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.

Efficacy

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

The notification "Genetic influence not available" 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)

Exposure



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.

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

For any questions regarding this report, contact Michel Cameron, PhD:

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biron.com/pgx-healthcare



Risk of atypical effect

PHARMACOGENOMIC REPORT

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Psychiatry, ADHD and Pain Management

To download the latest version, go here: secur.biron.com/login.

DO NOT DIST RIBUT E. 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.

ADMINISTRATIVE DATA		
Patient Name Test-Firstname Test-Lastname	Ordering Clinician Meredith Grey	Sample ID: BIO2107270327 Sample Type: test
Gender: Female Date of birth: 1999-01-01 Phone Number: (418) 999-9999 Email: test-sample.BIO2107270327@biron.local	Patient Address 1212 some street Ste-foy, Québec G2J 4M5	Date ordered: 2022-08-21 Date of sample reception: 2025-01-08 Date of report: 2025-01-08
Clinical Support Email: genetique@biron.com	Phone: 1-855-943-6379	Fax: (514) 317-2241

VARIANT(S) OF VERY IMPORTANT PHARMACOGENES (VIPS)

CYP1A2 IND, CYP2C9 IM, CYP2C19 IM, POR IA, CES1 IM, UGT1A1 IM, UGT1A4 IM, UGT2B7 variable, UGT2B15 IM.

Very Important Pharmacogenes (VIPs) - visit pharmgkb.org/vips for more information. Abbreviations - 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.

CAUTION	CAUTIONARY INFORMATION					
Medication	Identified risk	Recommendation				
Carbamazepine Tegretol®	Risk of hypersensitivity reaction due to the presence of the HLA-B*15:02 allele.	If patient is carbamazepine-naïve, avoid carbamazepine. If the patient has previously used carbamazepine consistently for longer than three months without incidence of cutaneous adverse reactions, cautiously consider using carbamazepine. 1				
Oxcarbazepine Trileptal®	Risk of hypersensitivity reaction due to the presence of the HLA-B*15:02 allele.	If patient is oxcarbazepine-naïve, avoid oxcarbazepine. If the patient has previously used oxcarbazepine consistently for longer than three months without incidence of cutaneous adverse reactions, cautiously consider use of oxcarbazepine. ¹				
Phenytoin Dilantin®	Risk of hypersensitivity reaction due to the presence of the HLA-B*15:02 allele.	If patient is phenytoin-naïve, avoid phenytoin. If the patient has previously used phenytoin consistently for longer than three months without incidence of cutaneous adverse reactions, cautiously consider use of phenytoïn.				
Clopid ogrel Plavix®	Reduced clopidogrel active metabolite formation; increased on-treatment platelet reactivity; increased risk for adverse cardiac and cerebrovascular events (CYP2C19 IM).	Cardiovascular indications: avoid standard dose (75mg/day) if possible; use prasugrel or ticagrelor at standard dose if no contraindication. Neurovascular:				

CAUTIONARY INFORMATION

Medication	ldentified risk	Recommendation
		consider alternative P2Y12 inhibitor at standard dose if clinically indicated and no contraindication. ²

PGx RECOMMENDATIONS - PSYCHIATRY

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- Dose increase may be required due to lower exposure to active metabolite(s) with standard dosing.
- Dose reduction may be required due to higher exposure to active metabolite(s) with standard dosing.
- Medication not recommended by peer-reviewed guidelines.

Greater potential for a poorer response or atypical effect.

Increased probability of a better response.

