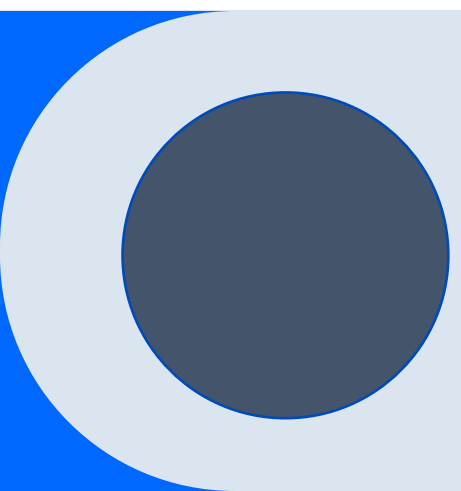
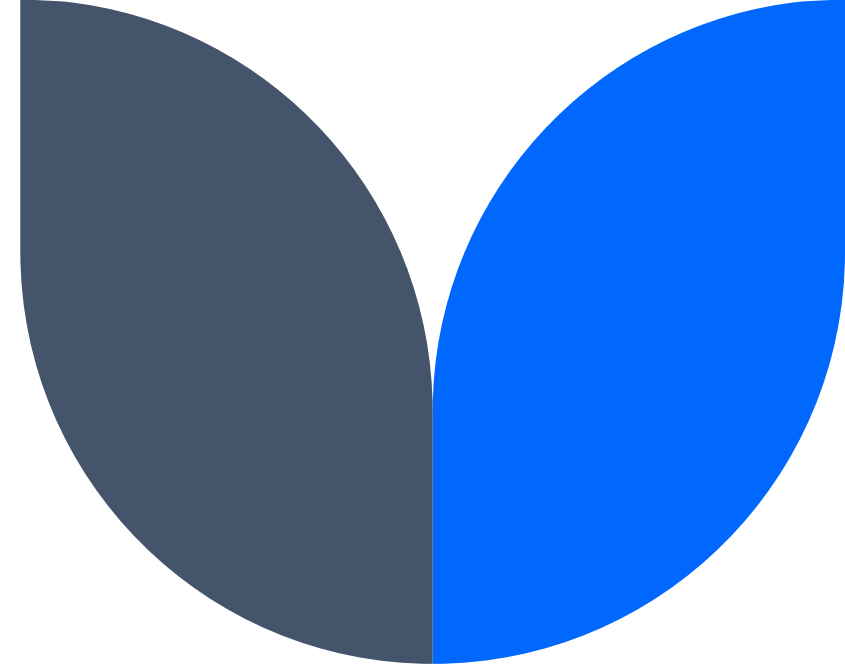




Pharmacogenomics: personalizing medications for our patients



Susan M. Smith, BS Pharm, PharmD, BCPS
Melissa Turner, PharmD

CPFI 2025 Annual Conference & National Student Retreat



Pharmacogenomics BINGO!

B I N G O

nomenclature	effective	polymorphisms	personalizing	precision
ultrarapid	variant	gene	transport	function
genetic	allele	JESUS	CYP2C9	toxicity
legal	foundational	PGx	FDA	PharmGKE
clopidogrel	normal	testing	pharmacokinetics	CPIO



This is your own unique bingo card.

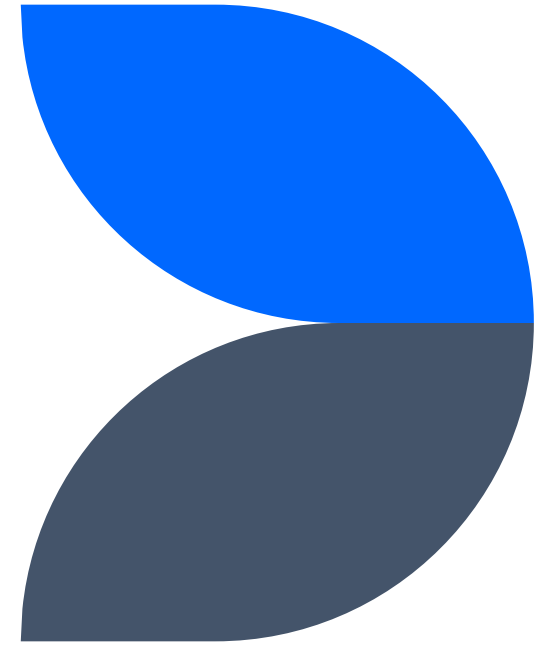
1. Click a box to mark it
2. Click a box again to unmark it
3. If you are playing this game over multiple days, bookmark this page (or email the URL to yourself), so you can get back to this card

OK

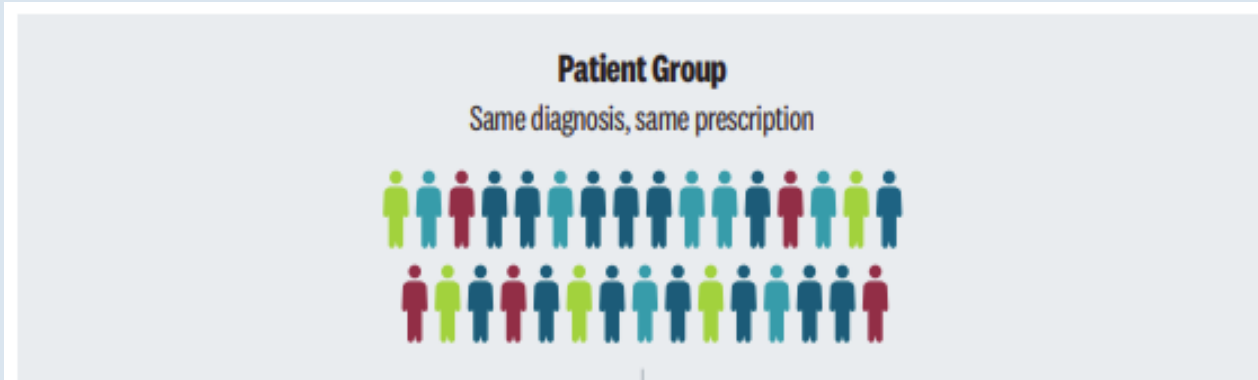
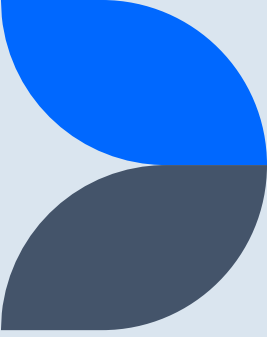
Learning Objectives

