

# Interpretation Guide

## Pharmacogenomic (PGx) Report

The following PGx report is a clinical decision support tool based on individual genetic results. It contributes to a better understanding and prediction of medication response and tolerability. This test does not predict the risk of any health problem. **Since response to medications is multifactorial, clinical judgment supersedes any recommendations provided.**

The report notifies you if the patient carries any genetic variant that can alter the following pharmacological parameters:









- pharmacokinetics: overall **exposure** to a medication depending on metabolic and efflux pump function;
- pharmacodynamics: the potential **efficacy** of a drug and whether the patient is predisposed to certain **atypical effects**.

These results do not change with age, but their interpretation can evolve as new data becomes available. Therefore, the Biron PGx reports are updated periodically. These results can also be useful for other medications, not covered by the report.

### How to use pharmacogenomic recommendations

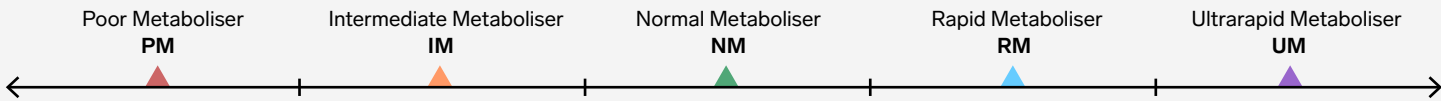
1. Only medications relevant for your patient need to be consulted.
2. Use the **Exposure** column to adjust doses for adequate plasma concentrations.
3. Use the **Efficacy** and **Risk of atypical effect** columns to choose the most compatible medication.

The **Exposure**, **Efficacy** and **Risk of atypical effect** columns are interpreted independently from each other. Medications are ordered by class with the most compatible options listed first within each class.

Exposure	Efficacy	Risk of atypical effect
 A higher dose may be required to achieve adequate plasma concentrations.	When choosing between multiple clinically appropriate medications, you may give preference to a medication in which a lower number of variants have been identified (e.g., 2/2 is better than 4/6), in terms of their association with an increased likelihood of a poorer response or an atypical effect.	
 A lower dose may be required to achieve adequate plasma concentrations.		
 Several metabolic pathways are involved, but their capacities are opposed (e.g., PM and UM). Thus, a calculation of dose adjustments is not possible based on current data and closer monitoring is recommended.	 Signifies the presence of a high-impact gene variant, which increases the probability of a poorer response.	 Signifies the presence of variants associated with an increased risk of particular side effects, compared to non-carriers.
 Drug not recommended by peer-reviewed guidelines due to a risk of toxicity or lack of efficacy.	 Signifies that all of the tested variants predict an increased likelihood of a better response, compared to non-carriers. This medication may be a good option.	 Medication not recommended by peer-reviewed guidelines due to a risk of severe side effects,

The notification Normal efficacy\* or Normal risk\* signifies that there is currently no available data allowing for a genetically-based prediction of medication effect.

### Nomenclature for enzyme phenotypes (e.g., cytochrome P450s or CYP)



**NM** is generally used to establish standard doses. This dose may be too high for **PM/IM** or too low for **RM/UM**, warranting a dose adjustments or the consideration of an alternative agent. For a pro-drug (e.g., clopidogrel, tramadol), phenotype variability will have the opposite effect.

Inducible Metaboliser (Ind) - Specific for CYP1A2, which can have increased function in the presence of an inducer, such as tobacco smoke, comparable to **RM/UM**.

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PHARMACOGENOMIC REPORT



Psychiatry and ADHD, Pain Management, Cardiology

To download the latest version, go here: [secur.biron.com/login](https://secur.biron.com/login)

DO NOT DISTRIBUTE. As per the Genetic Non-Discrimination Act, the information herein is intended for the patient and his/her healthcare providers. Without explicit patient consent, this report cannot be shared and/or used with other persons or organisations, including insurers and employers.

Many factors may affect the choice of optimal medication (e.g. diagnosis, environmental factors, epigenetics), hence this report is intended as a clinical decision support tool, does not constitute medical advice and does not supersede the professional judgement of the clinician. When supported by published guidelines, firm recommendations are provided and referenced, otherwise the evidence is directional in nature. For additional online ressources, visit [biron.com/pgxtest](https://biron.com/pgxtest).

ADMINISTRATIVE DATA		
<b>Patient Name</b> Michel The Great	<b>Ordering Clinician</b> N/A	<b>Sample ID:</b> bio022043 <b>Sample Type:</b> saliva
<b>Sex assigned at birth:</b> Male <b>Date of birth:</b> 1982-01-29 <b>Phone Number:</b> N/A <b>Email:</b> N/A	<b>Patient Address</b> 123 Rue des Tuileries Québec, QC G2G1K5	<b>Date ordered:</b> 2025-11-28 <b>Date of sample reception:</b> 2025-11-28 <b>Date of report:</b> 2025-11-28
<b>Clinical Support</b> <b>Email:</b> <a href="mailto:genetique@biron.com">genetique@biron.com</a>	<b>Phone:</b> 1-855-943-6379	<b>Fax:</b> (514) 317-2241

ATYPICAL PHENOTYPES
CYP1A2 IND, CYP2B6 IM, CYP4F2 PM, DPYD IM, POR PA, NUDT15 PM, SLCO1B1 Poor function, TPMT PM, UGT2B15 IM.
NM: Normal Metaboliser, IM: Intermediate Metaboliser, PM: Poor Metaboliser, RM: Rapid Metaboliser, UM: Ultrarapid Metaboliser, Ind: Inducible Metaboliser, NA: Normal Activity, IA: Intermediate Activity, PA: Poor Activity.

CAUTIONARY INFORMATION		
Medication	Identified risk	Recommendation
Capecitabine, 5-fluorouracil	Decreased DPD activity (DPD activity at 30% to 70% that of the normal population) and increased risk for severe or even fatal drug toxicity when treated with fluoropyrimidines.	Reduce starting dose by 50% followed by titration of dose based on toxicity or therapeutic drug monitoring, if available. Increase the dose in patients experiencing no or clinically tolerable toxicity in the first two cycles to maintain efficacy; decrease the dose in patients who do not tolerate the starting dose to minimize toxicities (DPYD IM, c.2846A>T/c.2846A>T). <sup>1</sup>
Azathioprine	Extremely high concentrations of TGN metabolites; fatal toxicity possible without dose decrease; no MeTIMP metabolites. Greatly increased risk of thiopurine-related leukopenia, neutropenia, myelosuppression.	For nonmalignant conditions, consider alternative nonthiopurine immunosuppressant therapy. For malignancy, start with drastically reduced doses (reduce daily dose by 10-fold and dose thrice weekly instead of daily) and adjust doses of azathioprine based on degree of myelosuppression and disease-specific guidelines. Allow 4–6 weeks to reach steady-state after each dose adjustment (TPMT PM, NUDT15 PM). <sup>2</sup>
Mercaptopurine	Extremely high concentrations of TGN metabolites; fatal toxicity possible without dose decrease; no MeTIMP metabolites. Greatly increased risk of thiopurine-related leukopenia, neutropenia, myelosuppression.	For malignancy, start with drastically reduced doses (reduce daily dose by 10-fold and reduce frequency to thrice weekly instead of daily (e.g., 10 mg/m²/day given just 3 days/week) and adjust doses of mercaptopurine based on degree of myelosuppression and disease-specific guidelines. Allow 4–6 weeks to reach steady-state after each dose adjustment. If myelosuppression occurs, emphasis should be on reducing mercaptopurine over other agents. For nonmalignant conditions, consider alternative nonthiopurine immunosuppressant therapy (TPMT PM, NUDT15 PM). <sup>2</sup>

