vortioxetine	Amitriptyline Clomipramine Desipramine Doxepin Nortriptyline Protriptyline
Antipsychotics	Aripiprazole Flupenthixol Olanzapine	Brexpiprazole Clozapine Haloperidol Iloperidone Perphenazine Pimozide Quetiapine Risperidone Thioridazine Zuclopenthixol	
Anxiolytics	Diazepam	Alprazolam Buspirone Clonazepam	
Beta Blockers	Propranolol		
Central Monoamine- Depleting Agents		Tetrabenazine	
Central Nervous System Agents		Dextromethorphan- Quinidine	
Cholinesterase Inhibitors		Galantamine	
Hypnotics		Eszopiclone	
Selective Serotonin Reuptake Inhibitors (SSRIs)	Citalopram Escitalopram Fluoxetine Sertraline	Fluvoxamine	Paroxetine

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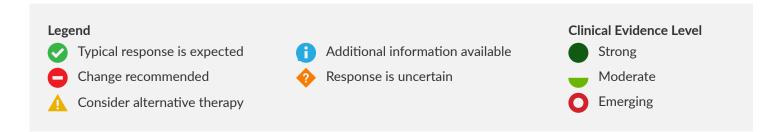


Surgery			
Therapeutic Class	Standard Precautions	A i Caution / Info	Change recommended
Anticholinergic Agents		Tolterodine	
Antiemetics			Ondansetron Tropisetron
Opioids	Fentanyl (OPRM1)	Fentanyl	
Other Drugs			
Therapeutic Class	Standard Precautions	A Caution / Info	Change recommended
Alpha-1 Blockers		Tamsulosin	
Anticholinergic Agents		Fesoterodine	
Antidiabetics	Gliclazide Glimepiride Glyburide Tolbutamide	Metformin Saxagliptin	
Anti-Retroviral Agents	Efavirenz Nevirapine		
Beta-3 Adrenergic Agonists	Mirabegron		
Cholinergic Agonists		Cevimeline	
Contraceptives			Estrogen-containing oral contraceptives
EGFR Inhibitors		Gefitinib	
Immunosuppressants		Sirolimus	
Vesicular monoamine transporter 2 inhibitor		Deutetrabenazine	
Xanthine Oxidase Inhibitor	Allopurinol		

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### Medication Report Details (by therapeutic class)

Drug		Finding	Recommendation	Concern	Evidence
Alpha-1 Blockers					
<b>Tamsulosin</b> (Flomax) FDA drug label: Actionable PGx		CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	-
Analgesics, Opioid					
Methadone (CYP2B6) (Dolophine) FDA drug label: Not established for PGx	<b>⊘</b>	CYP2B6: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		

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Drug		Finding	Recommendation	Concern	Evidence
Anti-ADHD Agents					
<b>Amphetamine</b> (Adzenys, Evekeo) FDA drug label: Not established for PGx	<b>⊘</b>	COMT(Val158Met): Normal function. Two normal function alleles.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		•
Atomoxetine (Strattera) FDA drug label: Actionable PGx		CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
<b>Clonidine</b> (Clonidine, Kapvay) FDA drug label: Not established for PGx		ADRA2A(C-1291G): Two variant alleles detected.	Individuals with rs1800544 GG genotype may be at an increased risk of therapeutic failure due to increased metabolic clearance.	Efficacy	
<b>Dexmethylphenidate</b> (Focalin) FDA drug label: Not established for PGx	<b>⊘</b>	COMT(Val158Met): Normal function. Two normal function alleles.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		
<b>Dextroamphetamine</b> (Zenzedi, Dexedrine) FDA drug label: Not established for PGx	<b></b>	COMT(Val158Met): Normal function. Two normal function alleles.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		
<b>Guanfacine</b> (Tenex, Intuniv) FDA drug label: Not established for PGx	•	CYP3A4: Ultrarapid metabolizer status. Two alleles showing increased function.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose.	Efficacy	
<b>Lisdexamfetamine</b> (Vyvanse) FDA drug label: Not established for PGx	<b></b>	COMT(Val158Met): Normal function. Two normal function alleles.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		
<b>Methylphenidate</b> (ADRA2A) (Concerta, Metadate, Ritalin, Ritalin LA, Quillivant, Daytrana, Methylin)		ADRA2A(C-1291G): Two variant alleles detected.	Individuals with rs1800544 GG genotype may be at an increased risk of therapeutic failure due to increased metabolic clearance.	Efficacy	
FDA drug label: Not established for PGx					