- Exposure is difficult to predict, insufficient data to calculate dose adjustments.
- Genetic Associations Identified Medications Efficacy Risk of atypical effect Exposure **Antidepressants** Selective serotonin reuptake inhibitors (SSRIs) CYP2D6 NM, CYP2C9 IM - Initiate with 4/6 variants: increased likelihood of a Genetic influence not available poorer response (SLC6A4 - association (Prozac®) recommended dose; a low dose may be demonstrated mainly in Caucasians, adequate. not observed in Asians, BDNF, FKBP5, HTR7); consider adding L-methylfolate 15mg/day (MTHFR). CYP2D6 NM, ABCB1 - Initiate with Fluvoxamine 4/6 variants: increased likelihood of a Genetic influence not available poorer response (SLC6A4 - association (Luvox®) recommended dose but may require a demonstrated mainly in Caucasians, higher dose. not observed in Asians, BDNF, FKBP5, HTR7); consider adding L-methylfolate 15mg/day (MTHFR). **Paroxetine** CYP2D6 NM, ABCB1 - Initiate therapy with 4/6 variants: increased likelihood of a Genetic influence not available poorer response (SLC6A4 - association (Paxil®) recommended starting dose but may require a higher dose. demonstrated mainly in Caucasians, not observed in Asians, BDNF, FKBP5, HTR7); consider adding L-methylfolate 15mg/day (MTHFR). CYP2C19 IM - Initiate with Citalopram 4/6 variants: increased likelihood of a 1/1 variant: increased risk of (Celexa®) recommended dose but consider a poorer response (SLC6A4 - association gastrointestinal side effects (HTR2A). slower titration schedule and do not demonstrated mainly in Caucasians, exceed the following daily doses: not observed in Asians, BDNF, FKBP5, 30mg for adults up to 65 yrs; 15mg for HTR7); consider adding L-methylfolate 15mg/day (MTHFR). adults 65 yrs or older. 3,4 CYP2C19 IM - Initiate with Escitalopram 4/6 variants: increased likelihood of a 1/1 variant: increased risk of (Cipralex®) recommended dose but consider a poorer response (SLC6A4 - association gastrointestinal side effects (HTR2A). slower titration schedule and do not demonstrated mainly in Caucasians, exceed the following daily doses: not observed in Asians, BDNF, FKBP5, 15mg for adults up to 65 yrs; 7.5mg for HTR7); consider adding L-methylfolate adults 65 yrs or older. 3,4 15mg/day (MTHFR). CYP2B6 NM, CYP2C19 IM - Initiate Sertraline 4/5 variants: increased likelihood of a 1/1 variant: increased risk of therapy with recommended starting poorer response (SLC6A4 - association gastrointestinal side effects (HTR2A). (Zoloft®) dose but with a slower titration demonstrated mainly in Caucasians, not observed in Asians, BDNF, FKBP5, schedule and a lower maintenance HTR7); consider adding L-methylfolate dose.3 15mg/day (MTHFR). Serotonin-norepinephrine reuptake inhibitors (SNRIs) Desvenlafaxine UGT1A1 IM, UGT2B15 IM - Initiate with 1/2 variants: increased likelihood of a Genetic influence not available recommended dose; a low dose may be poorer response (FKBP5); consider adding (Pristiq®) L-methylfolate 15mg/day (MTHFR). adequate. Levomilnacipran CYP3A4 NM, ABCB1 - Initiate therapy with 1/2 variants: increased likelihood of a Genetic influence not available (Fetzima®) recommended starting dose but may poorer response (FKBP5); consider adding L-methylfolate 15mg/day (MTHFR). require a higher dose.

Monoamine oxidase inhibitors (MAOs)

CYP2D6 NM - Normal exposure^{5,7}

Atomoxetine

(Strattera®)