1. Explain the terms precision medicine and precision pharmacotherapy
2. Discuss ways that pharmacogenomics affects pharmacodynamics (PD) and pharmacokinetics (PK) and how it can be used to improve prescribing and outcomes
3. Categorize the Core Pharmacists Competencies in Genomics into Foundational Genetic Concepts and Clinical Pharmacogenomics Concepts
4. Given a clinical scenario, formulate an appropriate medication regimen using pharmacogenomics resources

Precision medicine and precision pharmacotherapy



One drug does not fit all



Precision medicine and pharmacotherapy

Precision Medicine

- An approach to disease treatment and prevention that takes into account individual variability in genes, environment, and lifestyle for each person

Precision Pharmacotherapy

- Customize medications to subgroups of patients, categorized by shared molecular and cellular biomarkers, to improve treatment outcomes



Medication classes impacted by pharmacogenomics

Cardiology

Psychiatry

Neurology

Oncology

Pain
management

Infectious
disease

Rheumatology

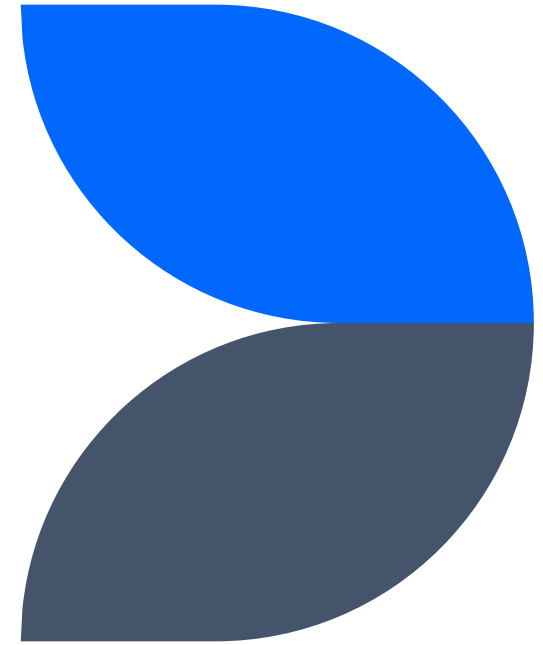
Solid organ
transplant

Respiratory

Immunology

Pharmacogenomics, Pharmacokinetics and Pharmacodynamics

PGx, PK, PD



What is pharmacogenomics (PGx)?

- Combines pharmacology and genomics to determine an individual's response to medications
- Can help healthcare providers better predict if a medication will be effective but not toxic for their patient
- PGx testing can reduce the risk of adverse effects by up to 30%

PGx, Pharmacokinetics (PK), and Pharmacodynamics (PD)

Pharmacodynamics

- Gene variants
- Adverse effects
- Receptor functions

Pharmacokinetics

- Absorption
 - Acid suppression
- Distribution
 - Bioavailability at target sites
- Metabolism
 - Metabolic rate
 - Hepatic function
 - Drug interactions
- Excretion
 - Renal function

Pharmacokinetics

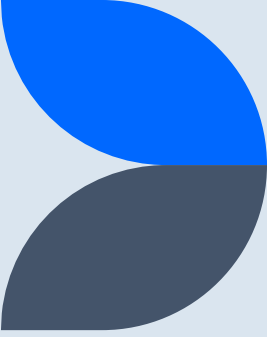
Types of metabolizer	Description	Example
Ultrarapid metabolizer (UM)	Increased enzyme activity compared to rapid metabolizers	<i>CYP2C19</i> *17/*17
Rapid metabolizer (RM)	Increased enzyme activity compared to normal metabolizers but less than ultra-rapid metabolizers	<i>CYP2C19</i> *1/*17
Normal metabolizer (NM)	Fully functional enzyme activity	<i>CYP2C19</i> *1/*1
Intermediate metabolizer (IM)	Decreased enzyme activity (activity between normal and poor metabolizer)	<i>CYP2C19</i> *1/*2
Poor metabolizer (PM)	Little to no enzyme activity	<i>CYP2C19</i> *2/*2

Pharmacodynamics

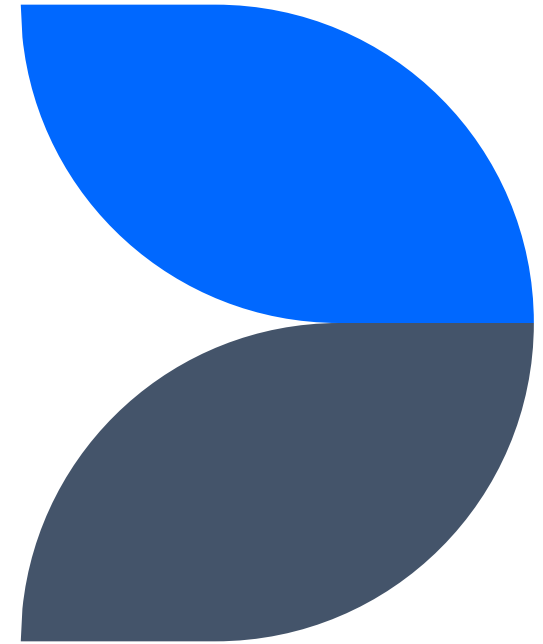
Transporter function	Description	Example
Increased function	Increased transport function compared to normal function	<i>SLCO1B1</i> *1/*14
Normal function	Fully functional transporter function	<i>SLCO1B1</i> *1/*1
Decreased function	Decreased transporter function (between normal and poor function)	<i>SLCO1B1</i> *1/*5
Poor function	Little to no transporter function	<i>SLCO1B1</i> *5/*5