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Drug	Finding	Recommendation	Concern	Evidence
Methylphenidate (COMT) (Concerta, Metadate, Ritalin, Ritalin LA, Quillivant, Daytrana, Methylin) FDA drug label: Not	COMT(Val158Met): Normal function. Two normal function alleles.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		
established for PGx				
Antiarrhythmics				
Flecainide(Tambocor)FDA drug label: Notestablished for PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
Propafenone (Rythmol)FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Monitor the patient's response to guide dosing, or consider using an alternative medication.	Efficacy	
Anticholinergic Agents	GY			
Fesoterodine (Toviaz)FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with lower plasma concentrations of the active medication. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	-
Tolterodine (Detrol)FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	-

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Drug	Finding	Recommendation	Concern	Evidence
Anticoagulants				
Acenocoumarol (Sintrom, Acitrom) FDA drug label: Not established for PGx	CYP2C9: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Warfarin (Coumadin)FDA drug label: Actionable PGx	Multigenic: VKORC1, CYP2C9: Two alleles showing normal activity.; Normal metabolizer. Two alleles showing normal activity.	Individuals with this combination of alleles may benefit from an increased dose of Warfarin. The FDA table recommends a therapeutic dose of 5-7 mg/day.	ADR & Efficacy	
Warfarin (CYP4F2) (Coumadin) FDA drug label: Actionable PGx	Multigenic: CYP2C9, VKORC1, CYP4F2: Normal metabolizer. Two alleles showing normal activity.; Two alleles showing normal activity.; Normal function. Two alleles with normal activity.	Individuals with this combination of alleles may benefit from a standard therapeutic dose of warfarin. Consider a regimen of 4.1-5.9 mg/day (29-41 mg/week).	ADR & Efficacy	
	S			

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Drug	Finding	Recommendation	Concern	Evidence
Anticonvulsants				
Brivaracetam FDA drug label: Actionable PGx	CYP2C19: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Carbamazepine (Tegretol) FDA drug label: Testing required	HLA-B*1502: Negative; Absence of *15:02 alleles.	Individuals with wild type alleles are expected to show typical response. No additional therapeutic recommendations.		
Clobazam (Onfi) FDA drug label: Actionable PGx	CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Oxcarbazepine (Trileptal) FDA drug label: Testing recommended	HLA-B*1502: Negative; Absence of *15:02 alleles.	Individuals with wild type alleles are expected to show typical response. No additional therapeutic recommendations.		
Phenytoin (Cerebyx)Image: Comparison of the second	CYP2C9: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Antidementia Agents	GY			
Donepezil (Aricept)FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	•

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Drug	Finding	Recommendation	Concern	Evidence
Antidepressants				
Amitriptyline (Elavil)FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	
Bupropion (Wellbutrin)FDA drug label: Not established for PGx	ANKK1: Normal function. Two alleles with normal activity.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		
Clomipramine (Anafranil) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	
Desipramine (Norpramin) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	
Doxepin (Deptran) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	
Duloxetine (Cymbalta) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	-
Imipramine (Tofranil) FDA drug label: Actionable PGx	Multigenic: CYP2C19, CYP2D6: CYP2C19: Normal metabolizer. Two normal function alleles.	Multiple results from uncorrelated genes. CYP2C19: Typical response is expected; no additional therapeutic recommendations.; CYP2D6: Consider alternative therapy		

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Drug	Finding	Recommendation	Concern	Evidence
Mirtazapine FDA drug label: Not established for PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Typical response expected. No additional therapeutic recommendations.		
Moclobemide (Manerix, Aurorix, Amira, Clobemix, Depnil) FDA drug label: Not established for PGx	CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Nortriptyline (Pamelor) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	
Protriptyline (Vivactil)Image: Comparison of the second s	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	-
Trazodone (Oleptro, Desyrel)FDA drug label: Not established for PGx	CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose, or using an alternative medication.	Efficacy	-
Venlafaxine (Effexor) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose, or using an alternative medication.	Efficacy	
Vortioxetine (Brintellix) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider adjusting the dose, or monitoring the patient's response to guide dosing.	Efficacy	