Genetic influence not available

Genetic influence not available

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	G	enetic Associations Identifi	ed
Medications	Exposure	Efficacy	Risk of atypical effect
Psychostimulants			
Amphetamine / Dextroamphetamine (Adderall XR®)	CYP2D6 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Dextroamphetamine (Dexed rine®)	CYP2D6 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Lisdexamfetamine (Vyvanse®)	CYP2D6 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Methylphenidate - Biphentin®	CES1 IM - Consider using a lower dose.	2/5 variants: increased likelihood of a poorer response (COMT, SLC6A2).	Genetic influence not available
Methylphenidate - Concerta®	CES1 IM - Consider using a lower dose.	2/5 variants: increased likelihood of a poorer response (COMT, SLC6A2).	Genetic influence not available
Methylphenidate - Foquest®	CES1 IM - Consider using a lower dose.	2/5 variants: increased likelihood of a poorer response (COMT, SLC6A2).	Genetic influence not available
Methylphenidate - Ritalin®	CES1 IM - Consider using a lower dose.	2/5 variants: increased likelihood of a poorer response (COMT, SLC6A2).	Genetic influence not available
Sedative-Hypnotic	es .		
Daridorexant (Quviviq®)	CYP3A4 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Diphenydramine (Benadryl®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Eszopicione (Lunesta®)	CYP3A4 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Lemborexant (Dayvigo®)	CYP3A4 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Melatonin	Genetic influence not available	Genetic influence not available	Genetic influence not available
Phenobarbital (Phenobarb®)	CYP2C9 IM - Initiate therapy with recommended starting dose; a low dose may be adequate.	Genetic influence not available	Genetic influence not available
Triazolam (Halcion®)	CYP3A4 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Zolpidem (Sublinox®)	CYP3A4 NM, CYP2C9 IM - Normal exposure	Genetic influence not available	Genetic influence not available
Zopiclone (Imovane®)	CYP3A4 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Wakefulness-prom	noting agents		
Modafinil (Alertec®)	CYP3A4 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Pitolisant (Wakix®)	CYP2D6 NM, CYP3A4 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Sodium oxybate (Xyrem®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Solriamfetol (Sunosi®)	Genetic influence not available	Genetic influence not available	Genetic influence not available

PGx RECOMMENDATIONS - PAIN MANAGEMENT

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

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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.

	G	enetic Associations Identific	ed
Medications	Exposure	Efficacy	Risk of atypical effect
Analgesic			
Acetaminophen (Tylenol®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Antimetabolite			
Methotrexate	ABCB1, MTHFR - Consider using a lower dose.	Genetic influence not available	Genetic influence not available
Cannabinoids			
Cannabidiol (CBD)	CYP3A4 NM, CYP2C9 IM - Normal exposure	Genetic influence not available	Genetic influence not available
Nabilone (Cesamet®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Tetrahydrocannabinol (THC)	CYP3A4 NM, CYP2C9 IM - Normal exposure	Genetic influence not available	1/2 variants: increased risk of cannabi use disorder (CNR1).
Muscle Relaxant			
Carisoprodol (Soma®)	CYP2C19 IM - Initiate therapy with recommended starting dose and use with caution; a low dose may be adequate.	Genetic influence not available	Genetic influence not available
Methocarbamol (Robaxin®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Cyclobenzaprine (Flexeril®)	CYP3A4 NM, CYP1A2 Ind - Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke.	Genetic influence not available	Genetic influence not available
Tizanidine (Zanaflex®)	CYP1A2 Ind - Initiate with recommended dose but may require a higher dose, especially with CYP1A2 inducers, such as smoke.	Genetic influence not available	Genetic influence not available
Nonsteroidal Anti-l	nflammatory Drugs (NSAID)		
Acetylsalicylic acid (Aspirin®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Diclofenac (Voltaren®)	UGT2B7 IM - Initiate with recommended dose; a low dose may be adequate.	Genetic influence not available	Genetic influence not available
Etodolac (Ultradol®)	CYP2C9 IM - Initiate with recommended dose; a low dose may be adequate.	Genetic influence not available	Genetic influence not available
Indomethacin (Indocid®)	CYP2C9 IM - Initiate with recommended dose; a low dose may be adequate.	Genetic influence not available	Genetic influence not available
Ketorolac (Toradol®)	CYP2C9 IM - Initiate with recommended dose; a low dose may be adequate.	Genetic influence not available	Genetic influence not available
Naproxen (Naprosyn®)	Genetic influence not available	Genetic influence not available	Genetic influence not available

poorer response.