CAUTIONARY INFORMATION

Medication	Identified risk	Recommendation
Thioguanine	Extremely high concentrations of TGN metabolites; fatal toxicity possible without dose decrease. Greatly increased risk of thiopurine-related leukopenia, neutropenia, myelosuppression.	Start with drastically reduced doses (reduce daily dose by 10-fold and dose thrice weekly instead of daily) and adjust doses of thioguanine based on degree of myelosuppression and disease-specific guidelines. Allow 4–6 weeks to reach steady-state after each dose adjustment. If myelosuppression occurs, emphasis should be on reducing thioguanine over other agents. For nonmalignant conditions, consider alternative nonthiopurine immunosuppressant therapy (TPMT PM, NUDT15 PM). <sup>2</sup>
Cisplatin	Increased risk of cisplatin-induced hearing loss (ototoxicity).	To reduce the risk of ototoxicity, monitor hearing loss more closely and consider the use of otoprotectants. Alternative treatments may be considered based on disease-specific guidelines (TPMT PM). <sup>3</sup>

## PGx RECOMMENDATIONS - PSYCHIATRY AND ADHD



Dose increase may be required due to lower exposure to active metabolite(s) with standard dosing.



Increased probability of a better response.



Dose reduction may be required due to higher exposure to active metabolite(s) with standard dosing.



Greater potential for a poorer response or atypical effect.



Exposure is difficult to predict, insufficient data to calculate dose adjustments.



Medication not recommended by peer-reviewed guidelines.

Normal exposure\*, Normal efficacy\* or Normal risk\*: Based on currently available genetic data, the efficacy or risk of an atypical effect is likely similar to that of most other individuals; further research is needed to better understand genetic influence.

### Genetic Associations Identified

Medications	Exposure	Efficacy	Risk of atypical effect
<b>Antidepressants</b>			
<b>Selective serotonin reuptake inhibitors (SSRIs)</b>			
<b>Fluoxetine</b> (Prozac®)	Normal exposure (CYP2D6 NM, CYP2C9 NM).	4/6 variants: increased likelihood of a poorer response (FKBP5, BDNF, GRIK4, HTR7).	Normal risk*
<b>Fluvoxamine</b> (Luvox®)	Initiate with recommended dose; a low dose may be adequate (CYP2D6 NM, ABCB1).	4/6 variants: increased likelihood of a poorer response (FKBP5, BDNF, GRIK4, HTR7).	Normal risk*
<b>Paroxetine</b> (Paxil®)	Initiate therapy with recommended starting dose; a low dose may be adequate (CYP2D6 NM, ABCB1).	4/6 variants: increased likelihood of a poorer response (FKBP5, BDNF, GRIK4, HTR7).	Normal risk*
<b>Citalopram</b> (Celexa®)	Initiate with recommended dose; a low dose may be adequate (CYP2C19 NM, CYP3A4 NM, ABCB1).	4/6 variants: increased likelihood of a poorer response (FKBP5, BDNF, GRIK4, HTR7).	1/1 variant: increased risk of gastrointestinal side effects (HTR2A).
<b>Escitalopram</b> (Ciprallex®)	Initiate with recommended dose; a low dose may be adequate (CYP2C19 NM, CYP3A4 NM, ABCB1).	4/6 variants: increased likelihood of a poorer response (FKBP5, BDNF, GRIK4, HTR7).	1/1 variant: increased risk of gastrointestinal side effects (HTR2A).
<b>Sertraline</b> (Zoloft®)	Initiate therapy with recommended starting dose but with a slower titration schedule and a lower maintenance dose (CYP2B6 IM, CYP2C19 NM). <sup>4</sup>	4/5 variants: increased likelihood of a poorer response (BDNF, FKBP5, GRIK4, HTR7)	1/1 variant: increased risk of gastrointestinal side effects (HTR2A).
<b>Serotonin–norepinephrine reuptake inhibitors (SNRIs)</b>			
<b>Desvenlafaxine</b> (Pristiq®)	Initiate with recommended dose (UGT1A1 NM, UGT2B15 IM, CYP3A4 NM).	2/2 variants: increased likelihood of a poorer response (FKBP5, GRIK4).	Normal risk*
<b>Levomilnacipran</b> (Fetzima®)	Initiate with recommended dose; a low dose may be adequate (CYP3A4 NM, ABCB1).	2/2 variants: increased likelihood of a poorer response (FKBP5, GRIK4).	Normal risk*
<b>Venlafaxine-XR</b> (Effexor XR®)	Initiate with recommended dose; a low dose may be adequate (CYP2D6 NM, ABCB1).	4/4 variants: increased likelihood of a poorer response (COMT, FKBP5, GRIK4, SLC6A2).	Normal risk*
<b>Duloxetine</b> (Cymbalta®)	Initiate therapy with recommended starting dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind, CYP2D6 NM).	2/3 variants: increased likelihood of a poorer response (FKBP5, GRIK4).	Normal risk*
<b>Other antidepressants</b>			
<b>Trazodone</b> (Desyrel®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Vilazodone</b> (Viibryd®)	Initiate with recommended dose; a low dose may be adequate (CYP3A4 NM, ABCB1).	Normal efficacy*	Normal risk*