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Drug	Finding	Recommendation	Concern	Evidence
Antidiabetics				
Gliclazide FDA drug label: Not established for PGx	CYP2C9: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Glimepiride FDA drug label: Not established for PGx	CYP2C9: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Glyburide (Glibenclamide) FDA drug label: Not established for PGx	CYP2C9: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Metformin (Glucophage®)	ATM: Enhanced response.	Increased drug efficacy likely.		
FDA drug label: Not established for PGx				
Saxagliptin (Onglyza) FDA drug label: Not established for PGx	CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose, or using an alternative medication.	Efficacy	0
Tolbutamide (Orinase) FDA drug label: Not established for PGx	CYP2C9: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Antiemetics				
Ondansetron (Zofran) FDA drug label: Informative PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	
<b>Tropisetron</b> (Navoban, Setrovel) FDA drug label: Not established for PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	•

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Drug		Finding	Recommendation	Concern	Evidence
Antifungals					
<b>Ketoconazole</b> (Nizoral) FDA drug label: Not established for PGx	0	CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	0
<b>Voriconazole</b> (Vfend) FDA drug label: Actionable PGx		CYP2C19: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Antineoplastic Agen	its				
<b>Methotrexate</b> (Rheumatrex, Trexall) FDA drug label: Not established for PGx	•?	MTHFR: Uncertain Allele	No recommendation for Methotrexate is available for this combination of variants/ alleles.		
Antiplatelet Agents					
<b>Clopidogrel</b> FDA drug label: Actionable PGx	<b>⊘</b>	CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
<b>Prasugrel</b> FDA drug label: Informative PGx	<b>⊘</b>	CYP2C19: Normal metabolizer. Two alleles showing normal activity.	Typical response expected. No additional therapeutic recommendations.		•
<b>Ticagrelor</b> (Brilinta) FDA drug label: Not established for PGx		CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	0

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Drug	Finding	Recommendation	Concern	Evidence
Antipsychotics				
Aripiprazole (Abilify) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Brexpiprazole (Rexulti) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
Clozapine	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
FlupenthixolFDA drug label: Not established for PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Typical response expected. No additional therapeutic recommendations.		
Haloperidol (Haldol) FDA drug label: Not established for PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
Iloperidone         FDA drug label:         Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	•
Olanzapine (Zalasta, Zyprexa) FDA drug label: Not established for PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Typical response expected. No additional therapeutic recommendations.		

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Drug	Finding	Recommendation	Concern	Evidence
Perphenazine (Trilafon) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele	thus an increased risk of pharmacotherapy	Efficacy	-
<b>Pimozide</b> (Orap) FDA drug label: Testing required	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele	thus an increased risk of pharmacotherapy	Efficacy	
Quetiapine (Seroquel) FDA drug label: Not established for PGx	CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose, or using an alternative medication.	Efficacy	0
<b>Risperidone</b> (Risperdal) FDA drug label: Informative PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele	thus an increased risk of pharmacotherapy	Efficacy	
Thioridazine FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele	thus an increased risk of pharmacotherapy	Efficacy	•
Zuclopenthixol FDA drug label: Not established for PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele	thus a significantly increased risk of	Efficacy	

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Drug	Finding	Recommendation	Concern	Evidence
Anti-Retroviral Agent	ts			
<b>Efavirenz</b> (Sustiva) FDA drug label: Actionable PGx	CYP2B6: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
<b>Nevirapine</b> (Viramune) FDA drug label: Not established for PGx	<ul> <li>CYP2B6: Normal metabolizer. Two normal function alleles.</li> </ul>	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Anxiolytics				
Alprazolam (Xanax, Niravam) FDA drug label: Not established for PGx	CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider adjusting the dose, or using an alternative medication.	Efficacy	-
<b>Buspirone</b> (Buspar) FDA drug label: Not established for PGx	CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose, or using an alternative medication.	Efficacy	0
<b>Clonazepam</b> (Klonopin) FDA drug label: Not established for PGx	CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Monitor the patient's response to guide dosing, or consider using an alternative medication.	Efficacy	0
<b>Diazepam</b> FDA drug label: Actionable PGx	CYP2C19: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Beta-3 Adrenergic Ag	gonists			
<b>Mirabegron</b> (Myrbetriq) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with lower plasma concentrations of the active medication. No additional therapeutic recommendations.		-