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	G	enetic Associations Identifi	ed
Medications	Exposure	Efficacy	Risk of atypical effect
Hydromorphone (Dilaudid®)	Genetic influence not available	0/1 variant: no increased likelihood of a poorer response.	Genetic influence not available
Meperidine (Demerol®)	CYP2B6 NM, POR, CYP3A4 NM - Initiate therapy with recommended starting dose; a low dose may be adequate.	Genetic influence not available	Genetic influence not available
Methadone	CYP2B6 NM, CYP3A4 NM, POR - Initiate therapy with recommended starting dose; a low dose may be adequate.	Genetic influence not available	Genetic influence not available
Nalbuphine (Nubain®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Oxycodone (Supeudol®)	CYP3A4 NM, CYP2D6 NM - Normal exposure	0/1 variant: no increased likelihood of a poorer response.	Genetic influence not available
Remifentanil (Ultiva®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Sufentanil (Sufenta®)	CYP3A4 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Tapentadol (Nucynta®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Tramadol (Ultram®)	CYP2D6 NM - Normal exposure	0/1 variant: no increased likelihood of a poorer response.	Genetic influence not available
Codeine	CYP2D6 NM, UGT2B7 RM - Initiate with recommended dose but may require a higher dose.	0/1 variant: no increased likelihood of a poorer response.	Genetic influence not available
Morphine	UGT2B7 RM - Initiate with recommended dose but may require a higher dose.	0/1 variant: no increased likelihood of a poorer response.	0/1 variant: no increased risk of gastrointestinal side effects (FAAH).
Opioid antagonist	s		
Naltrexone (Revia®)	Genetic influence not available	1/1 variant: increased likelihood of a poorer response when used in combination with bupropion for weight loss (ANKK1).	Genetic influence not available
Naloxone (Narcan®)	Genetic influence not available	1/1 variant: increased likelihood of a poorer response (OPRM1).	Genetic influence not available

PGx RECOMMENDATIONS - COMPLEMENTARY TREATMENTS

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

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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.

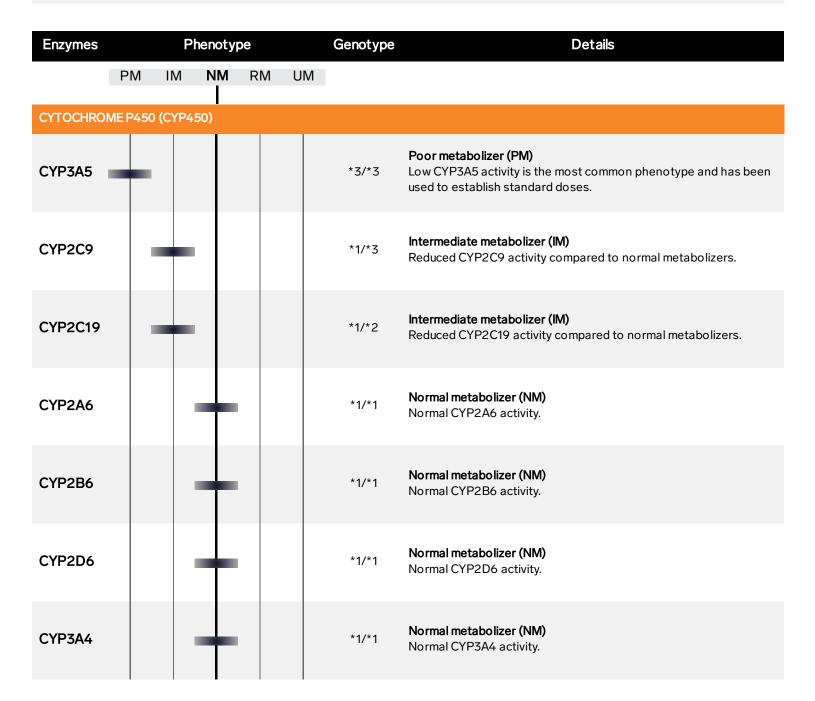
Exposure is difficult to predict, insufficient data to calculate dose adjustments.

Medication not recommended by peer-reviewed guidelines.