My face during

Our first pharmacogenomics class



Role of the Pharmacist in Pharmacogenomics



Role of the Pharmacist in PGx

- Foundational Genetic Concepts: 6
- Clinical Pharmacogenomics: 24
- Last updated 2021

Foundational Genetic Concepts

1. Explain basic genetics concepts using appropriate nomenclature
2. Recognize the 4 factors in the manifestation of disease and drug response
3. Differentiate between the clinical diagnosis of disease informed by genetics and the identification of genetic predisposition of disease
4. Assess differences in genetic testing technologies
5. Recognize the legal protections against discrimination based on genetic test results

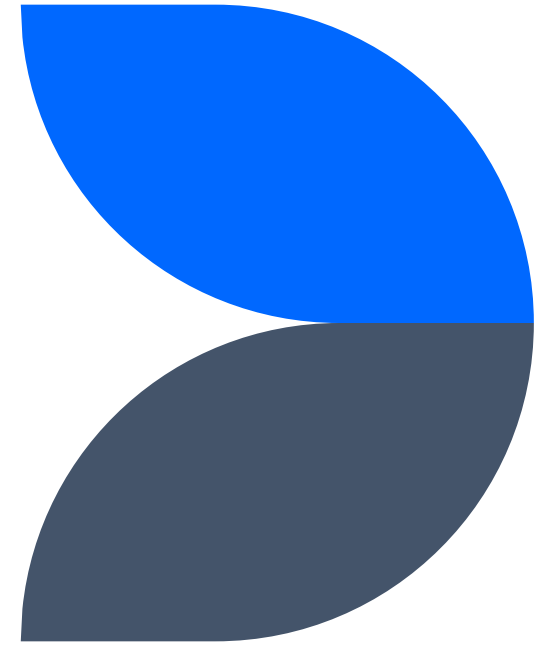
Clinical Pharmacogenomics Concepts

1. Identify PGx test results that are relevant to a patient's care
2. Interpret PGx test results
3. Determine the impact of genetic variation on pharmacokinetics and/or pharmacodynamics
4. Understand the influence of ethnicity in genetic polymorphisms and associations of polymorphisms with drug response

Beware that...

- Phenotype can change if a new medication is added =
PHENOCONVERSION

PGx Resources



Lexicomp: Pharmacogenomics section

Sertraline (Lexi-Drugs)

Outline **Alphabetical** < **Monograph** Images Adult Patient Education

[Expand All](#) ⌵

- Medication Safety Issues
- Medication Guide and/or Vaccine Information Statement (VIS)
- > Warnings & Precautions
- > Reproduction, Pregnancy, & Lactation
- > Adverse Reactions
- > Interactions
- > Pharmacogenomics**
 - Gene Testing May Be Considered**
 - Genes of Interest
- > Patient & Therapy Management
- > Preparations
- > Pharmacology & Pharmacokinetics

Gene Testing May Be Considered

- [CYP2B6 - Sertraline](#)

Genes of Interest

- [CYP2C19 - Sertraline](#)
- [HTR2A - Sertraline](#)
- [SLC6A4 - Sertraline](#)

Monitoring Parameters

Weight, height, BMI (longitudinal monitoring); closely monitor behavior (eg, anxiety, agitation, panic attacks, insomnia, irritat the initial 1 to 2 months of therapy or during periods of dosag

Nursing Practice Points

Boxed warning information: This drug has a boxed warning.

Nursing care and monitoring:

Lexicomp: Pharmacogenomics section

Abacavir (Lexi-Drugs)

Outline Alphabetical Expand All ⌵

> Uses

Clinical Practice Guidelines

> Administration and Storage Issues

Patient Counseling Points

Medication Guide and/or Vaccine Information Statement (VIS)

> Warnings & Precautions

> Reproduction, Pregnancy, & Lactation

> Adverse Reactions

> Interactions

▼ Pharmacogenomics

Gene Testing Required

Monograph Images Adult Patient Education

Gene Testing Required

- [HLA-B - Abacavir](#)

Monitoring Parameters

CBC with differential, CD4 count, HIV RNA plasma levels, serum tr genotype status prior to initiation of therapy and prior to reinitiat

Nursing Practice Points

Previous exposure/allergy to abacavir and risk factors for heart di symptoms of hypersensitivity) are available in each bottle and pat can occur within hours or at any time and may be fatal (can also c Teach patient proper timing of multiple medications and importa

Dosage Forms: US

Excipient information presented when available (limited, particula

Solution, Oral:

Ziagen: 20 mg/mL (240 mL) [contains methylparaben, propyle

Lexicomp: Pharmacogenomics section

HLA-B - Abacavir (Pharmacogenomics)

Outline [Alphabetical](#)

[Testing Recommendation](#)
Evidence Rating
Management
Related Information
Index Terms
Discussion
References

Monograph

Testing Recommendation
Testing required
See also [Definitions and Criteria for Testing Recommendations and Ratings](#).

Evidence Rating
Efficacy/Safety Outcome – Hypersensitivity Reaction: Overall Evidence Quality: Excellent (Evidence of Association: Strong; Evidence of Testing Benefit: Strong)
**Outcome with the highest overall evidence quality is displayed here. The discussion below describes evidence related to all outcomes for which there are meaningful pu*

Management
Prior to initiating abacavir, abacavir-naïve patients should be screened for *HLA-B*57:01*. Abacavir is contraindicated in *HLA-B*57:01* allele carriers due to an increased risk of experiencing a hypersensitivity reaction (HSR) (DPWG 2019; Martin 2014; Ziagen prescribing information). Discontinue abacavir treatment if an HSR is suspected. Do not reinitiate on abacavir therapy who do not have a genetic testing result for *HLA-B*57:01* carrier status should also be screened for *HLA-B*57:01*.