## Genetic Associations Identified

Medications	Exposure	Efficacy	Risk of atypical effect
<b>Vortioxetine</b> (Trintellix®)	Normal exposure (CYP2D6 NM, CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Mirtazapine</b> (Remeron®)	⬆️ Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind, CYP2D6 NM, CYP3A4 NM).	2/2 variants: increased likelihood of a poorer response (FKBP5, TPH2).	Normal risk*
<b>Bupropion</b> (Wellbutrin®)	⬇️ Consider using a lower dose (CYP2B6 IM, POR).	0/1 variant: no increased likelihood of a poorer response for treatment of depressive symptoms (HTR2A).	Normal risk*
<b>Esketamine</b> (Spravato®)	⬇️ Consider using a lower dose (CYP2B6 IM, POR).	0/1 variant: no increased likelihood of a poorer response.	0/1 variant: no increased risk of emergent hypertension.
<b>Ketamine</b> (Ketalar®)	⬇️ Consider using a lower dose (CYP2B6 IM, POR).	0/1 variant: no increased likelihood of a poorer response.	0/1 variant: no increased risk of emergent hypertension.
<b>Tricyclic antidepressants (TCAs)</b>			
<b>Amitriptyline</b> (Elavil®)	CYP2C19 NM, CYP2D6 NM, ABCB1 - Initiate with recommended dose; a low dose may be adequate.	1/1 variant: increased likelihood of a poorer response (TPH2)	Genetic influence not available
<b>Clomipramine</b> (Anafranil®)	CYP2C19 NM, CYP2D6 NM - Normal exposure <sup>5</sup>	1/1 variant: increased likelihood of a poorer response (TPH2)	Genetic influence not available
<b>Desipramine</b> (Norpramin®)	CYP2D6 NM, ABCB1 - Initiate with recommended dose; a low dose may be adequate.	1/1 variant: increased likelihood of a poorer response (TPH2)	Genetic influence not available
<b>Doxepin</b> (Sinequan®)	CYP2C19 NM, CYP2D6 NM, ABCB1 - Initiate with recommended dose; a low dose may be adequate.	1/1 variant: increased likelihood of a poorer response (TPH2)	Genetic influence not available
<b>Imipramine</b> (Tofranil®)	CYP2C19 NM, CYP2D6 NM - Normal exposure <sup>5</sup>	1/1 variant: increased likelihood of a poorer response (TPH2)	Genetic influence not available
<b>Nortriptyline</b> (Aventyl®)	CYP2D6 NM, ABCB1 - Initiate with recommended dose; a low dose may be adequate.	1/1 variant: increased likelihood of a poorer response (TPH2)	Genetic influence not available
<b>Trimipramine</b> (Surmontil®)	CYP2C19 NM, CYP2D6 NM, ABCB1 - Initiate with recommended dose; a low dose may be adequate.	1/1 variant: increased likelihood of a poorer response (TPH2)	Genetic influence not available
<b>Monoamine oxidase inhibitors (MAOs)</b>			
<b>Moclobemide</b> (Manerix®)	Normal exposure (CYP2D6 NM, CYP2C19 NM).	Normal efficacy*	Normal risk*
<b>Phenelzine</b> (Nardil®)	Normal exposure*	Normal efficacy*	Normal risk*
<b>Tranlycypromine</b> (Parnate®)	Normal exposure*	Normal efficacy*	Normal risk*
<b>Antipsychotics</b>			
<b>1st generation antipsychotics</b>			
<b>Chlorpromazine</b> (Largactil®)	Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP2D6 NM, CYP1A2 Ind).	Normal efficacy*	Normal risk*
<b>Fluphenazine</b> (Moditen®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*
<b>Haloperidol</b> (Haldol®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	0/1 variant: no increased risk of extrapyramidal side effects.
<b>Perphenazine</b> (Trilafon®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*

## Genetic Associations Identified

Medications	Exposure	Efficacy	Risk of atypical effect
<b>Pimozide</b> (Orap®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*
<b>Zuclopenthixol</b> (Clopixol®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*
<b>Loxapine</b> (Loxapac®)	⬆️ Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind).	Normal efficacy*	Normal risk*
<b>Trifluoperazine</b> (Stelazine®)	⬆️ Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind).	Normal efficacy*	Normal risk*


## 2nd generation antipsychotics

<b>Aripiprazole</b> (Abilify®)	Normal exposure (CYP2D6 NM).	1/1 variant: increased likelihood of a poorer response (ANKK1).	Normal risk*
<b>Brexpiprazole</b> (Rexulti®)	Normal exposure (CYP2D6 NM, CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Cariprazine</b> (Vraylar®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Lurasidone</b> (Latuda®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Paliperidone</b> (Invega®)	Initiate with recommended dose; a low dose may be adequate (CYP3A4 NM, ABCB1).	Normal efficacy*	Normal risk*
<b>Quetiapine</b> (Seroquel®)	Initiate with recommended dose; a low dose may be adequate (CYP3A4 NM, ABCB1).	Normal efficacy*	1/2 variants: increased risk of antipsychotic-induced weight gain (MC4Ra).
<b>Risperidone</b> (Risperdal®)	Normal exposure (CYP2D6 NM).	1/1 variant: increased likelihood of a poorer response (ANKK1).	1/6 variants: increased risk of antipsychotic-induced weight gain (MC4R).
<b>Ziprasidone</b> (Zeldox®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Asenapine</b> (Saphris®)	⬆️ Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind, UGT1A4 NM).	Normal efficacy*	Normal risk*
<b>Clozapine</b> (Clozaril®)	⬆️ Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind, CYP2D6 NM).	Normal efficacy*	1/6 variants: increased risk of antipsychotic-induced weight gain (MC4R).
<b>Olanzapine</b> (Zyprexa®)	⬆️ Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind).	Normal efficacy*	1/6 variants: increased risk of antipsychotic-induced weight gain (MC4R).

## Anxiolytics

<b>Alprazolam</b> (Xanax®)	Normal exposure (CYP3A4 NM, CYP3A5 PM).	Normal efficacy*	Normal risk*
<b>Buspirone</b> (Buspar®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Chlordiazepoxide</b> (Librium®)	Initiate with recommended dose; a low dose may be adequate (CYP3A4 NM, UGT2B15 IM).	Normal efficacy*	Normal risk*
<b>Clobazam</b> (Frisium®)	Normal exposure (CYP2C19 NM).	Normal efficacy*	Normal risk*
<b>Clonazepam</b> (Rivotril®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Clorazepate</b> (Tranxene®)	Initiate with recommended dose; a low dose may be adequate (CYP3A4 NM, UGT2B15 IM).	Normal efficacy*	Normal risk*


## Genetic Associations Identified

Medications	Exposure	Efficacy	Risk of atypical effect
<b>Diazepam</b> (Valium®)	Normal exposure (CYP2C19 NM, CYP3A4 NM, UGT2B15 IM).	Normal efficacy*	Normal risk*
<b>Flurazepam</b> (Dalmane®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Hydroxyzine</b> (Atarax®)	Normal exposure (CYP3A4 NM, CYP3A5 PM).	Normal efficacy*	Normal risk*
<b>Lorazepam</b> (Ativan®)	Initiate with recommended dose; a low dose may be adequate (UGT2B15 IM).	Normal efficacy*	Normal risk*
<b>Midazolam</b> (Versed®)	Normal exposure (CYP2C19 NM, CYP3A5 PM, CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Nitrazepam</b> (Mogadon®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Oxazepam</b> (Serax®)	Initiate with recommended dose; a low dose may be adequate (UGT2B15 IM).	Normal efficacy*	Normal risk*
<b>Temazepam</b> (Restoril®)	Initiate with recommended dose; a low dose may be adequate (UGT2B15 IM).	Normal efficacy*	Normal risk*
<b>Bromazepam</b> (Lectopam®)	 Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind).	Normal efficacy*	Normal risk*

## Central alpha-adrenergic agonists

<b>Clonidine</b> (Catapres®)	Normal exposure (CYP2D6 NM).	1/1 variant: increased likelihood of a poorer response (GNB3).	Normal risk*
<b>Guanfacine</b> (Intuniv XR®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*

## Mood Stabilizers

<b>Carbamazepine</b> (Tegretol®)	Normal exposure (CYP3A4 NM, CYP3A5 PM, UGT2B7 NM, ABCB1).	Normal efficacy*	HLA-A*31:01 negative, HLA-B*15:02 negative - Normal risks of cutaneous adverse reactions.
<b>Gabapentin</b> (Neurontin®)	Normal exposure (ABCB1).	Normal efficacy*	Normal risk*
<b>Lamotrigine</b> (Lamictal®)	Normal exposure (ABCG2, UGT2B7 NM).	Normal efficacy*	HLA-B*15:02 negative - Normal risks of cutaneous adverse reactions.
<b>Levetiracetam</b> (Keppra®)	Normal exposure (ABCB1).	Normal efficacy*	Normal risk*
<b>Lithium</b> (Carbolith®)	Normal exposure*	0/1 variant: no increased likelihood of a poorer response (CACNG2); 1/1 variant: increased likelihood of relapse after successful lithium therapy (lncRNA) - for bipolar disorder.	Normal risk*
<b>Oxcarbazepine</b> (Trileptal®)	Normal exposure (UGT2B7 NM).	Normal efficacy*	HLA-B*15:02 negative - Normal risks of cutaneous adverse reactions.
<b>Phenytoin</b> (Dilantin®)	Normal exposure (CYP2C9 NM).	Normal efficacy*	HLA-B*15:02 negative - normal risks of cutaneous adverse reactions.
<b>Pregabalin</b> (Lyrica®)	Normal exposure*	Normal efficacy*	Normal risk*
<b>Topiramate</b> (Topamax®)	Genetic influence not available	1/1 variant: increased likelihood of a poorer response for treatment of alcohol-related disorders (GRIK1).	Normal risk*
<b>Valproic acid, Divalproex</b> (Depakene®, Epival®)	Normal exposure (CYP2A6 NM, CYP2C9 NM).	Normal efficacy*	 1/1 variant: increased likelihood of weight gain (ANKK1).