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Drug	Finding	Recommendation	Concern	Evidence
Beta Blockers				
Carvedilol (Coreg) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose, or using an alternative medication.	Efficacy	
Metoprolol (Lopressor) FDA drug label: Informative PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose, or using an alternative medication.	Efficacy	
Nebivolol (Bystolic) FDA drug label: Informative PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Typical response expected. No additional therapeutic recommendations.		•
Propranolol (Inderal) FDA drug label: Informative PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Typical response expected. No additional therapeutic recommendations.		-
<b>Timolol</b> (Blocadren) FDA drug label: Not established for PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy.	Efficacy	-
Central Monoamine-D	Pepleting Agents			
<b>Tetrabenazine</b> (Xenazine) FDA drug label: Testing required	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
Central Nervous Syste	em Agents			
Dextromethorphan- Quinidine (Nuedexta) FDA drug label: Testing recommended	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	

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Drug		Finding	Recommendation	Concern	Evidence
Cholinergic Agonist	S				
<b>Cevimeline</b> (Evoxac) FDA drug label: Actionable PGx		CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
Cholinesterase Inhit	oitors				
Galantamine (Razadyne, Razadyne ER, Nivalin, Lycoremine, Reminyl) FDA drug label: Informative PGx		CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	-
Contraceptives					
Estrogen-containing oral contraceptives FDA drug label: Not established for PGx	•	F5: Two Leiden alleles	Individuals with these homozygous variant alleles frequently present with significantly increased risk of side effects. This medication should be avoided.	ADR	
EGFR Inhibitors		Λ			
<b>Gefitinib</b> (Iressa) FDA drug label: Actionable PGx		CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
Endocrine-Metaboli	c Age	ents			
<b>Eliglustat</b> FDA drug label: Testing required	•	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	

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Drug		Finding	Recommendation	Concern	Evidence
Hypnotics					
<b>Eszopiclone</b> (Lunesta) FDA drug label: Not established for PGx		CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Monitor the patient's response to guide dosing, or consider using an alternative medication.	Efficacy	0
lmmunosuppressar	nts				
<b>Azathioprine</b> (Imuran) FDA drug label: Testing recommended	<b>⊘</b>	TPMT: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
<b>Cyclosporine</b> (Gengraf, Neoral) FDA drug label: Not established for PGx		CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
Mercaptopurine (Purinethol) FDA drug label: Testing recommended	<b></b>	TPMT: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
<b>Sirolimus</b> (Rapamune) FDA drug label: Not established for PGx		CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	0
<b>Tacrolimus</b> (Prograf, Hecoria) FDA drug label: Not established for PGx		CYP3A5: Two alleles showing normal activity.	Normal metabolizers of this medication may present with lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose; monitor the patient's response to guide dosing.	Efficacy	•
<b>Thioguanine</b> (6-TG, Tabloid, Lanvis) FDA drug label: Testing recommended	<b>S</b>	TPMT: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•

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Drug		Finding	Recommendation	Concern	Evidence
Muscle Relaxants					
<b>Carisoprodol</b> (Soma)	Ø	CYP2C19: Normal metabolizer. Two	Normal metabolizers of this medication are expected to show typical response. No		
FDA drug label: Actionable PGx		normal function alleles.	additional therapeutic recommendations.		