	G	enetic Associations Identif	ied
Medications	Exposure	Efficacy	Risk of atypical effect
Antiemetics			
Dimenhydrinate (Gravol®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Granisetron (Kytril®)	CYP3A4 NM, CYP3A5 PM - Normal exposure	Genetic influence not available	Genetic influence not available
Ondansetron (Zofran®)	CYP2D6 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Palonosetron (Aloxi®)	CYP2D6 NM - Normal exposure	Genetic influence not available	Genetic influence not available
Proton pump inhib	itors (PPI)		
Esomeprazole (Nexium®)	Genetic influence not available	Genetic influence not available	Genetic influence not available
Dexlansoprazole (Dexilant®)	CYP2C19 IM - Initiate therapy with standard dose but for chronic therapy (> 12 weeks) and efficacy achieved, consider reducing the daily dose by 50% and monitor for continued efficacy.	Genetic influence not available	Genetic influence not available
Lansoprazole (Prevacid®)	CYP2C19 IM - Initiate therapy with standard dose but for chronic therapy (> 12 weeks) and efficacy achieved, consider reducing the daily dose by 50% and monitor for continued efficacy.9	Genetic influence not available	Genetic influence not available
Omeprazole (Losec®)	CYP2C19 IM - Initiate therapy with standard dose but for chronic therapy (> 12 weeks) and efficacy achieved, consider reducing the daily dose by 50% and monitor for continued efficacy.	Genetic influence not available	Genetic influence not available
Pantoprazole (Pantoloc®)	CYP2C19 IM - Initiate therapy with standard dose but for chronic therapy (> 12 weeks) and efficacy achieved, consider reducing the daily dose by 50% and monitor for continued efficacy.	Genetic influence not available	Genetic influence not available

PGx ASSOCIATIONS - EXPOSURE

METABOLISM	The level of activit	The level of activity of the enzymes listed below influence the exposure to active metabolite(s).		
	CYP450	POR	CES	UGT
Variants detected / tested	3/8	1/1	1/1	4/4



DISTRIBUTION		The level of activity of the efflux pumps listed below influence the exposure to medications at the site of action (e.g., central nervous system).		
	ABCB1	ABCG2		
Variant(s) detected / tested	1/1	1/1		

Efflux Pump		Phenotype		Genotype	Details
	Poor activity	Intermediate activity	Normal activity		
EFFLUX PUN	/IPS (P-GLY	COPROTEIN, MU	JLT IDRUG RI	ESISTANCE PF	ROTEIN)
ABCB1				AAT/ACT	Poor activity (PA) Reduced efflux pump activity may result in higher substrate concentrations, increasing the likelihood of response but also increasing the risk of adverse drug reactions.
ABCG2		+		G/T	Intermediate activity (IA) Reduced efflux pump activity may result in higher substrate concentrations, increasing the likelihood of response but also increasing the risk of adverse drug reactions.

PGx ASSOCIATIONS - EFFICACY

The variants listed below have been associated with a higher likelihood of a poorer response to medications, compared to non-carriers.

Poorer response more likely

Variant(s) detected / tested

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Poore	Poorer response more likely		
Gene	Reference SNP cluster ID	Variant detected	Associated medication(s)
ANKK1	rs1800497	yes	aripiprazole, risperidone, naltrexone
BDNF	rs6265	yes	citalopram, escitalopram, sertraline, fluoxetine, fluvoxamine, paroxetine
COMT	rs4680	yes	venlafaxine (AA), methylphenidate (GG)
FKBP5	rs4713916	yes	citalopram, desvenlafaxine, duloxetine, escitalopram, levomilnacipran, sertraline, fluoxetine, fluoxamine, mirtazapine, paroxetine, venlafaxine
GNB3	rs5443	yes	clonidine
HTR7	rs7905446	yes	citalopram, escitalopram, sertraline, fluoxetine, fluvoxamine, paroxetine
OPRM1	rs1799971	yes	naloxone
SLC6A2	rs2242446	yes	venlafaxine
SLC6A2	rs28386840	yes	methylphenidate
SLC6A4	5-HTTLPR	yes	citalopram, escitalopram, sertraline, fluoxetine, fluvoxamine, paroxetine
TPH2	rs1487278	yes	amitriptyline, clomipramine, desipramine, doxepin, imipramine, mirtazapine, nortriptyline, trimipramine
ADRA2A	rs1800544	no	methylphenidate
ANKK1	rs1800497	no	bupropion
BDNF	rs6265	no	esketamine, ketamine
CACNG2	rs2283967	no	lithium
DRD3	rs963468	no	duloxetine
GRIK1	rs2832407	no	topiramate
GRIK4	rs1954787	no	citalopram, desvenlafaxine, duloxetine, escitalopram, fluoxetine, fluvoxamine, levomilnacipran, paroxetine, sertraline, venlafaxine