## Norepinephrine Reuptake Inhibitor



Genetic Associations Identified			
Medications	Exposure	Efficacy	Risk of atypical effect
Atomoxetine (Strattera®)	Normal exposure (CYP2D6 NM). <sup>6,7</sup>	Genetic influence not available	Genetic influence not available
Psychostimulants			
Amphetamine / Dextroamphetamine (Adderall XR®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*
Dextroamphetamine (Dexedrine®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*
Lisdexamfetamine (Vyvanse®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*
Methylphenidate - Biphentin®	Normal exposure (CES1 NM).	3/5 variants: increased likelihood of a poorer response (ADRA2A, SLC6A2a, SLC6A2b).	Normal risk*
Methylphenidate - Concerta®	Normal exposure (CES1 NM).	3/5 variants: increased likelihood of a poorer response (ADRA2A, SLC6A2a, SLC6A2b).	Normal risk*
Methylphenidate - Focquest®	Normal exposure (CES1 NM).	3/5 variants: increased likelihood of a poorer response (ADRA2A, SLC6A2a, SLC6A2b).	Normal risk*
Methylphenidate - Quillivant®	Normal exposure (CES1 NM).	3/5 variants: increased likelihood of a poorer response (ADRA2A, SLC6A2a, SLC6A2b).	Normal risk*
Methylphenidate - Ritalin®	Normal exposure (CES1 NM).	3/5 variants: increased likelihood of a poorer response (ADRA2A, SLC6A2a, SLC6A2b).	Normal risk*
Sedative-Hypnotics			
Daridorexant (Quviviq®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
Diphenhydramine (Benadryl®)	Normal exposure*	Normal efficacy*	Normal risk*
Eszopiclone (Lunesta®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
Lemborexant (Dayvigo®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
Melatonin	Normal exposure*	Normal efficacy*	Normal risk*
Phenobarbital (Phenobarb®)	Normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
Triazolam (Halcion®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
Zolpidem (Sublinox®)	Normal exposure (CYP3A4 NM, CYP2C9 NM).	Normal efficacy*	Normal risk*
Zopiclone (Imovane®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
Wakefulness-promoting agents			
Modafinil (Alertec®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
Pitolisant (Wakix®)	Normal exposure (CYP2D6 NM, CYP3A4 NM).	Normal efficacy*	Normal risk*
Sodium oxybate (Xyrem®)	Normal exposure*	Normal efficacy*	Normal risk*
Solriamfetol (Sunosi®)	Normal exposure*	Normal efficacy*	Normal risk*



PGx RECOMMENDATIONS - PAIN MANAGEMENT

- ↑

Dose increase may be required due to lower exposure to active metabolite(s) with standard dosing.

✓

Increased probability of a better response.
- ↓

Dose reduction may be required due to higher exposure to active metabolite(s) with standard dosing.

!

Greater potential for a poorer response or atypical effect.
- 📉

Exposure is difficult to predict, insufficient data to calculate dose adjustments.

✗

Medication not recommended by peer-reviewed guidelines.

Normal exposure\*, Normal efficacy\* or Normal risk\*: Based on currently available genetic data, the efficacy or risk of an atypical effect is likely similar to that of most other individuals; further research is needed to better understand genetic influence.


Genetic Associations Identified			
Medications	Exposure	Efficacy	Risk of atypical effect
Analgesic			
Acetaminophen (Tylenol®)	Normal exposure*	Normal efficacy*	Normal risk*
Antimetabolite			
Methotrexate	↓ Consider using a lower dose (ABCB1, MTHFR).	Normal efficacy*	Normal risk*
Cannabinoids			
Cannabidiol (CBD)	Normal exposure (CYP3A4 NM, CYP2C9 NM).	Normal efficacy*	Normal risk*
Nabilone (Cesamet®)	Normal exposure*	Normal efficacy*	Normal risk*
Tetrahydrocannabinol (THC)	Normal exposure (CYP3A4 NM, CYP2C9 NM).	Normal efficacy*	! 1/2 variants: increased risk of cannabis use disorder (CNR1).
Muscle Relaxant			
Carisoprodol (Soma®)	Normal exposure (CYP2C19 NM).	Normal efficacy*	Normal risk*
Methocarbamol (Robaxin®)	Normal exposure*	Normal efficacy*	Normal risk*
Cyclobenzaprine (Flexeril®)	↑ Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP3A4 NM, CYP1A2 Ind).	Normal efficacy*	Normal risk*
Tizanidine (Zanaflex®)	↑ Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind).	Normal efficacy*	Normal risk*
Nonsteroidal Anti-Inflammatory Drugs (NSAID)			
Acetylsalicylic acid (Aspirin®)	Normal exposure*	Normal efficacy*	Normal risk*
Celecoxib (Celebrex®)	Normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
Diclofenac (Voltaren®)	Normal exposure (UGT2B7 NM).	Normal efficacy*	Normal risk*
Etodolac (Ultradol®)	Normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
Flurbiprofen (Ansaid®)	Normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*