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Drug		Finding	Recommendation	Concern	Evidence
Non-drug					
ABCG2	0	ABCG2: Normal function. Two normal function alleles.	Typical response is expected; no additional therapeutic recommendations.		
ADRA2A(C-1291G)	•	ADRA2A(C-1291G): Two variant alleles detected.	Uncertain phenotype		
ANKK1	0	ANKK1: Normal function. Two alleles with normal activity.	Normal function. Two alleles with normal activity.		
АроЕ	0	ApoE: Often associated with normal lipid metabolism.	Typical cardiovascular disease risk expected.		
COMT(Val158Met)	2	COMT(Val158Met): Normal function. Two normal function alleles.	No recommendation for COMT(Val158Met) is available for this combination of variants/ alleles.		
CYP1A2	0	CYP1A2: Normal metabolizer. Two alleles showing normal activity.	No additional therapeutic recommendations.		
CYP2B6	0	CYP2B6: Normal metabolizer. Two normal function alleles.	No additional therapeutic recommendations.		
CYP2C8	<b>⊘</b>	CYP2C8: Normal metabolizer. Two alleles showing normal activity.	Typical response is expected; no additional therapeutic recommendations.		
CYP4F2	0	CYP4F2: Normal function. Two alleles with normal activity.	Normal function. Two alleles with normal activity.		
GRIK4	0	GRIK4: Altered function. Two altered function alleles.	Altered function. Two alleles with altered activity.		
HTR2A	0	HTR2A: Normal function. Two alleles with normal activity.	Normal function. Two alleles with normal activity.		
HTR2C(-759C>T)	•	HTR2C(-759C>T): C/ C	Uncertain phenotype		

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Drug		Finding	Recommendation	Concern	Evidence
IFNL3	<b>⊘</b>	IFNL3: Normal function. Two normal function alleles.	Typical response is expected; no additional therapeutic recommendations.		
MTHFR (A1298C)	•	MTHFR (A1298C): Uncertain Allele	No recommendation for MTHFR (A1298C) is available for this combination of variants/ alleles.		
MTHFR (C677T)	•	MTHFR (C677T): Uncertain Allele	No recommendation for MTHFR (C677T) is available for this combination of variants/ alleles.		
OPRM1(A118G)	0	OPRM1(A118G): Normal function. Two alleles with normal activity.	Normal function. Two alleles with normal activity.		

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Drug	Finding	Recommendation	Concern	Evidence
Nonsteroidal Anti-Inflam	natory Drugs (NSAIDs	5)		
Celecoxib (Celebrex)FDA drug label: Actionable PGx	CYP2C9: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Diclofenac (Cataflam) FDA drug label: Not established for PGx	CYP2C9: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Flurbiprofen (Ocufen)FDA drug label: Actionable PGx	CYP2C9: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Ibuprofen (Motrin, Advil) FDA drug label: Not established for PGx	CYP2C9: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Lornoxicam (Xefo) FDA drug label: Not established for PGx	CYP2C9: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Meloxicam (Mobic) FDA drug label: Actionable PGx	CYP2C9: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Piroxicam (Feldene) FDA drug label: Actionable PGx	CYP2C9: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•

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Drug	Finding	Recommendation	Concern	Evidence
Opioids				
Alfentanil (Rapifen, Alfenta) FDA drug label: Not established for PGx	OPRM1(A118G): Normal function. Tv alleles with normal activity.	Individuals with normal function of this yo gene are expected to show typical response. No additional therapeutic recommendations.		•
<b>Buprenorphine</b> (Butrans, Buprenex) FDA drug label: Not established for PGx	CYP3A4: Ultrarapid metabolizer. Two increased function alleles.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	0
<b>Codeine</b> FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function alle and one duplicated normal function alle	frequently present with notably higher plasma concentrations of the active medication, thus a significantly increased	ADR	•
<b>Fentanyl</b> (Duragesic, Sublimaze) FDA drug label: Not established for PGx	CYP3A4: Ultrarapid metabolizer status. Two alleles showing increased function.	may present with notably lower plasma	Efficacy	-
Fentanyl (OPRM1) (Duragesic, Sublimaze) FDA drug label: Not established for PGx	OPRM1(A118G): Normal function. Tv alleles with normal activity.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		•
<b>Hydrocodone</b> FDA drug label: Not established for PGx	CYP2D6: Ultrarapid metabolizer. One normal function alle and one duplicated normal function alle	may present with notably higher plasma concentrations of the active medication, thus an increased risk of side effects. Be	ADR	•
<b>Hydromorphone</b> (Dilaudid) FDA drug label: Not established for PGx	OPRM1(A118G): Normal function. Tv alleles with normal activity.	Individuals with normal function of this yo gene are expected to show typical response. No additional therapeutic recommendations.		
<b>Morphine</b> (MS-IR) FDA drug label: Not established for PGx	OPRM1(A118G): Normal function. Tv alleles with normal activity.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		