Lexicomp: Pharmacogenomics section

Abacavir (Lexi-Drugs)

Outline

Alphabetical

Expand All

> Brand Names

Pharmacologic Category

> Dosages

> Uses

Clinical Practice Guidelines

> Administration and Storage Issues

Patient Counseling Points

Medication Guide and/or Vaccine Information Statement (VIS)

> Warnings & Precautions

Contraindications

Monograph

Images

Adult Patient Education

Pediatric Patient Education

Contraindications

Hypersensitivity to abacavir or any component of the formulation; patients who are positive for the HLA-B*5701 allele;

Warnings/Precautions

Concerns related to adverse effects:

Hypersensitivity reactions: [US Boxed Warning]: Serious and sometimes fatal hypersensitivity reactions have at a higher risk for a hypersensitivity reaction to abacavir, although hypersensitivity reactions have occurred. All patients should be screened for the HLA-B*5701 allele prior to initiating or reinitiation of therapy. Discontinue abacavir if a hypersensitivity reaction is suspected. Abacavir should not be used in patients who are positive for the HLA-B*5701 allele or in patients with a prior hypersensitivity reaction to abacavir. Reintroduction of any abacavir product may result in severe or fatal hypersensitivity reactions, even in patients who have no history of hypersensitivity to abacavir. Hypersensitivity to abacavir should be documented in the medical record of allele-positive patients. Reactions usually occur within 6 weeks of initiation of therapy, although these reactions may occur at any time during therapy (HHS [ARV adult] 2023). These reactions usually include fever, skin rash, constitutional symptoms (malaise, fatigue, aches), respiratory symptoms (eg, pharyngitis, cough, dyspnea), and/or gastrointestinal symptoms (eg, diarrhea, nausea, vomiting). Other signs and symptoms include lethargy, headache, myalgia, edema, and laboratory abnormalities (eg, eosinophilia, leukopenia, neutropenia, hemoglobinuria, renal impairment, and liver enzyme abnormalities). Patients who develop a hypersensitivity reaction should be managed according to the manufacturer's recommendations. Patients who develop a severe or fatal hypersensitivity reaction should be managed according to the manufacturer's recommendations.

https://online-lexi-com.eu1.proxy.openathens.net/lco/action/doc/retrieve/docid/patch_f/6256?cesid=70YesdPX1Ax&searchUrl=%2Ffco%2Faction%2Fsearch%3Fq%3Dabacavir%26t%3Dname%26acs%3Dtrue%26acq%3Dabacav#genereqlist

Accessed 3.4.25

Pharmacogenomics databases

PharmGKB

- Pharmacogenomics knowledge base
- <https://www.pharmgkb.org/>

Clinical Pharmacogenetics Implementation Consortium Guidelines (CPIC)

- Guidelines – typically summarized in “Table 2”
- <https://cpicpgx.org/>

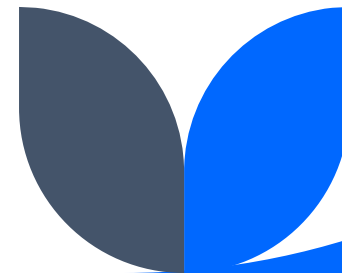


PharmGKB

Search for medications, genes, or specific genetic variants

- FDA Drug label annotations
- Drug label annotations
- Curated pathways
- Clinical guideline annotations

"Comprehensive resource that curates knowledge about the impact of genetic variation on drug response for clinicians and researchers."



CPIC

CPIC creates and posts peer-reviewed, evidence-based, updatable, and detailed gene/drug clinical practice guidelines

"CPIC is an international consortium of individual volunteers and a small dedicated staff who are interested in facilitating use of pharmacogenetic tests for patient care."

CPIC

[Guidelines](#)

[Genes-Drugs](#)

[Alleles](#)

[Publications](#)

[Meetings](#)

[Resources](#)

[Working Groups](#)

[Members](#)

[Contact](#)

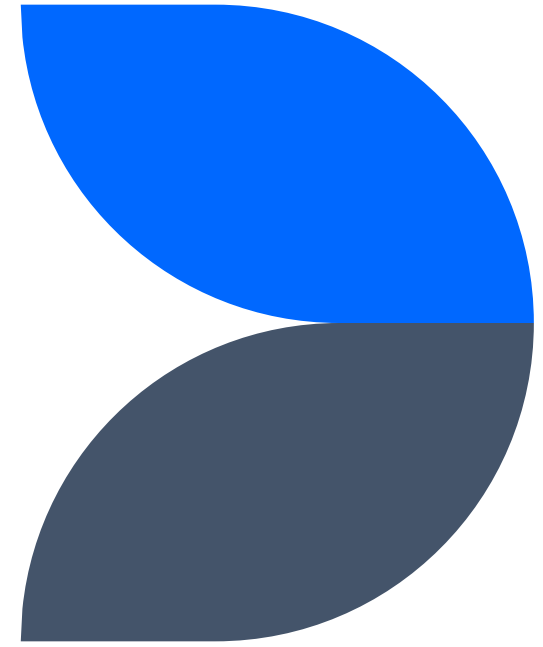


Search CPIC Website



Active Learning

Case #1



Active Learning - Case #1

LP is a 59-year-old woman with ST segment elevation myocardial infarction (STEMI) who is undergoing percutaneous coronary intervention (PCI). Pharmacogenetic testing reveals that her genotype is *CYP2C19* *17/*17. Using the information in the next 2 slides, which of the following is the MOST appropriate recommendation regarding clopidogrel use in this patient (assuming no contraindications)?

- a. do not administer clopidogrel
- b. administer clopidogrel at standard doses
- c. increase the clopidogrel dose by 50%
- d. decrease the clopidogrel dose by 50%



Active Learning - Case #1

CPIC – Guidelines

Example: clopidogrel and *CYP2C19*

CPIC UPDATE

Clinical Pharmacogenetics Implementation Consortium Guideline for *CYP2C19* Genotype and Clopidogrel Therapy: 2022 Update

Craig R. Lee¹, Jasmine A. Luzum², Katrin Sangkuhl³, Roseann S. Gammal^{4,5}, Marc S. Sabatine⁶, Charles Michael Stein⁷, David F. Kisor⁸, Nita A. Limdi⁹, Yee Ming Lee¹⁰, Stuart A. Scott^{11,12}, Jean-Sébastien Hulot¹³, Dan M. Roden¹⁴, Andrea Gaedigk¹⁵, Kelly E. Caudle⁵, Teri E. Klein³, Julie A. Johnson¹⁶ and Alan R. Shuldiner^{17,*}

CYP2C19 catalyzes the bioactivation of the antiplatelet prodrug clopidogrel, and *CYP2C19* genotype impacts clopidogrel active metabolite formation. *CYP2C19* intermediate and poor metabolizers who receive clopidogrel experience reduced platelet inhibition and increased risk for major adverse cardiovascular and cerebrovascular events. This guideline is an update to the 2013 Clinical Pharmacogenetics Implementation Consortium (CPIC) guideline for the use of clopidogrel based on *CYP2C19* genotype and includes expanded indications for *CYP2C19* genotype-guided antiplatelet therapy, increased strength of recommendation for *CYP2C19* intermediate metabolizers, updated *CYP2C19* genotype to phenotype translation, and evidence from an expanded literature review (updates at www.cpicpgx.org).