## Genetic Associations Identified


Medications	Exposure	Efficacy	Risk of atypical effect
<b>Ibuprofen</b> (Advil®)	Normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
<b>Indomethacin</b> (Indocid®)	Normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
<b>Ketorolac</b> (Toradol®)	Normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
<b>Meloxicam</b> (Mobicox®)	Normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
<b>Naproxen</b> (Naprosyn®)	Normal exposure*	Normal efficacy*	Normal risk*
<b>Piroxicam</b> (Feldene®)	Initiate with recommended dose; normal exposure (CYP2C9 MM).	Normal efficacy*	Normal risk*
<b>Tenoxicam</b> (Mobiflex®)	Initiate with recommended dose; normal exposure (CYP2C9 MM).	Normal efficacy*	Normal risk*
<b>Nabumetone</b> (Relafen®)	↓ Initiate with recommended dose but monitor tolerance more closely; may require a lower dose., especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind, CYP2C9 NM).	Normal efficacy*	Normal risk*
Opioids			
<b>Buprenorphine</b> (Butrans®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Butorphanol</b> (Stadol®)	Normal exposure*	Normal efficacy*	Normal risk*
<b>Fentanyl</b> (Duragesic®)	Normal exposure (CYP3A4 NM, CYP3A5 PM).	Normal efficacy*	Normal risk*
<b>Nalbuphine</b> (Nubain®)	Normal exposure*	Normal efficacy*	Normal risk*
<b>Remifentanyl</b> (Ultiva®)	Normal exposure*	Normal efficacy*	Normal risk*
<b>Sufentanil</b> (Sufenta®)	Normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Tapentadol</b> (Nucynta®)	Normal exposure*	Normal efficacy*	Normal risk*
<b>Meperidine</b> (Demerol®)	↓ Consider using a lower dose (CYP2B6 IM, POR, CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Methadone</b>	↓ Consider using a lower dose (CYP2B6 IM, POR, CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Codeine</b>	Normal exposure (CYP2D6 NM, UGT2B7 NM).	! 1/1 variant: increased likelihood of a poorer response (OPRM1).	Normal risk*
<b>Hydrocodone</b> (Hycodan®)	Normal exposure (CYP2D6 NM).	! 1/1 variant: increased likelihood of a poorer response (OPRM1).	Normal risk*
<b>Hydromorphone</b> (Dilaudid®)	Normal exposure*	! 1/1 variant associated with an increased likelihood of a poorer response (OPRM1).	Normal risk*
<b>Morphine</b> (Statex®)	Normal exposure (UGT2B7 NM).	! 1/1 variant: increased likelihood of a poorer response (OPRM1).	0/1 variant: no increased risk of gastrointestinal side effects (FAAH).
<b>Oxycodone</b> (Supeudol®)	Normal exposure (CYP3A4 NM, CYP2D6 NM).	! 1/1 variant: increased likelihood of a poorer response (OPRM1).	Normal risk*
<b>Tramadol</b> (Ultram®)	Normal exposure (CYP2D6 NM).	! 1/1 variant: increased likelihood of a poorer response (OPRM1).	Normal risk*
Opioid antagonists			
<b>Naloxone</b> (Narcan®)	Normal exposure*	0/1 variant: no increased likelihood of a poorer response.	Normal risk*


Genetic Associations Identified			
Medications	Exposure	Efficacy	Risk of atypical effect
Naltrexone (Revia®)	Normal exposure*	1/1 variant: increased likelihood of a poorer response when used in combination with bupropion for weight loss (ANKK1).	Normal risk*


PGx RECOMMENDATIONS - CARDIOLOGY


- 

Dose increase may be required due to lower exposure to active metabolite(s) with standard dosing.
- 

Increased probability of a better response.
- 


Dose reduction may be required due to higher exposure to active metabolite(s) with standard dosing.
- 

Greater potential for a poorer response or atypical effect.
- 

Exposure is difficult to predict, insufficient data to calculate dose adjustments.
- 


Medication not recommended by peer-reviewed guidelines.

Normal exposure\*, Normal efficacy\* or Normal risk\*: Based on currently available genetic data, the efficacy or risk of an atypical effect is likely similar to that of most other individuals; further research is needed to better understand genetic influence.

Genetic Associations Identified			
Medications	Exposure	Efficacy	Risk of atypical effect
Angiotensin-Converting Enzyme Inhibitors (ACEI)			
Benazepril (Lotensin®)	Initiate with recommended starting dose, normal exposure (CES1 NM).	Normal efficacy*	Normal risk*
Captopril (Capoten®)	Normal exposure*	Normal efficacy*	Normal risk*
Cilazapril (Inhibace®)	Initiate with recommended starting dose, normal exposure (CES1 NM).	Normal efficacy*	Normal risk*
Fosinopril (Monopril®)	Initiate with recommended starting dose, normal exposure (CES1 NM).	Normal efficacy*	Normal risk*
Lisinopril (Prinivil/Zestril®)	Normal exposure*	Normal efficacy*	Normal risk*
Perindopril (Coversyl®)	Initiate with recommended starting dose, normal exposure (CES1 NM).	Normal efficacy*	Normal risk*
Quinapril (Accupril®)	Initiate with recommended starting dose, normal exposure (CES1 NM).	Normal efficacy*	Normal risk*
Ramipril (Altace®)	Initiate with recommended starting dose, normal exposure (CES1 NM).	Normal efficacy*	Normal risk*
Trandolapril (Mavik®)	Initiate with recommended starting dose, normal exposure (CES1 NM).	Normal efficacy*	Normal risk*
Enalapril (Vasotec®)	<div> Initiate with recommended starting dose but monitor side effects and the risk of hypotension more closely (CES1 NM, SLCO1B1 poor function).</div>	Normal efficacy*	Normal risk*
Antiarrhythmics			
Amiodarone (Pacerone®/ Cordarone®)	Initiate therapy with recommended starting dose (CYP3A4 NM).	Normal efficacy*	Normal risk*
Digoxin (Lanoxin®)	Initiate with recommended dose; a low dose may be adequate (ABCB1).	Normal efficacy*	Normal risk*
Disopyramide (Rythmodan®)	Initiate therapy with recommended starting dose (CYP3A4 NM).	Normal efficacy*	Normal risk*
Flecainide (Tambocor®)	Initiate therapy with recommended starting dose, normal exposure (CYP2D6 NM). <sup>8</sup>	Normal efficacy*	Normal risk*
Propafenone (Rythmol®)	Initiate therapy with recommended starting dose, normal exposure (CYP2D6 NM). <sup>8</sup>	Normal efficacy*	Normal risk*
Anticoagulants			

Genetic Associations Identified			
Medications	Exposure	Efficacy	Risk of atypical effect
Acenocoumarol- Nicoumalone (Sintrom®)	Initiate with recommended dose, normal exposure (VKORC1 -1639 GG). <sup>8</sup>	Normal efficacy*	Normal risk*
Apixaban (Eliquis®)	Initiate with recommended starting dose, normal exposure (CYP3A4 NM, ABCB1).	Normal efficacy*	Normal risk*
Dabigatran (Pradaxa®)	Initiate with recommended starting dose (CES1 NM, UGT2B15 IM).	Normal efficacy*	Normal risk*
Edoxaban (Lixiana®/Savaysa®)	⚠ Initiate with recommended starting dose but monitor more closely for potential bleeding risks (SLCO1B1 PA).	Normal efficacy*	Normal risk*
Rivaroxaban (Xarelto®)	⚠ Initiate with recommended starting dose but monitor more closely for potential bleeding risks (ABCB1).	Normal efficacy*	Normal risk*
Warfarin (Coumadin®)	⚠ A dose adjustment may be required; go to warfarindosing.org to calculate the average recommended dose using a validated pharmacogenetic algorithm (CYP2C9 NM, VKORC1 -1639 GG, CYP4F2 TT, rs12777823 GG). <sup>9</sup>	Normal efficacy*	Normal risk*
Antiplatelets			
Clopidogrel (Plavix®)	Initiate with recommended starting dose, normal exposure (CYP2C19 NM). <sup>10</sup>	Normal efficacy*	Normal risk*
Prasugrel (Effient®)	Normal exposure*	Normal efficacy*	Normal risk*
Ticagrelor (Brilinta®)	Normal exposure*	Normal efficacy*	Normal risk*
Angiotensin II Receptor Blockers (ARBs)			
Azilsartan (Edarbi®)	Initiate with recommended starting dose, normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
Irbesartan (Avapro®)	Initiate with recommended starting dose, normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
Losartan (Cozaar®)	Initiate with recommended starting dose, normal exposure (CYP2C9 NM).	Normal efficacy*	Normal risk*
Olmesartan (Olmotec®/Benicar®)	Normal exposure*	Normal efficacy*	Normal risk*
Telmisartan (Micardis®)	Normal exposure*	Normal efficacy*	Normal risk*
Valsartan (Diovan®)	Initiate with recommended starting dose, normal exposure (ABCB1).	Normal efficacy*	Normal risk*
Beta-blockers			
Acebutolol (Sectral®/Monitan®)	Initiate with recommended dose; normal exposure (CYP2C19 NM).	Normal efficacy*	Normal risk*
Atenolol (Tenormin®)	Normal exposure; eliminated largely unchanged by the kidneys.	Normal efficacy*	Normal risk*
Bisoprolol (Monocor®/Zebeta®)	Initiate with recommended starting dose, normal exposure (CYP2D6 NM, CYP3A4 NM, CYP3A5 PM).	Normal efficacy*	Normal risk*
Carvedilol (Coreg®)	Initiate with recommended starting dose, normal exposure (CYP2D6 NM, CYP2C9 NM, UGT1A1 NM).	Normal efficacy*	Normal risk*
Labetalol (Trandate®)	CYP2C19 NM - Initiate with recommended dose; normal exposure.	Normal efficacy*	Normal risk*