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Drug	Finding	Recommendation	Concern	Evidence
Oxycodone (Oxycontin) FDA drug label: Not established for PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably higher plasma concentrations of the active medication, thus a significantly increased risk of side effects. This medication should be avoided.	ADR	-
Oxycodone (CYP3A5) (Oxycontin) FDA drug label: Not established for PGx	CYP3A5: Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		•
Sufentanil (Sufenta) FDA drug label: Not established for PGx	OPRM1(A118G): Normal function. Two alleles with normal activity.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		•
<b>Tramadol</b> (Ultracet, Ultram) FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably higher plasma concentrations of the active medication, thus a significantly increased risk of side effects. This medication should be avoided.	ADR	
	SP			

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Drug		Finding	Recommendation	Concern	Evidence
Proton Pump Inhibi	tors (F	PPIs)			
<b>Dexlansoprazole</b> (Kapidex, Dexilant) FDA drug label: Actionable PGx		CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication may present with lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Monitor the patient's response to guide dosing; consider increasing the dose.	Efficacy	•
<b>Esomeprazole</b> (Nexium) FDA drug label: Actionable PGx	<b>~</b>	CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Lansoprazole (Prevacid) FDA drug label: Informative PGx		CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication may present with lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Monitor the patient's response to guide dosing; consider increasing the dose.	Efficacy	•
<b>Omeprazole</b> (Prilosec, Zegerid) FDA drug label: Actionable PGx		CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication may present with lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Monitor the patient's response to guide dosing; consider increasing the dose.	Efficacy	•
<b>Pantoprazole</b> (Protonix) FDA drug label: Actionable PGx		CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication may present with lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Monitor the patient's response to guide dosing; consider increasing the dose.	Efficacy	•
<b>Rabeprazole</b> (Aciphex) FDA drug label: Actionable PGx	<b>⊘</b>	CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		

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Drug	Finding	Recommendation	Concern	Evidence
Selective Serotonin Reup	otake Inhibitors (SSRI	5)		
Citalopram (Celexa)FDA drug label: Actionable PGx	CYP2C19: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Escitalopram (Lexapro) FDA drug label: Actionable PGx	CYP2C19: Normal metabolizer. Two alleles showing normal activity.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Fluoxetine (Prozac)Image: Comparison of the second	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Typical response expected. No additional therapeutic recommendations.		
Fluvoxamine (Luvox)FDA drug label: Actionable PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. Be alert to lack of efficacy; consider alternative medication.	Efficacy	•
Paroxetine (Paxil)FDA drug label: Informative PGx	CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication frequently present with notably lower plasma concentrations of the active medication, thus a significantly increased risk of pharmacotherapy failure. This medication should be avoided.	Efficacy	
Sertraline (Zoloft) FDA drug label: Not established for PGx	CYP2C19: Normal metabolizer. Two normal function alleles.	Normal metabolizers of this medication are expected to show typical response. No additional therapeutic recommendations.		
Statins				
Atorvastatin (Lipitor, Caduet) FDA drug label: Not established for PGx	CYP3A4: Ultra-rapid metabolizer. Two alleles showing increased activity.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Consider increasing the dose, or using an alternative medication.	Efficacy	-
Simvastatin (Zocor) FDA drug label: Informative PGx	SLCO1B1: Normal liver uptake activity.	Individuals with normal SLCO1B1 liver uptake activity are expected to have a typical response to a standard dose of simvastatin.		