This document is an update to the 2013 Clinical Pharmacogenetics Implementation Consortium (CPIC) guideline on the clinical use of *CYP2C19* genotype test results for patients requiring antiplatelet therapy.¹ Since the previous update, results from prospective randomized clinical trials, multicenter pragmatic studies, and meta-analyses on *CYP2C19* genotype-guided antiplatelet therapy with clinical outcomes data have demonstrated the utility of this approach.^{2–6} The purpose of this guideline is to provide clinicians with information that facilitates the interpretation of clinical *CYP2C19* genotype test results to guide clopidogrel prescribing. As in the previous guideline, recommendations for use of other laboratory tests, such as platelet function monitoring, as well as cost-effectiveness analyses, are beyond the scope of this document. The guideline does not focus on demographic and other clinical variables, such as adherence to therapy, age, diabetes mellitus, obesity, smoking, and concomitant use of other drugs that may influence clopidogrel efficacy and clinical decision making. CPIC guidelines are periodically updated at www.cpicpgx.org/guidelines/.

FOCUSED LITERATURE REVIEW

A systematic literature review focused on *CYP2C19* genotype and clopidogrel response was conducted (see **Supplementary Material** for more details). Evidence is summarized in **Tables S1–S4**.

GENE: *CYP2C19*

Background

The *CYP2C19* gene is highly polymorphic; the Pharmacogene Variation Consortium (PharmVar) has currently defined over 35 star (*) allele haplotypes,⁷ including rare gene deletions (<https://www.pharmvar.org/gene/CYP2C19>; see *CYP2C19* Allele Definition Table online^{8,9}). The frequencies of these star (*) alleles significantly differ across ancestrally diverse populations (see *CYP2C19* Allele Frequency Table online^{8,9}). Alleles are categorized into functional groups as follows: normal function (e.g., *CYP2C19*1*), decreased function (e.g., *CYP2C19*9*), no function (e.g., *CYP2C19*2* and *3), and increased function (e.g., *CYP2C19*17*). Clinical allele function, as described in the

Table 1. Assignment of predicted *CYP2C19* phenotype based on genotype

Predicted phenotype	Genotype	Examples of <i>CYP2C19</i> diplotypes ^a
<i>CYP2C19</i> ultrarapid metabolizer	An individual carrying two increased function alleles	*17/*17
<i>CYP2C19</i> rapid metabolizer	An individual carrying one normal function allele and one increased function allele	*1/*17
<i>CYP2C19</i> normal metabolizer	An individual carrying two normal function alleles	*1/*1
<i>CYP2C19</i> likely intermediate metabolizer ^b	An individual carrying one normal function allele and one decreased function allele or one increased function allele and one decreased function allele or two decreased function alleles	*1/*9, *9/*17, *9/*9
<i>CYP2C19</i> intermediate metabolizer	An individual carrying one normal function allele and one no function allele or one increased function allele and one no function allele	*1/*2, *1/*3, *2/*17, *3/*17
<i>CYP2C19</i> likely poor metabolizer ^b	An individual carrying one decreased function allele and one no function allele	*2/*9, *3/*9
<i>CYP2C19</i> poor metabolizer	An individual carrying two no function alleles	*2/*2, *3/*3, *2/*3
Indeterminate metabolizer	An individual carrying one or two uncertain function alleles	*1/*12, *2/*12, *12/*14

^aPlease refer to the *CYP2C19* Diplotype-Phenotype Table online for a complete list. For allele functions and population-specific allele and phenotype frequencies, please refer to the *CYP2C19* Allele Functionality Table and the *CYP2C19* Allele Frequency Table online.^{8,9}

^bThere are limited data to characterize the function of decreased function alleles.

Active Learning – Case #1

CPIC – Guidelines Example: clopidogrel and CYP2C19

Table 2 Antiplatelet therapy recommendations based on CYP2C19 phenotype when considering clopidogrel for cardiovascular indications

CYP2C19 phenotype ^a	Implications for phenotypic measures	Therapeutic recommendation	Classification of recommendation ^b - ACS and/or PCI ^c	Classification of recommendation ^b - non-ACS, non-PCI cardiovascular indications ^d
CYP2C19 ultrarapid metabolizer	Increased clopidogrel active metabolite formation; lower on-treatment platelet reactivity; no association with higher bleeding risk	If considering clopidogrel, use at standard dose (75 mg/day)	Strong	No recommendation
CYP2C19 rapid metabolizer	Normal or increased clopidogrel active metabolite formation; normal or lower on-treatment platelet reactivity; no association with higher bleeding risk	If considering clopidogrel, use at standard dose (75 mg/day)	Strong	No recommendation
CYP2C19 normal metabolizer	Normal clopidogrel active metabolite formation; normal on-treatment platelet reactivity	If considering clopidogrel, use at standard dose (75 mg/day)	Strong	Strong
CYP2C19 likely intermediate metabolizer	Reduced clopidogrel active metabolite formation; increased on-treatment platelet reactivity; increased risk for adverse cardiac and cerebrovascular events	Avoid standard dose clopidogrel (75 mg) if possible. Use prasugrel or ticagrelor at standard dose if no contraindication	Strong ^e	No recommendation ^e
CYP2C19 intermediate metabolizer	Reduced clopidogrel active metabolite formation; increased on-treatment platelet reactivity; increased risk for adverse cardiac and cerebrovascular events	Avoid standard dose (75 mg) clopidogrel if possible. Use prasugrel or ticagrelor at standard dose if no contraindication	Strong	No recommendation
CYP2C19 likely poor metabolizer	Significantly reduced clopidogrel active metabolite formation; increased on-treatment platelet reactivity; increased risk for adverse cardiac and cerebrovascular events	Avoid clopidogrel if possible. Use prasugrel or ticagrelor at standard dose if no contraindication	Strong ^e	Moderate ^e
CYP2C19 poor metabolizer	Significantly reduced clopidogrel active metabolite formation; increased on-treatment platelet reactivity; increased risk for adverse cardiac and cerebrovascular events	Avoid clopidogrel if possible. Use prasugrel or ticagrelor at standard dose if no contraindication	Strong	Moderate

ACS, acute coronary syndrome; PCI, percutaneous coronary intervention.

