

# Pharmacogenomics Knowledgebase



&

# Pharmacogenomics Clinical Annotation Tool



Michelle Whirl-Carrillo, PhD

Director & MPI, PharmGKB

Co-Investigator, PharmCAT & CPIC

Stanford University

# The Team That Makes These Projects Possible

## Scientific Team

Li Gong, PhD  
Katrin Sangkuhl, PhD  
Ingrid Keseler, PhD  
Binglan Li, PhD  
Caroline Thorn, PhD  
Rachel Huddart, PhD

## Developers

Ryan Whaley  
Mark Woon  
Isa Reinert

Clarissa Klein  
Tiffany Murray  
Cindy Paulazzo  
Matt Wright, PhD  
Stuart Scott, PhD  
Roxana Daneshjou,  
MD, PhD  
Teri Klein, PhD  
Michelle Whirl-Carrillo,  
PhD



PharmGKB: NIH/NHGRI/NICHD/NIDA U24 HG010615 PharmCAT: NIH/NHGRI U24 HG013077



Search for a molecule, gene, variant, or combination



# Pharmacogenomics Knowledgebase

## Want Personalized PGx Recommendations?



Try out our new [Genotype Selection Interface](#) (GSI) to access and compare pharmacogenomic prescribing information from CPIC, DPWG, and FDA based on the genotypes you enter.

## Interested in Pediatric Pharmacogenomics?



Read about pediatrics on PharmGKB through the [Pediatric Dashboard](#). Switch Pediatric Focus "on" using the Focus link at the top right-hand corner of any page to see relevant information highlighted, if available. See [Pediatric Help](#) for more information.

Clinical Guideline  
Annotations

 201

Drug Label  
Annotations

 1,062

FDA Drug Label  
Annotations

 458

Curated  
Pathways

 242



## WHAT IS PHARMACOGENOMICS?

The study of the relationship between genetic variations and how our body responds to medications.

[Pretty cool right? Tell me more...](#)



## PHARMACOGENOMICS. KNOWLEDGE. IMPLEMENTATION.

**PharmGKB** is a comprehensive resource that curates knowledge about the impact of genetic variation on drug response for clinicians and researchers.

[Learn more about PharmGKB](#)

<https://www.pharmgkb.org>

# Usage & Content

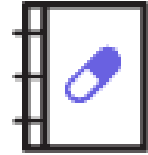


550,000-650,00  
page views/month  
50,000-55,000  
sessions/month

**Pharmacokinetic &  
pharmacodynamic pathways**

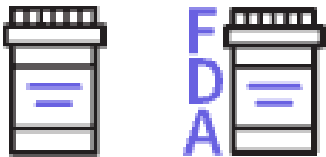
 >240

**PGx guideline  
annotations**



>190 drugs

**Drug label  
annotations**



>1000 >450 FDA

**Variant  
annotations**



>27,600



>800 >8,120 >18,000

**Clinical  
annotations**



>5,170



>3,250 >640



# New Content, Features, Tools



## Curated literature

Past year:

About 1,000 new variant annotations

About 40 new and many updated  
clinical annotations



## Staying current with updates

CPIC, DPWG, PharmVar, FDA

E.g. update all NAT2 annotations to new  
PharmVar nomenclature

Coordinate PharmVar updates across  
PharmGKB, CPIC, PharmCAT

# Annotate labels with **gene** or **variant** information

Levels of PGx information

Testing required

Testing recommended

Actionable PGx

Informative PGx

Definitions on PharmGKB: <https://www.pharmgkb.org/page/drugLabelLegend>

# Annotate labels with **gene** or **variant** information

## Levels of PGx information

Testing required

Testing recommended

Actionable PGx

Informative PGx

Contains information a clinician could consider when prescribing medication, including

- changes in efficacy, dosage, metabolism or toxicity OR
- contraindication of the drug