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Drug		Finding	Recommendation	Concern	Evidence
Vesicular monoamine transporter 2 inhibitor					
<b>Deutetrabenazine</b> (Austedo) FDA drug label: Actionable PGx		CYP2D6: Ultrarapid metabolizer. One normal function allele and one duplicated normal function allele.	Ultrarapid metabolizers of this medication may present with notably lower plasma concentrations of the active medication, thus an increased risk of pharmacotherapy failure. Be alert to lack of efficacy; monitor the patient's response to guide dosing.	Efficacy	
Xanthine Oxidase Ir	hibit	or			
<b>Allopurinol</b> (Zyloprim)	<b>⊘</b>	ABCG2: Normal function. Two normal function alleles.	Individuals with normal function of this gene are expected to show typical response. No additional therapeutic recommendations.		

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### **Clinical Evidence Levels**

### Strong

- Includes gene-drug pairs approved by the Global Research Institute for Medical Research Pharmacogenomics Advisory Group.
- Includes gene-drug pairs supported by multiple studies documenting consistent effects of specific genetic variant(s) on clinical outcomes.
- Includes gene-drug pairs approved by the Dutch Pharmacogenetics Working Group (DPWG) and/or guidelines
  published in Clinical Pharmacology and Therapeutics by the Clinical Pharmacogenetics Implementation Consortium
  (CPIC).

Moderate

- Includes gene-drug pairs supported by pharmacokinetic, pharmacodynamic, or molecular/cellular functional studies showing consistent effects of genetic variant(s).
- Includes Drug product information (e.g. This interpretation is based on guidance available in the FDA (Food and Drug Administration) drug label for ABILIFY® (10/2013).
- Includes gene-drug pairs for which potential clinical outcomes are inferred from similar gene-drug interactions approved by the Dutch Pharmacogenetics Working Group (DPWG), and/or guidelines published in Clinical Pharmacology and Therapeutics by the Clinical Pharmacogenetics Implementation Consortium (CPIC), and/or pharmacogenomic reports and submission from the Global Research Institute for Medical Research.

### Emerging

• Includes gene-drug pairs supported by published studies of the drug, related drug, or a probing compound of interest involving limited data and/or inconsistent findings.

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### **Patient Information Card**

This card contains an abbreviated genetic summary.

It is not intended as a replacement for the complete GeneDose<sup>™</sup> report.

it is not interface as	arcpideenier	it for the complete defields			
r			CYP3A4	*1B *1B	Ultrarapid metabolizer
GRI: https://www.	gri-labs.com	1	СҮРЗА5	*1D *1D; or *1A *1A; or *1A *1D	Normal metabolizer
	-		CYP4F2	*1 *1	Normal metabolizer
Patient:	Doe, Jane		Factor V Leiden	Variant	See full GeneDose report
DOB:	1990-01-01		GRIK4	T T	Altered function
Sample ID:	ex225		HLA-B*1502	WT WT	Negative
This card shows info	ormation abo	ut your genetics that relate	HTR2A	AAA	Normal function
		r doctors before being	HTR2C(-759C>T)	C/C	Unknown Metabolizer
prescribed new me	dications.		IFNL3	C C	Normal function
Pha ABCG2	rmacogenom G G	ic Summary Normal function	MTHFR	Uncertain Allele	See full GeneDose report
ADRA2A(C-1291G	) G G	Unknown Metabolizer	MTHFR (A1298C)	Uncertain Allele	See full GeneDose report
ANKK1	G G	Normal function		Uncertain	
АроЕ	ε3 ε3	See full GeneDose report	MTHFR (C677T)	Allele	See full GeneDose report
ATM	C C	Increased likelihood of treatment success when	OPRM1(A118G)	AIA	Normal function
	010	taking metformin	Prothrombin (F2)	Normal	See full GeneDose report
COMT(Val158Met)	G G	Normal function	SLCO1B1	*1 *1	Normal liver uptake
CYP1A2	*1A *1A	Normal metabolizer			activity
CYP2B6	*1A *1A	Normal metabolizer	ТРМТ	*1 *1	Normal metabolizer
CYP2C19	*1 *1	Normal metabolizer	VKORC1	*1 *1	Normal (with respect to
CYP2C8	YP2C8 *1 *1 Normal metabolizer				Warfarin) ed by GRI
CYP2C9	*1 *1	Normal metabolizer		Powere	
CYP2D6	*1Ax2 *1A	Ultrarapid metabolizer			
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