^aThe online CYP2C19 Allele Frequency Table provides phenotype frequencies for major race/ethnic groups, and the online CYP2C19 Diplotype-Phenotype Table provides a complete list of possible diplotypes and phenotype assignments.^{8,9}

^bRating scheme described in the **Supplementary Material** online.

^cACS and/or PCI includes patients undergoing PCI for an ACS or non-ACS (elective) indication.

^dNon-ACS, non-PCI cardiovascular indications include peripheral arterial disease and stable coronary artery disease following a recent myocardial infarction outside the setting of PCI.

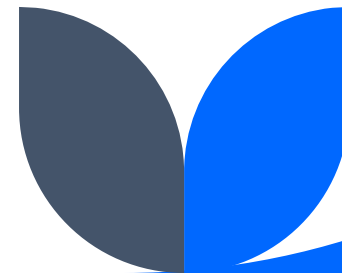
^eThe strength of recommendation for "likely" phenotypes are the same as their respective confirmed phenotypes. "Likely" indicates the uncertainty in the phenotype assignment, but it is reasonable to apply the recommendation for the confirmed phenotype to the corresponding "likely" phenotype.

<https://files.cpicpgx.org/data/guideline/publication/clopidogrel/2022/35034351.pdf> Accessed 3.4.25

Active Learning - Case #1 - ANSWER

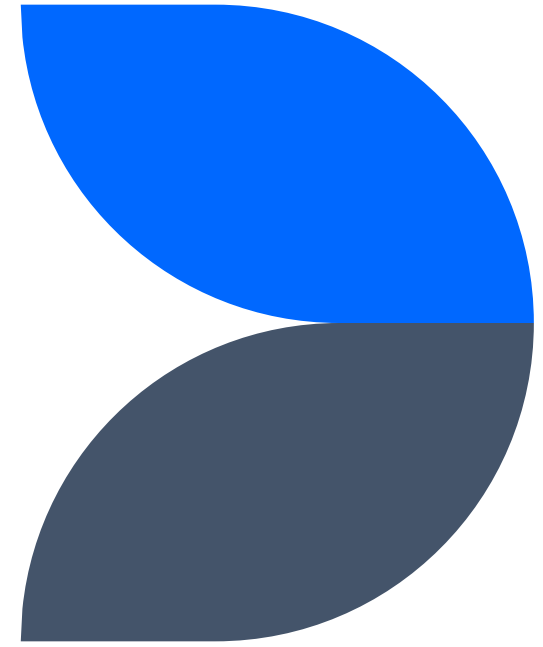
LP is a 59-year-old woman with ST segment elevation myocardial infarction (STEMI) who is undergoing percutaneous coronary intervention (PCI). Pharmacogenetic testing reveals that her genotype is *CYP2C19* *17/*17. Which of the following is the MOST appropriate recommendation regarding clopidogrel use in this patient (assuming no contraindications)?

- a. do not administer clopidogrel
- ✓ b. administer clopidogrel at standard doses
- c. increase the clopidogrel dose by 50%
- d. decrease the clopidogrel dose by 50%



Active Learning

Case #2



Active Learning - Case #2

LC is a 34-year-old African American female who was recently diagnosed with major depressive disorder. She had PGx testing done and the providers asks if paroxetine is still an appropriate option. Her genotype is *CYP2D6* *1/*1x3. Using the information in the next 2 slides, which of the following is the MOST appropriate recommendation regarding escitalopram use in this patient (assuming no contraindications)?

- a. Do not administer paroxetine; select an alternative
- b. Administer paroxetine at standard doses
- c. Consider a lower starting dose and slower titration
- d. Consider a 50% reduction in recommended starting dose, slower titration schedule, and a 50% lower maintenance dose

Active Learning - Case #2

Table 1 Assignment of predicted phenotypes based on diplotypes

Phenotype	Activity score range	Activity score/diplotypes	Examples of CYP2D6 diplotypes ^a
Assignment of predicted CYP2D6 phenotypes based on diplotypes			
CYP2D6 ultrarapid metabolizer	>2.25	>2.25	*1/*1xN, *1/*2xN, *2/*2xN
CYP2D6 normal metabolizer	$1.25 \leq x \leq 2.25$	1.25	*1/*10, *1/*9, *1/*41
		1.5	*1/*17, *1/*29
		1.75	*1/*10x3
		2.0	*1/*1, *1/*2
		2.25	*2x2/*10
CYP2D6 intermediate metabolizer	$0 < x < 1.25$	0.25	*4/*10, *4/*41
		0.5	*10/*10, *10/*41
		0.75	*10/*29, *9/*14, *17/*41
		1	*1/*5, *1/*4, *1/*5
CYP2D6 poor metabolizer	0	0	*3/*4, *4/*4, *5/*5, *5/*6