## Genetic Associations Identified

Medications	Exposure	Efficacy	Risk of atypical effect
<b>Metoprolol</b> (Lopresor/Betaloc®/ Toprol XL®)	Initiate with recommended starting dose, normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*
<b>Nadolol</b> (Corgard®)	Initiate with recommended starting dose but monitor bradycardia more closely; a low dose may be adequate (ABCB1).	Normal efficacy*	Normal risk*
<b>Sotalol</b> (Sotacor®/Betapace®)	Normal exposure; eliminated largely unchanged by the kidneys.	Normal efficacy*	Normal risk*
<b>Propranolol</b> (Inderal®)	 Initiate therapy with recommended starting dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke (CYP2D6 NM, CYP1A2 Ind).	Normal efficacy*	Normal risk*

## Calcium channel blockers (CCBs)

<b>Amlodipine</b> (Norvasc®)	Initiate with recommended starting dose, normal exposure (CYP3A4 NM, CYP3A5 PM).	Normal efficacy*	Normal risk*
<b>Diltiazem</b> (Cardizem®/Tiazac®)	Initiate with recommended starting dose, normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Felodipine</b> (Plendil®)	Initiate with recommended starting dose, normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Nifedipine</b> (Adalat/Procardia®)	Initiate with recommended starting dose, normal exposure (CYP3A4 NM, CYP3A5 PM).	Normal efficacy*	Normal risk*
<b>Verapamil</b> (Isoptin®/Verelan®/ Calan®)	Initiate with recommended starting dose, normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*

## Diuretics

<b>Acetazolamide</b> (Diamox®)	Normal exposure; eliminated largely unchanged by the kidneys.	Normal efficacy*	Normal risk*
<b>Amiloride</b> (Midamor®)	Normal exposure; eliminated largely unchanged by the kidneys.	Normal efficacy*	Normal risk*
<b>Chlorthalidone</b> (Hygroton®)	Normal exposure; eliminated largely unchanged by the kidneys.	Normal efficacy*	Normal risk*
<b>Eplerenone</b> (Inspra®)	Initiate with recommended starting dose, normal exposure (CYP3A4 NM, CYP3A5 PM).	Normal efficacy*	Normal risk*
<b>Furosemide</b> (Lasix®)	Normal exposure*	Normal efficacy*	Normal risk*
<b>Hydrochlorothiazide (HCTZ)</b> (Hydrodiuril®/ Microzide®)	Normal exposure; eliminated largely unchanged by the kidneys.	Normal efficacy*	Normal risk*
<b>Indapamide</b> (Iozide®/Lozol®)	Initiate with recommended starting dose, normal exposure (CYP3A4 NM).	Normal efficacy*	Normal risk*
<b>Spironolactone</b> (Aldactone®)	Normal exposure; eliminated largely unchanged by the kidneys.	Normal efficacy*	Normal risk*
<b>Triamterene</b> (Dyrenium®)	 Initiate with recommended starting dose but monitor efficacy more closely, especially with CYP1A2 inducers, such as smoke (CYP1A2 Ind).	Normal efficacy*	Normal risk*


## Lipid-lowering agent


<b>Fenofibrate</b> (Lipidil®/Tricor®)	Normal exposure*	Normal efficacy*	Normal risk*
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Genetic Associations Identified			
Medications	Exposure	Efficacy	Risk of atypical effect
Selective cholesterol-absorption inhibitor			
<b>Ezetimibe</b> (Ezetrol®/Zetia®)	Normal exposure*	Normal efficacy*	Normal risk*
Statins			
<b>Atorvastatin</b> (Lipitor®/Caduet®)	↓ Prescribe ≤20 mg as a starting dose and adjust doses based on disease-specific guidelines. If dose >20 mg is needed for desired efficacy, consider rosuvastatin or combination therapy (i.e., atorvastatin plus nonstatin guideline-directed medical therapy) (SLCO1B1 poor function). <sup>11</sup>	Normal efficacy*	Normal risk*
<b>Fluvastatin</b> (Lescol®)	↓ Prescribe ≤40 mg per day as a starting dose and adjust doses based on disease-specific guidelines. If patient is tolerating 40 mg per day but higher potency is needed, a higher dose (>40 mg) or an alternative statin or combination therapy (i.e., fluvastatin plus nonstatin guideline-directed medical therapy) could be considered. Be aware of possible increased risk for myopathy with fluvastatin especially with doses >40 mg per day (SLCO1B1 poor function, CYP2C9 NM). <sup>11</sup>	Normal efficacy*	Normal risk*
<b>Pitavastatin</b> (Livalo®)	↓ Prescribe ≤1 mg as a starting dose and adjust doses based on disease-specific guidelines. If dose >1 mg needed for desired efficacy, consider an alternative statin or combination therapy (i.e., pitavastatin plus nonstatin guideline-directed medical therapy) (SLCO1B1 poor function). <sup>11</sup>	Normal efficacy*	Normal risk*
<b>Pravastatin</b> (Pravachol®)	↓ Prescribe ≤40 mg as a starting dose and adjust doses based on disease-specific guidelines. If patient is tolerating 40-mg dose but higher potency is needed, a higher dose (>40 mg) or an alternative statin or combination therapy (i.e., pravastatin plus nonstatin guideline-directed medical therapy) could be considered. Be aware of possible increased risk for myopathy especially with pravastatin doses >40 mg (SLCO1B1 poor function). <sup>11</sup>	Normal efficacy*	Normal risk*
<b>Rosuvastatin</b> (Crestor®)	↓ Prescribe ≤20 mg as a starting dose and adjust doses based on disease-specific and population-specific guidelines. If dose >20 mg needed for desired efficacy, consider combination therapy (i.e., rosuvastatin plus nonstatin guideline-directed medical therapy) (SLCO1B1 poor function, ABCG2 normal function). <sup>5</sup>	Normal efficacy*	Normal risk*
<b>Lovastatin</b> (Mevacor®)	✗ Prescribe an alternative statin depending on the desired potency (SLCO1B1 poor function). <sup>11</sup>	Normal efficacy*	Normal risk*
<b>Simvastatin</b> (Zocor®)	✗ Prescribe an alternative statin depending on the desired potency (SLCO1B1 poor function). <sup>11</sup>	Normal efficacy*	Normal risk*





PGx RECOMMENDATIONS - COMPLEMENTARY TREATMENTS


- 

Dose increase may be required due to lower exposure to active metabolite(s) with standard dosing.
- 

Increased probability of a better response.
- 

Dose reduction may be required due to higher exposure to active metabolite(s) with standard dosing.
- 

Greater potential for a poorer response or atypical effect.
- 

Exposure is difficult to predict, insufficient data to calculate dose adjustments.
- 

Medication not recommended by peer-reviewed guidelines.