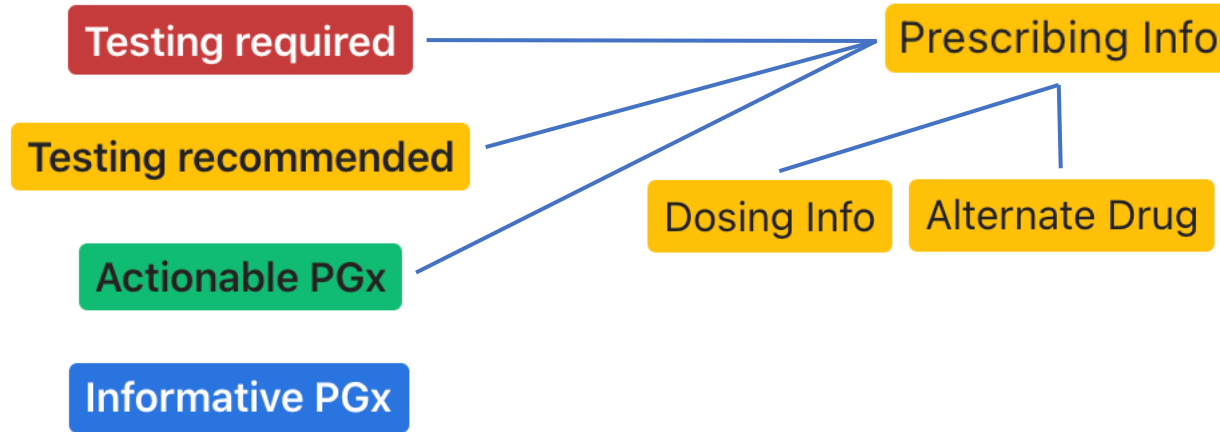
in a subset of patients with certain variants, genotypes, or phenotypes.

All other labels on the FDA Biomarker table that do not meet the other criteria.

# Drug Label Annotation Updates

Levels of PGx information

PGx guidance on the label



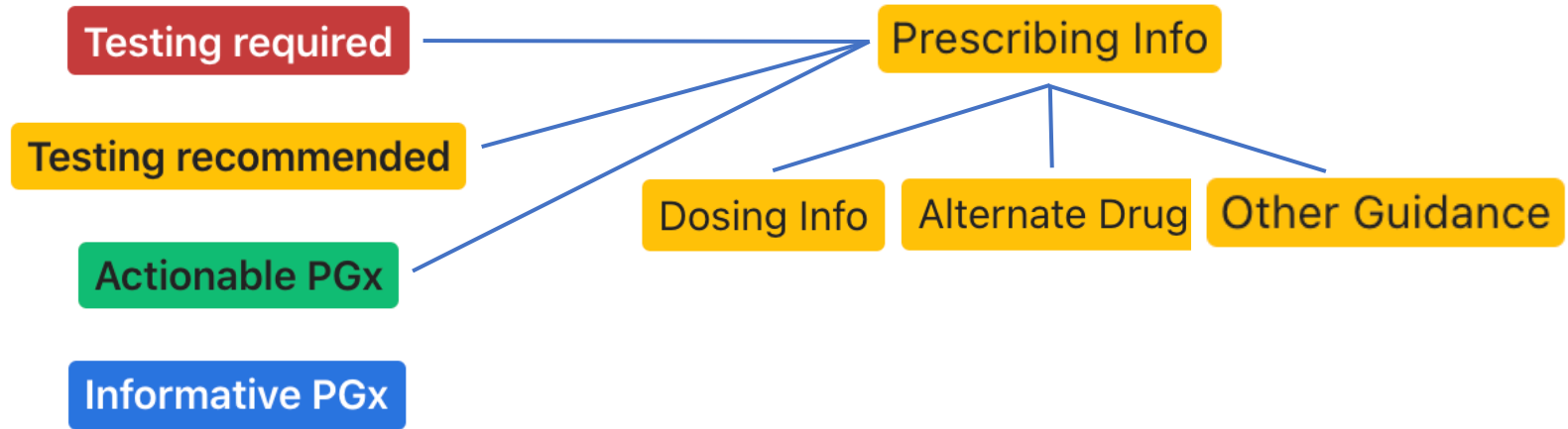
Definitions on PharmGKB: <https://www.pharmgkb.org/page/drugLabelLegend>