Active Learning - Case #2

Table 2 Dosing recommendations antidepressants based on CYP2D6 phenotype

Phenotype	Implication	Therapeutic recommendation	Classification of recommendation ^a	Considerations
(a) Dosing recommendations for paroxetine based on CYP2D6 phenotype				
CYP2D6 ultrarapid metabolizer	Increased metabolism of paroxetine to less active compounds when compared with CYP2D6 normal metabolizers. Lower plasma concentrations decrease the probability of clinical benefit. The extent to which ultrarapid metabolizers phenoconvert to normal, intermediate, or poor metabolizers due to paroxetine autoinhibition of CYP2D6 is unclear	Select alternative drug not predominantly metabolized by CYP2D6	Moderate	Drug–drug interactions and other patient characteristics (e.g., age, renal function, liver function) should be considered when selecting an alternative therapy
CYP2D6 normal metabolizer	Normal metabolism of paroxetine to less active compounds. Paroxetine-associated phenoconversion of normal metabolizers to intermediate or poor metabolizers due to CYP2D6 autoinhibition may occur and is dose-dependent and greater at steady state concentrations	Initiate therapy with recommended starting dose	Strong	
CYP2D6 intermediate metabolizer	Reduced metabolism of paroxetine to less active compounds when compared with CYP2D6 normal metabolizers when starting treatment or at lower doses. Higher plasma concentrations may increase the probability of side effects. Paroxetine-associated phenoconversion of intermediate metabolizers to poor metabolizers due to CYP2D6 autoinhibition may occur and is dose-dependent and greater at steady-state concentrations	Consider a lower starting dose and slower titration schedule as compared with normal metabolizers	Optional	Drug–drug interactions and other patient characteristics (e.g., age, renal function, liver function) should be considered when adjusting dose or selecting an alternative therapy
CYP2D6 poor metabolizer	Greatly reduced metabolism when compared with CYP2D6 normal metabolizers. Higher plasma concentrations may increase the probability of side effects. The impact of paroxetine-associated autoinhibition of CYP2D6 is minimal in poor metabolizers	Consider a 50% reduction in recommended starting dose, slower titration schedule, and a 50% lower maintenance dose as compared with normal metabolizers	Moderate	Drug–drug interactions and other patient characteristics (e.g., age, renal function, liver function) should be considered when adjusting dose or selecting an alternative therapy

Active Learning - Case #2 - ANSWER

LC is a 34-year-old African American female who was recently diagnosed with major depressive disorder. She had PGx testing done and the providers asks if paroxetine is still an appropriate option. Her result for *CYP2D6* *1/*1x3.

Which of the following is the MOST appropriate recommendation regarding escitalopram use in this patient (assuming no contraindications)?



- a. Do not administer paroxetine; select an alternative
- b. Administer paroxetine at standard doses
- c. Consider a lower starting dose and slower titration
- d. Consider a 50% reduction in recommended starting dose, slower titration schedule, and a 50% lower maintenance dose

Education for pharmacists

- Continuing Education (CE) impact – 1 hour CE, 8-hour certification course
- RxGenomix Clinical Pearls videos
- Books on Amazon
- Join CPIC (free) – monthly meeting on the first Thursday of every month at 11:00 am EST
 - Update on new guidelines or guidelines in progress
 - Presentation – someone who is implementing PGx in clinical practice/ambulatory care/hospital and the results

Summary

- Utilize PGx data to avoid the “trial and error” approach to pharmacotherapy management.
- Include PGx along with PK and PD considerations in your patient monitoring practices
- Recall Core Pharmacists competencies as foundational genetic concepts and clinical pharmacogenomics concepts
- Refer to PGx-specific resources to assist in formulating an appropriate medication regimen