Normal exposure\*, Normal efficacy\* or Normal risk\*: Based on currently available genetic data, the efficacy or risk of an atypical effect is likely similar to that of most other individuals; further research is needed to better understand genetic influence.

Genetic Associations Identified			
Medications	Exposure	Efficacy	Risk of atypical effect
Antiemetics			
Dimenhydrinate (Gravol®)	Normal exposure*	Normal efficacy*	Normal risk*
Granisetron (Kytril®)	Normal exposure (CYP3A4 NM, CYP3A5 PM).	Normal efficacy*	Normal risk*
Ondansetron (Zofran®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*
Palonosetron (Aloxi®)	Normal exposure (CYP2D6 NM).	Normal efficacy*	Normal risk*
Proton pump inhibitors (PPI)			
Dexlansoprazole (Dexilant®)	Initiate therapy with standard dose but consider increasing the dose by 50-100% for the treatment of H. pylori infection and erosive esophagitis. Daily dose may be given in divided doses. Monitor for efficacy (CYP2C19 NM). <sup>12</sup>	Normal efficacy*	Normal risk*
Esomeprazole (Nexium®)	Normal exposure*	Normal efficacy*	Normal risk*
Lansoprazole (Prevacid®)	Initiate therapy with standard dose but consider increasing the dose by 50-100% for the treatment of H. pylori infection and erosive esophagitis. Daily dose may be given in divided doses. Monitor for efficacy (CYP2C19 NM). <sup>12</sup>	Normal efficacy*	Normal risk*
Omeprazole (Losec®)	Initiate therapy with standard dose but consider increasing the dose by 50-100% for the treatment of H. pylori infection and erosive esophagitis. Daily dose may be given in divided doses. Monitor for efficacy (CYP2C19 NM). <sup>12</sup>	Normal efficacy*	Normal risk*
Pantoprazole (Pantoloc®)	Initiate therapy with standard dose but consider increasing the dose by 50-100% for the treatment of H. pylori infection and erosive esophagitis. Daily dose may be given in divided doses. Monitor for efficacy (CYP2C19 NM). <sup>12</sup>	Normal efficacy*	Normal risk*

PGx ASSOCIATIONS - EXPOSURE

Enzymes	Phenotype					Genotype	Details
	PM	IM	NM	RM	UM		
CYTOCHROME P450 (CYP450)							
CYP4F2	<div></div>					*3/*3	<b>Poor metabolizer (PM)</b> Total loss of CYP4F2 function.
CYP3A5	<div></div>					*3/*3	<b>Poor metabolizer (PM)</b> Low CYP3A5 activity is the most common phenotype and has been used to establish standard doses.
CYP2B6		<div></div>				*1/*6 or *4/*9	<b>Intermediate metabolizer (IM)</b> Reduced CYP2B6 activity compared to normal metabolizers.
CYP2A6			<div></div>			*1/*1	<b>Normal metabolizer (NM)</b> Normal CYP2A6 activity.
CYP2C9			<div></div>			*1/*1	<b>Normal metabolizer (NM)</b> Normal CYP2C9 activity.
CYP2C19			<div></div>			*1/*1	<b>Normal metabolizer (NM)</b> Normal CYP2C19 activity.
CYP2D6			<div></div>			*2/*41	<b>Normal metabolizer (NM)</b> Normal CYP2D6 activity.
CYP3A4			<div></div>			*1/*1	<b>Normal metabolizer (NM)</b> Normal CYP3A4 activity.
CYP1A2			<div></div>			*1F/*1F	<b>Inducible metabolizer (Ind)</b> Increased CYP1A2 activity compared to normal metabolizers if the patient is exposed to CYP1A2 inducers, such as smoke.

Enzymes	Phenotype					Genotype	Details
	PM	IM	NM	RM	UM		
OTHER METABOLIC ENZYMES AND FACTORS							
POR	<div></div>					A/A	<b>Poor activity (PA)</b> Greatly reduced cytochrome P450 oxidoreductase (POR) activity compared to normal phenotypes (POR participates in CYP2B6 activation).
SLCO1B1	<div></div>					*5/*5	<b>Poor function</b> Greatly reduced solute carrier organic anion transporter family member 1B1 (SLCO1B1) activity compared to a normal function.
TPMT	<div></div>					*3C/*3C	<b>Poor metabolizer (PM)</b> Greatly reduced thiopurine S-methyltransferase (TPMT) activity compared to normal metabolizers.
NUDT15	<div></div>					*3/*3	<b>Poor metabolizer (PM)</b> Greatly reduced or absent nudix hydrolase 15 (NUDT15) activity compared to normal metabolizers.
DPYD		<div></div>				c.868A>G/c.868A>G	<b>Intermediate metabolizer (IM)</b> Reduced dihydropyrimidine dehydrogenase (DPYD) activity compared to normal metabolizers.
CES1			<div></div>			C/C	<b>Normal metabolizer (NM)</b> Normal carboxylesterase-1 (CES1) activity.
UDP-GLUCURONOSYLTRANSFERASES (UGT)							
UGT2B15		<div></div>				*1/*2	<b>Intermediate metabolizer (IM)</b> Reduced UGT2B15 activity compared to normal metabolizers.
UGT1A1			<div></div>			*1/*1	<b>Normal metabolizer (NM)</b> Normal UGT1A1 activity.
UGT1A4			<div></div>			*1/*1	<b>Normal metabolizer (NM)</b> Normal UGT1A4 activity.

Enzymes		Phenotype					Genotype		Details	
		PM	IM	NM	RM	UM				
UGT2B7				<div></div>			*1/*1		Normal metabolizer (NM) Normal UGT2B7 activity.	

ANALYTICAL RESULTS

The following analytical results were used to generate the pharmacogenomic interpretations found in this report. Technical limitations inherent with the methods used to produce these results may hinder the attribution of a definitive phenotype (see "TEST METHODOLOGY AND LIMITATIONS").