Gene	Reference SNP cluster ID	Variant detected	Associated medication(s)	
HTR2A	rs2770296	no	bupropion	
HTR2A	rs6311	no	citalopram, escitalopram, fluoxetine, fluvoxamine, paroxetine	
IncRNA	rs74795342	no	lithium	
IncRNA	rs75222709	no	lithium	
OPRM1	rs1799971	no	codeine, hydrocodone, hydromorphone, morphine, tramadol	
SLC6A2	rs5569	no	methylphenidate	
TH	rs2070762	no	methylphenidate	

PGx ASSOCIATIONS - ADVERSE DRUG REACTIONS

The variants listed below have been associated with a higher likelihood of specific adverse drug reactions, compared to non-carriers.

Increased risk of adverse drug reactions

Variant(s) detected / tested

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Increased risk of adverse drug reactions						
Gene	Reference SNP cluster ID	Variant detected	Associated medication(s)			
ANKK1	rs1800497	yes	weight gain: valproic acid			
CNR1	rs806380	yes	cannabis use disorder: THC			
HLA-B	*15:02	yes	cutaneous adverse reactions: carbamazepine, oxcarbazepine, phenytoin			
HTR2A	rs6311	yes	gastrointestinal side effects (nausea, vomiting): citalopram, escitalopram, sertraline			
HTR2C	rs3813929	yes	antipsychotic-induced weight gain: clozapine, olanzapine, risperidone			
MC4R	rs489693	yes	antipsychotic-induced weight gain: clozapine, olanzapine, quetiapine, risperidone			
ADRA2A	rs1800544	no	antipsychotic-induced weight gain: clozapine, olanzapine, risperidone			
DRD2	rs6275	no	antipsychotic-induced weight gain: clozapine, olanzapine, risperidone			
FAAH	rs324420	no	gastrointestinal side effects (nausea, vomiting): morphine (A allele); cannabis use disorder: THC (AA)			
GNB3	rs5443	no	antipsychotic-induced weight gain: clozapine, olanzapine, risperidone			
HLA-A	*31:01	no	cutaneous adverse reactions: carbamazepine			
INSIG2	rs17047764	no	antipsychotic-induced weight gain: clozapine, olanzapine, risperidone			
MC4R	rs17782313	no	antipsychotic-induced weight gain: quetiapine			
SLC6A2	rs28386840	no	hypertension: esketamine			
SLC6A5	rs2298826	no	extrapyramidal side effects: haloperidol			

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 Detai	ls (GRCH38.p12)	Result
ABCB1	rs1045642	chr7:87509329	A A
	rs2032582	chr7:87531302	AIC
	rs2032583	chr7:87531245	T T
ABCG2	rs2231142	chr4:88131171	G T
ADRA2A	rs1800544	chr10:111076745	C G
ANKK1	rs1800497	chr11:113400106	GIG
BDNF	rs6265	chr11:27658369	CIC
CACNG2	rs2283967	chr22:36567486	CIC
			<u> </u>
CES1	rs71647871	chr16:55823658	C T
CNR1	rs806380	chr6:88154934	AA
COMT	rs4680	chr22:19963748	A G
CYP1A2	rs762551	chr15:74749576	CIA
	rs2069514	chr15:74745879	G G
CYP2A6	rs1801272	chr19:40848628	AIA
	rs5031017	chr19:40843845	GIG
	rs28399433	chr19:40850474	AA
CYP2B6	rs2279343	chr19:41009358	G G
	rs3745274	chr19:41006936	GIG
	rs28399499	chr19:41012316	T T
CYP2C9	rs1057910	chr10:94981296	CIA
	rs1799853	chr10:94942290	C C
	rs7900194	chr10:94942309	GļG
	rs9332131	chr10:94949282-94949283	AIA
	rs9332239	chr10:94989020	CIC
	rs28371685	chr10:94981224	CIC
	rs28371686	chr10:94981301	C C
	rs72558187 rs72558190	chr10:94941958 chr10:94947782	T T C C
CYP2C19			GIA
CIPZCI9	rs4244285 rs4986893	chr10:94781859 chr10:94780653	G G
	rs6413438	chr10:94781858	CIC
	rs12248560	chr10:94761900	CIC
	rs12769205	chr10:94775367	GIA
	rs17884712	chr10:94775489	GG
	rs28399504	chr10:94762706	AIA
	rs41291556	chr10:94775416	тіт
	rs56337013	chr10:94852738	cic
	rs72552267	chr10:94775453	GĠ
	rs72558186	chr10:94781999	T T
CYP2D6	rs16947	chr22:42127941	G G
	rs1065852	chr22:42130692	G G
	rs1135840	chr22:42126611	CIC
	rs3892097	chr22:42128945	CIC
	rs5030655	chr22:42129084	AļA
	rs5030656	chr22:42128174-42128178	AA
	rs5030862	chr22:42130668	C C
	rs5030865	chr22:42129033	C C
	rs5030867	chr22:42127856	T T
	rs28371725 rs28371706	chr22:42127803 chr22:42129770	C C G G
	rs28371706 rs35742686	chr22:42129770 chr22:42128242	T T
	rs59421388	chr22:42127608	C C
	rs72549354	chr22:42128815-42128817	
	rs774671100	chr22:42130555-42130755	GIG
	rs201377835	chr22:42129910	CIC
	Gene Deletion	n/a	Not Detected
	Gene Duplication	n/a	Not Detected