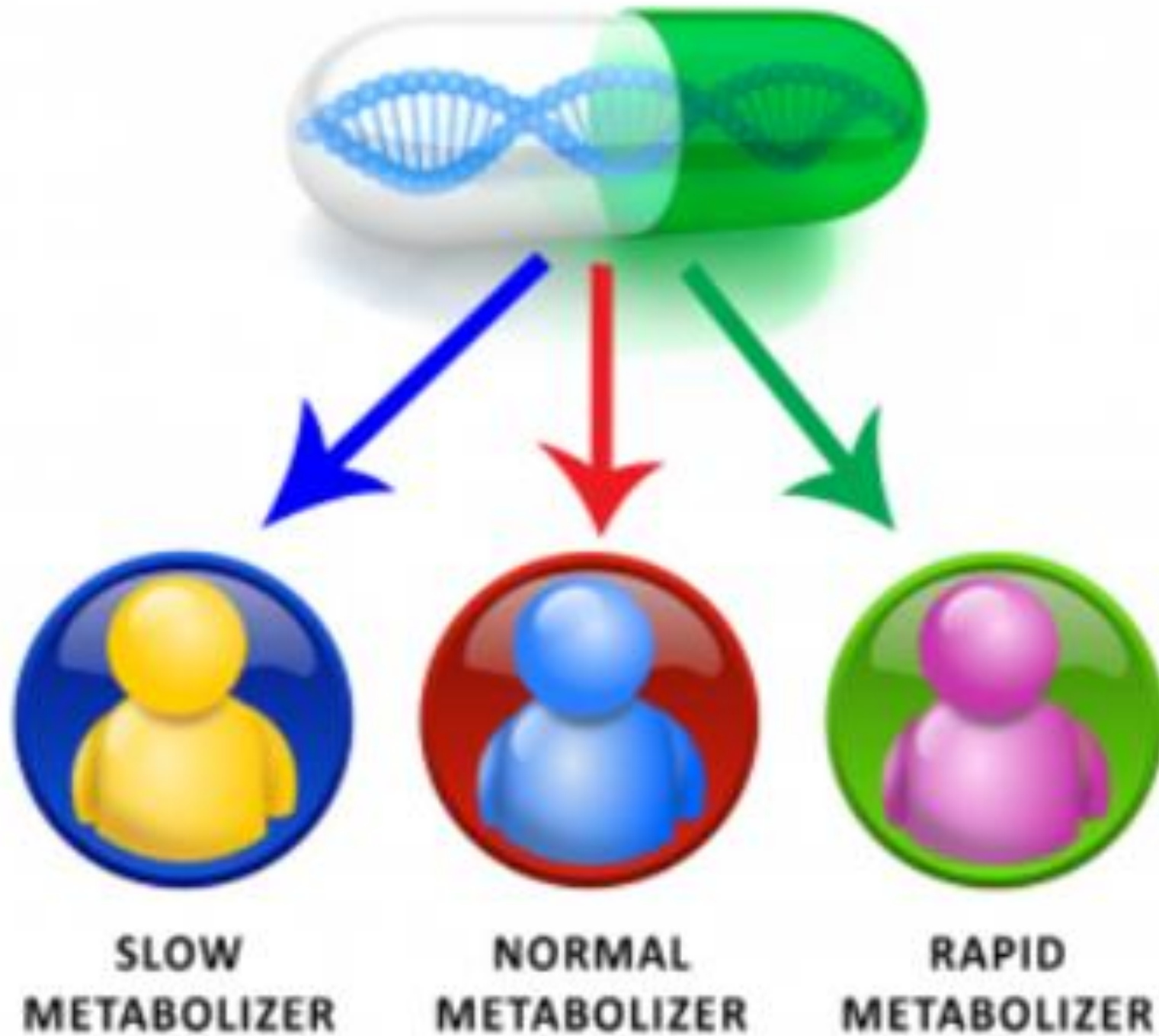
Thank you

Susan M. Smith

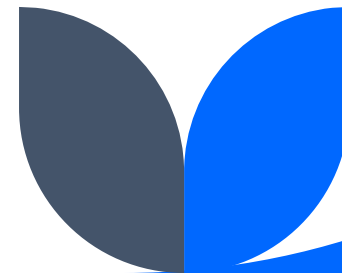
su.smith@wingate.edu

Melissa Turner

melissa@tarheelpgxconsulting.com



Questions?



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Company	Testing Options & Costs	Genes Tested	Purchase Method
ClarityX https://clarityxdna.com/products	Mindwell (\$299.25): Evaluates response to depression medications, analyzing 135+ medications MaxRX (\$374.25): Assesses response to 275+ medications across 21 therapeutic areas	Mainly <i>CYP2D6</i> , <i>CYP2C19</i> , and <i>MTHFR</i> for the therapeutic tests Other genes available: <i>ABCB1</i> , <i>GLP1R</i> , <i>CYP1A2</i> , <i>CYP3A5</i> , <i>CYP3A4</i> , <i>ANKK1</i> , <i>OPRM1</i> , <i>DRD2</i> , <i>COMT</i> , <i>VKORC1</i> , <i>APOE</i> , etc.	Patients can purchase directly from their website. Providers can order test kits to be sent to their office to test patients
OneOme https://oneome.com/rightmed-test/	RightMed® Test: Analyzes a patient's DNA to provide insights on over 100 medications. Self-pay cost is \$349, with financial assistance reducing the cost to \$199 for eligible patients.	100 alleles that covers 27 genes (<i>CYP1A2</i> , <i>CYP2B6</i> , <i>CYP2C cluster</i> , <i>CYP2C9</i> , <i>CYP2C19</i> , <i>CYP2D6</i> , <i>CYP3A4</i> , <i>CYP3A5</i> , <i>CYP4F2</i> , <i>COMT</i> , <i>DPYD</i> , <i>DRD2</i> , <i>F2</i> , <i>F5</i> , <i>GRIK4</i> , <i>HLA-A</i> , <i>HLA-B</i> , <i>HTR2A</i> , <i>HTR2C</i> , <i>IL28B</i> , <i>MTHFR</i> , <i>NUDT15</i> , <i>OPRM1</i> , <i>SLC6A4</i> , <i>SLCO1B1</i> , <i>TPMT</i> , <i>UGT1A1</i> , <i>VKROC1</i>)	Ordered through healthcare providers and pharmacists. Patients can also request the test directly from OneOme.
23andMe https://shorturl.at/SxYFg	Health + Ancestry Service (\$199): Includes ancestry and trait reports, health predisposition reports and wellness reports	Tests some alleles through saliva for <i>CYP2C19</i> , <i>SLCO1B1</i> , and <i>DPYD</i> *Health predisposition reports include those approved and not approved by the FDA *Testing on the pharmacogenetics report should be confirmed by an independent test prescribed by a healthcare provider	Direct-to-consumer; kits can be purchased online
GENETWORx https://genetworx.com/services/effectiverx-comprehensive-tests/	Comprehensive Panel: includes all genes listed Neuro Panel also available for patients only looking to test genes for mental health medications Medicaid & Medicare Part B- should be \$0. Contact for estimated cost for uninsured or commercial insurance	<i>MTHFR</i> , <i>CYP2C9</i> , <i>CYP2C19</i> , <i>CYP3A4/5</i> , <i>VKORC1</i> , <i>ADRA2A</i> , <i>ANKK1</i> , <i>COMT</i> , <i>CYP1A2</i> , <i>CYP2B6</i> , <i>CYP2D6</i> , <i>OPRM1</i> , <i>SLC6A4</i> , <i>HTR2C/2A</i> , <i>GRIK4</i> , <i>APOE</i> , <i>F2/5</i> , <i>DPYD</i> , <i>IFNL3</i> , <i>ITGB3</i> , <i>OPRK1</i> , <i>UGT1A1</i> , <i>UGT2B15</i>	Ordered through provider for patient
GeneSight https://shorturl.at/su4hm	Psychotropic Test for patients only looking to test genes for mental health medications MTHFR Test- only tests for <i>MTHFR</i> gene Cost estimated \$330 or less. Medicaid and Medicare Part B \$0	<i>MTHFR</i> , <i>CYP2D6</i> , <i>CYP2C19</i> , <i>CYP3A4</i> , <i>CYP2B6</i> , <i>CES1A1</i> , <i>CYP1A2</i> , <i>UGT1A4</i> , <i>UGT2B15</i> , <i>CYP2C9</i> , <i>SLC6A4</i> , <i>HLA-A</i> , <i>HLA-B</i> , <i>ADRA2A</i> , <i>HTR2A</i> , <i>COMT</i>	Ordered by healthcare providers. Test can be sent to the patient's home or to the healthcare provider's office

Please note that all pricing, available tests, genes, alleles, and ordering options are subject to change. We strive to provide the most accurate and up-to-date information, but changes may occur based on laboratory updates or provider availability. Final details should be confirmed prior to ordering.

QR Code to
access testing
options