Genes	Variant Details (GRCH38.p12)		Result
ABCB1	rs1045642	chr7:87509329	A G
	rs2032582	chr7:87531302	A C
	rs2032583	chr7:87531245	C T
ABCG2	rs2231142	chr4:88131171	G G
ADRA2A	rs1800544	chr10:111076745	C C
ANKK1	rs1800497	chr11:113400106	G G
BDNF	rs6265	chr11:27658369	C C
CACNG2	rs2283967	chr22:36567486	C C
CES1	rs71647871	chr16:55823658	C C
CNR1	rs806380	chr6:88154934	A A
COMT	rs4680	chr22:19963748	G G
CYP1A2	rs762551	chr15:74749576	A A
	rs2069514	chr15:74745879	G G
CYP2A6	rs1801272	chr19:40848628	A A
	rs28399433	chr19:40850474	A A
CYP2B6	rs2279343	chr19:41009358	G A
	rs3745274	chr19:41006936	G T
	rs28399499	chr19:41012316	T T
CYP2C cluster	rs12777823	chr10:94645745	1 4
CYP2C9	rs1057910	chr10:94981296	A A
	rs1799853	chr10:94942290	C C
	rs7900194	chr10:94942309	G G
	rs9332131	chr10:94949282-94949283	A A
	rs9332239		C C
	rs28371685	chr10:94981224	C C
	rs28371686	chr10:94981301	C C
	rs72558187	chr10:94941958	T T
	rs72558190	chr10:94947782	C C
	rs4244285	chr10:94781859	G G
	rs4986893	chr10:94780653	G G
	rs6413438	chr10:94781858	C C
CYP2C19	rs12248560	chr10:94761900	C C
	rs12769205	chr10:94775367	A A
	rs17884712	chr10:94775489	G G
	rs28399504	chr10:94762706	A A
	rs41291556	chr10:94775416	T T
	rs56337013	chr10:94852738	C C
	rs72552267	chr10:94775453	G G
	rs72558186	chr10:94781999	T T
	rs16947	chr22:42127941	A A
	rs1065852	chr22:42130692	G G
	rs1135840	chr22:42126611	G G
	rs3892097	chr22:42128945	C C
CYP2D6	rs5030655	chr22:42129084	A A
	rs5030656	chr22:42128174-42128178	A A
	rs5030862		C C
	rs5030865	chr22:42129033	C C
	rs5030867	chr22:42127856	T T
	rs28371725	chr22:42127803	C T
	rs28371706	chr22:42129770	G G
	rs35742686	chr22:42128242	T T
	rs59421388	chr22:42127608	C C
	rs774671100	chr22:42130555-42130755	G G
	rs201377835		C C
	Gene Deletion	n/a	Not Detected
	Gene Duplication	n/a	Not Detected
CYP3A4	rs4986907	chr7:99769804	C C
	rs35599367	chr7:99768693	G G
	rs55785340	chr7:99768360	A A
	rs67666821	chr7:99758184-99758188	D D
	rs72552799		C C

Genes	Variant Details (GRCH38.p12)		Result
CYP3A5	rs776746	chr7:99672916	C C
	rs10264272	chr7:99665212	C C
	rs41303343	chr7:99652771	D D
CYP4F2	rs2108622	chr19:15879621	1 2
DPYD	rs75017182	chr1:97579893	2 2
	rs55886062	chr1:97515787	1 1
	rs3918290	chr1:97450058	0 0
	rs112766203	chr1:97305279	5 5
	rs67376798	chr1:97082391	6 6
	rs115232898	chr1:97699474	3 3
	rs146356975	chr1:97595149	4 4
DRD2	rs6275	chr11:113412755	G G
DRD3	rs963468	chr3:114144040	A G
FAAH	rs324420	chr1:46405089	C C
FKBP5	rs4713916	chr6:35702206	G G
GNB3	rs5443	chr12:6845711	C C
GRIK1	rs2832407	chr21:29595188	A C
GRIK4	rs1954787	chr11:120792654	C T
HLA-A*31:01	rs1061235	chr6:29945521	A A
HLA-B*15:02	rs144012689	chr6:31355003	T T
HTR2A	rs6311	chr13:46897343	C C
	rs6313	chr13:46895805	G G
	rs2770296	chr13:46866425	C C
HTR2C	rs3813929	chrX:114584047	C T
HTR7	rs7905446	chr10:90859404	G T
INSIG2	rs17047764	chr2:118111006	C G
long non-coding (lnc) RNA	rs74795342	chr21:18954018	G G
MC4R	rs489693	chr18:60215554	A C
	rs17782313	chr18:60183864	T T
MTHFR	rs1801131	chr1:11794419	G T
	rs1801133	chr1:11796321	A G
NUDT15	rs116855232	chr13:48045719	1 0
OPRM1	rs1799971	chr6:154039662	A G
POR	rs2868177	chr7:75960585	A A
SLC6A2	rs5569	chr16:55697923	A G
	rs2242446	chr16:55656513	T T
	rs28386840	chr16:55652906	A A
SLC6A4	5-HTTLPR	chr17:30190154-30240133	S L
SLC6A5	rs2298826	chr11:20638211	G G
SLCO1B1	rs4149056	chr12:21178615	1 1
TH	rs2070762	chr11:2165105	A G
TPH2	rs1487278	chr12:72007071	C T
TPMT	rs1800462	chr6:18143724	7 7
	rs1800460	chr6:18138997	8 8
	rs1142345	chr6:18130687	9 9
UGT1A1	rs4148323	chr2:233760498	G G
	rs34815109	chr2:233760234-233760248	6 6
UGT1A4	rs2011425	chr2:233718962	T T
UGT2B7	rs7439366	chr4:69098620	T T
UGT2B15	rs1902023	chr4:68670366	C A
VKORC1	rs9923231	chr16:31096368	1 3

## TEST METHODOLOGY AND LIMITATIONS

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The Biron pharmacogenomic test for psychiatry and pain management is a MALDI-TOF-based single nucleotide primer extension genotyping test; laboratory developed and validated test (LDT), not approved by Health Canada. Nucleic acid amplification techniques may be subject to general interference by factors such as reaction inhibitors and low quality or quantity of extracted DNA. Factors influencing the amount and quality of extracted DNA include but are not limited to patient oral hygiene, collection technique and presence of dietary or microbial source of nucleic acids and nuclease. When present, these interferents typically yield no result rather than an inaccurate one. Risk of suboptimal DNA quantity or quality is significantly reduced by automated DNA extraction which uses chemistry without PCR inhibitors (magnetic beads) and systematic dilution, quantitation and normalization of DNA before nucleic acid amplification. Very infrequent variants or polymorphisms occurring in primer-binding regions may also affect testing and could produce an erroneous result or assay failure. The test does not detect all known and unknown variations in the genes tested, nor does absence of a detectable variant (typically reported as \*1 for metabolic enzymes) rule out the presence of other, non-detected variants. The test detects CYP2D6 deletion and duplication but cannot differentiate duplication in the presence of deletion. CYP2D6 deletion and duplication assays can translate into equivocal phenotype results where a range of enzyme activity level must be reported. Test results and clinical interpretation may be inaccurate for individuals who have undergone or are receiving non-autologous blood transfusions, tissue, and/or organ transplant therapies.

## DISCLAIMER

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Biron Health Group developed this pharmacogenomic report. This test does not diagnose any disorder, condition or disease. The interpretations and recommendations provided in this report are intended as a clinical support tool (DST) to be used solely by a healthcare professional. Treatment decisions for the patient remain the sole responsibility of the treating healthcare provider. The interpretations of the results provided by this report were determined by Biron's data curation protocol, which were established as per the current available scientific evidence available at the time this report version was created. As more evidence becomes available in the future, these interpretations may change. Some variants tested may not be used to provide report interpretations due to a lack of clear gene-drug association as determined by Biron's data curation protocol. The presence of a notification within the "Exposure", "Efficacy" or "Adverse Drug Reactions" categories for a given drug indicates that an associated genetic variant was detected. The lack of a notification within these categories for a given drug does not eliminate the requirement for dose adjustments for optimal dosage, does not guarantee effective drug therapy and does not eliminate the risks of adverse drug reactions. Commercial names are indicated as examples and do not consist an exhaustive list.

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For the full list of references, contact [pgxinfo@biron.com](mailto:pgxinfo@biron.com)

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