# Drug Label Annotation Updates

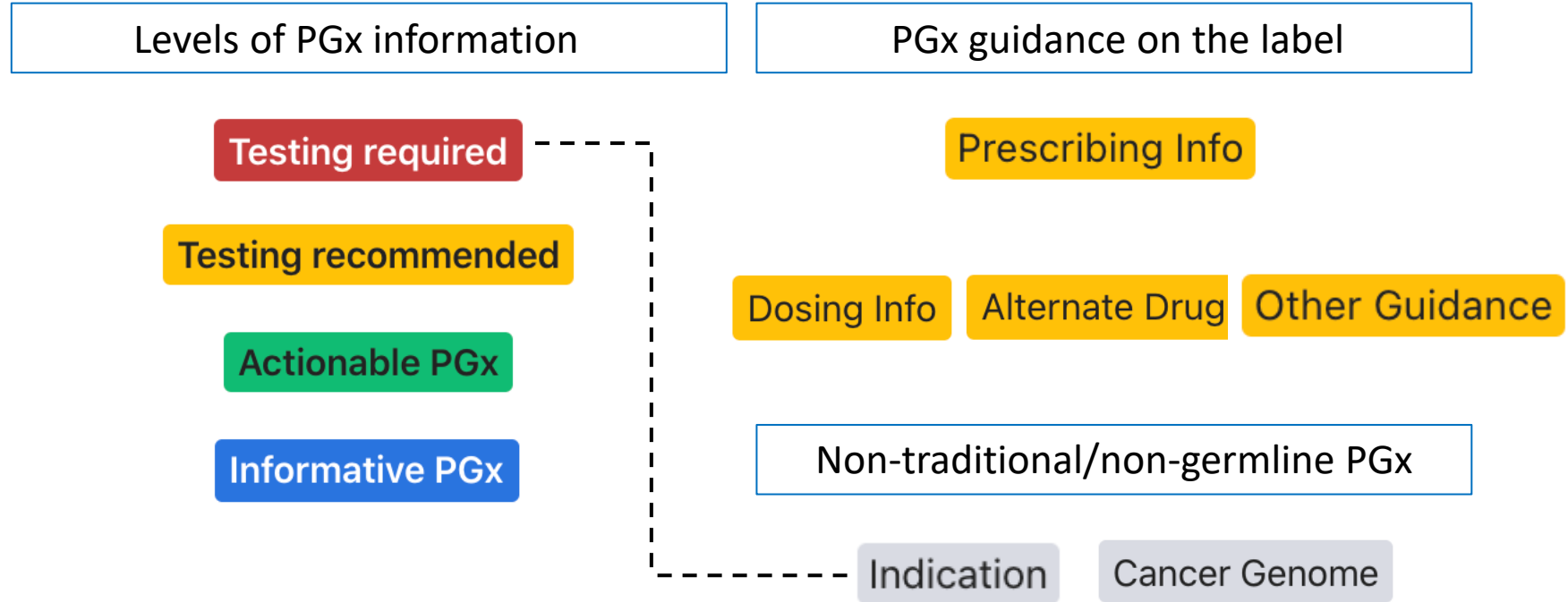
Levels of PGx information

PGx guidance on the label



Definitions on PharmGKB: <https://www.pharmgkb.org/page/drugLabelLegend>

# Annotate labels with **gene** or **variant** information



Definitions on PharmGKB: <https://www.pharmgkb.org/page/drugLabelLegend>

# Annotate labels with **gene** or **variant** information

Levels of PGx information

PGx guidance on the label

Testing required

Prescribing Info

Testing recommended

Dosing Info

Alternate Drug

Other Guidance

Actionable PGx

Informative PGx

Gene/variant not applicable or not clinically relevant

Non-traditional/non-germline PGx

Criteria Not Met

All other labels on the FDA Biomarker table that do not meet the other criteria.

Indication

Cancer Genome

Definitions on PharmGKB: <https://www.pharmgkb.org/page/drugLabelLegend>

# Table of Pharmacogenetic Associations

Share Post LinkedIn Email Print

Precision Medicine

Table of Pharmacogenetic Associations

FDA Recognition of

Pharmacogenetic tests, along with other information about patients disease or condition, can play an important role in drug therapy. When a care provider is considering prescribing a drug, knowledge of a patient's genotype may be used to aid in determining a therapeutic strategy, an appropriate dosage, or assessing the likelihood of benefit or toxicity.

## On this page:

## Section 2: Pharmacogenetic Associations for which the Data Indicate a Potential Impact on Safety or Response

Drug	Gene	Affected Subgroups+	Description of Gene-Drug Interaction
Allopurinol	HLA-B	*58:01 allele positive	Results in higher adverse reaction risk (severe skin reactions).
Carbamazepine	HLA-A	*31:01 allele	Results in higher adverse reaction risk (severe skin reactions).

## Section 1: Pharmacogenetic Associations for which the Data Support Therapeutic Management Recommendations

Drug	Gene	Affected Subgroups+	Description of Gene-Drug Interaction
Abacavir	HLA-B	*57:01 allele positive	Results in higher adverse reaction risk (hypersensitivity reactions). Do not use abacavir in patients positive for HLA-B*57:01.
Abrocitinib	CYP2C19	poor metabolizers	Results in higher systemic concentrations and may result in higher adverse reaction risk. Dosage adjustment is recommended. Refer to FDA labeling for specific dosing

## Section 3: Pharmacogenetic Associations for which the Data Demonstrate a Potential Impact on Pharmacokinetic Properties Only

The impact of these genetic variants or genetic variant inferred phenotypes on the safety or response of the corresponding drug has not been established.