Genes	Variant Deta	ils (GRCH38.p12)	Result
CYP3A4	rs4986907	chr7:99769804	C C
	rs35599367	chr7:99768693	GIG
	rs55785340	chr7:99768360	AlA
	rs67666821	chr7:99758184-99758188	DD
	rs72552799	chr7:99770165	C C
CYP3A5	rs776746	chr7:99672916	C C
	rs10264272 rs28365083	chr7:99665212 chr7:99652613	C C G G
	rs28383479	chr7:99660516	CIC
	rs41303343	chr7:99652771	DD
	rs55817950	chr7:99676198	gig
	rs56411402	chr7:99665237	T T
DRD2	rs6275	chr11:113412755	G G
DRD3	rs963468	chr3:114144040	A G
FAAH	rs324420	chr1:46405089	C C
FKBP5	rs4713916	chr6:35702206	A G
GNB3	rs5443	chr12:6845711	C T
GRIK1	rs2832407	chr21:29595188	C C
GRIK4	rs1954787	chr11:120792654	C C
HLA- A*31:01	rs1061235	chr6:29945521	A A
HLA- B*15:02	rs144012689	chr6:31355003	A A
HTR2A	rs6311	chr13:46897343	C C
	rs6313	chr13:46895805	G G
	rs2770296	chr13:46866425	C C
HTR2C	rs3813929	chrX:114584047	C C
HTR7	rs7905446	chr10:90859404	G T
INSIG2	rs17047764	chr2:118111006	G G
long non-	rs74795342	chr21:18954018	AIG
coding (Inc) RNA	rs75222709	chr21:18955109	G T
MC4R	rs489693	chr18:60215554	AIC
	rs17782313	chr18:60183864	T T
MTHFR	rs1801131	chr1:11794419	G G
	rs1801133	chr1:11796321	G G
OPRM1	rs1799971	chr6:154039662	AIA
POR	rs2868177	chr7:75960585	A G
SLC6A2	rs5569	chr16:55697923	GIG
	rs2242446 rs28386840	chr16:55656513 chr16:55652906	T T A A
CLCCA			
SLC6A4	5-HTTLPR	chr17:30190154-30240133	SIS
SLC6A5	rs2298826	chr11:20638211	G G
TH	rs2070762	chr11:2165105	AIG
TPH2	rs1487278	chr12:72007071	C T
UGT1A1	rs4148323 rs34815109	chr2:233760498 chr2:233760234-233760248	G G 6 7
UGT1A4	rs2011425	chr2:233718962	G T
UGT2B7	rs7439366	chr4:69098620	T C
UGT2B15	rs1902023	chr4:68670366	CIA

TEST METHODOLOGY AND LIMITATIONS

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

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