Drug	Gene	Affected Subgroups+	Description of Gene-Drug Interaction
Amitriptyline	CYP2D6	ultrarapid, intermediate, or poor metabolizers	May alter systemic concentrations.

# FDA Table of Pharmacogenetic Associations

The FDA Table of Pharmacogenetic Associations was launched in 2020 and is periodically updated. PharmGKB has reprinted the information from this table below as a searchable, sortable list with links to gene and drug pages on the PharmGKB website.

[Original source at FDA.](#)

## Sections

- [Pharmacogenetic Associations for which the Data Support Therapeutic Management Recommendations](#) (n=62)
- [Pharmacogenetic Associations for which the Data Indicate a Potential Impact on Safety or Response](#) (n=22)
- [Pharmacogenetic Associations for which the Data Demonstrate a Potential Impact on Pharmacokinetic Properties Only](#) (n=40)

## Pharmacogenetic Associations for which the Data Support Therapeutic Management Recommendations

CHEMICAL ↕	GENE ↕	AFFECTED SUBGROUPS	DESCRIPTION OF GENE-DRUG INTERACTION
<a href="#">mivacurium</a>	<a href="#">BCHE</a>	intermediate or poor metabolizers	Results in higher systemic concentrations and higher adverse reaction risk (prolonged neuromuscular blockade). Avoid use in poor metabolizers.
<a href="#">succinylcholine</a>	<a href="#">BCHE</a>	intermediate or poor metabolizers	Results in higher systemic concentrations and higher adverse reaction risk (prolonged neuromuscular blockade). Avoid use in poor metabolizers. May administer test dose to assess sensitivity and administer cautiously via slow infusion.
<a href="#">pantoprazole</a>	<a href="#">CYP2C19</a>	intermediate or poor metabolizers	Results in higher systemic concentrations. Consider dosage reduction in children who are

# Prescribing Info

PharmGKB PGx Prescribing Info encompasses annotations of [clinical pharmacogenomic guidelines](#), prescribing information from [FDA approved drug labels](#) and information from the [FDA Table of Pharmacogenetic Associations](#). This list indicates which drugs have Prescribing Info available from these three sources. The list is categorized in a drug hierarchy based on the [Anatomical Therapeutic Chemical](#) classification system, with each level prefixed by the ATC identifier, then the name.

## Tag Legend

**Guideline** ⓘ = has been annotated as part of a [Clinical Guideline Annotation](#).

**Label** ⓘ = has prescribing information annotated as part of an [FDA Drug Label Annotation](#).

**PGx Assoc** ⓘ = is listed on the [FDA Table of Pharmacogenetic Associations](#).

## A ALIMENTARY TRACT AND METABOLISM DRUGS

### A02 Drugs For Acid Related Disorders

#### A02B Drugs For Peptic Ulcer And Gastro-oesophageal Reflux Disease (gord)

##### A02BC Proton pump inhibitors for peptic ulcer and GORD

- A02BC01 [omeprazole](#) **Guideline** ⓘ **PGx Assoc** ⓘ
- A02BC02 [pantoprazole](#) **Guideline** ⓘ **Label** ⓘ **PGx Assoc** ⓘ
- A02BC03 [lansoprazole](#) **Guideline** ⓘ **PGx Assoc** ⓘ
- A02BC04 [rabeprazole](#) **PGx Assoc** ⓘ
- A02BC05 [esomeprazole](#) **PGx Assoc** ⓘ
- A02BC06 [dexlansoprazole](#) **Guideline** ⓘ **PGx Assoc** ⓘ

### A03 Drugs For Functional Gastrointestinal Disorders

#### A03F Propulsives

##### A03FA Propulsives

- A03FA01 [metoclopramide](#) **Label** ⓘ **PGx Assoc** ⓘ

### A04 Antiemetics And Antinauseants



Search for a molecule, gene, variant, or combination

### Want Personalized PGx Recommendations?



Try out our new [Genotype Selection Interface](#) (GSI) to access and compare pharmacogenomic prescribing information from CPIC, DPWG, and FDA based on the genotypes you enter.

### Interested in Pediatric Pharmacogenomics?



Read about pediatrics on PharmGKB through the [Pediatric Dashboard](#). Switch Pediatric Focus "on" using the Focus link at the top right-hand corner of any page to see relevant information highlighted, if available. See [Pediatric Help](#) for more information.

Clinical Guideline  
Annotations

 201 ▾

Drug Label  
Annotations

 1,062 ▾

FDA Drug Label  
Annotations

 458 ▾

Curated  
Pathways

 242 ▾



## WHAT IS PHARMACOGENOMICS?

The study of the relationship between genetic variations and how our body responds to medications.

[Pretty cool right? Tell me more...](#)



## PHARMACOGENOMICS. KNOWLEDGE. IMPLEMENTATION.

**PharmGKB** is a comprehensive resource that curates knowledge about the impact of genetic variation on drug response for clinicians and researchers.

[Learn more about PharmGKB](#)

# GSI: Genotype Selection Interface

PharmGKB's **Genotype Selection Interface (GSI)** allows users to access and compare pharmacogenomic guideline recommendations from the [Clinical Pharmacogenetics Implementation Consortium \(CPIC\)](#) and the [Dutch Pharmacogenetics Working Group \(DPWG\)](#), and [wording from FDA's approved drug labels and Table of Pharmacogenetic Associations](#), based on individual genotypes.

- Enter genotypes or phenotypes for one or more genes below and then click the "Make Report" button to see genotype-specific drug dosing recommendations from CPIC, DPWG and FDA.
- Two alleles are required for all genes except MT-RNR1, HLA-A/HLA-B, and G6PD.

Read the [GSI documentation](#) for more information. Major changes to the content are documented [here](#).

Pick genotypes to see specific information

Gene	Diplotype		or	Phenotype	Reset
ABCG2	--	▼ --	▼ or	--	⏪ Reset
CACNA1S	--	▼ --	▼ or	--	⏪ Reset
CFTR	--	▼ --	▼ or	--	⏪ Reset
CYP2B6	--	▼ --	▼ or	--	⏪ Reset
CYP2C19	--	▼ --	▼ or	--	⏪ Reset
CYP2C9	--	▼ --	▼ or	--	⏪ Reset
CYP2D6	--	▼ --	▼ or	--	⏪ Reset
CYP3A4	--	▼ --	▼ or	--	⏪ Reset



# GSI: Genotype Selection Interface

PharmGKB's **Genotype Selection Interface (GSI)** allows users to access and compare pharmacogenomic guideline recommendations from the [Clinical Pharmacogenetics Implementation Consortium \(CPIC\)](#), and the [Dutch Pharmacogenetics Working Group \(DPWG\)](#), and wording from [FDA](#)'s approved drug labels and Table of Pharmacogenetic Associations, based on individual genotypes.

- Enter genotypes or phenotypes for one or more genes below and then click the "Make Report" button to see genotype-specific drug dosing recommendations from CPIC, DPWG and FDA.
- Two alleles are required for all genes except MT-RNR1, HLA-A/HLA-B, and G6PD.

Read the [GSI documentation](#) for more information. Major changes to the content are documented [here](#).

Pick genotypes to see specific information

Gene	Diplotype		or	Phenotype	Reset
ABCG2	--	▼	--	▼	↺ Reset
CACNA1S	--	▼	--	▼	↺ Reset
CFTR	--	▼	--	▼	↺ Reset
CYP2B6	--	▼	--	▼	↺ Reset
CYP2C19	--	▼	--	▼	↺ Reset
CYP2C9	<div style="border: 2px solid red; padding: 2px;">           ✓ --         </div>	▼	*9	▼	↺ Reset
CYP2D6	*1	▼	--	▼	↺ Reset
CYP3A4	*2	▼	--	▼	↺ Reset
	*3	▼	--	▼	↺ Reset

# GSI: Genotype Selection Interface

PharmGKB's **Genotype Selection Interface (GSI)** allows users to access and compare pharmacogenomic guideline recommendations from the [Clinical Pharmacogenetics Implementation Consortium \(CPIC\)](#), and the [Dutch Pharmacogenetics Working Group \(DPWG\)](#), and wording from [FDA](#)'s approved drug labels and Table of Pharmacogenetic Associations, based on individual genotypes.

- Enter genotypes or phenotypes for one or more genes below and then click the "Make Report" button to see genotype-specific drug dosing recommendations from CPIC, DPWG and FDA.
- Two alleles are required for all genes except MT-RNR1, HLA-A/HLA-B, and G6PD.

Read the [GSI documentation](#) for more information. Major changes to the content are documented [here](#).

Pick genotypes to see specific information

Gene	Diplotype		or	Phenotype	Reset
ABCG2	--	▼	--	▼	↻ Reset
CACNA1S	--	▼	--	▼	↻ Reset
CFTR	--	▼	--	▼	↻ Reset
CYP2B6	--	▼	--	▼	↻ Reset
CYP2C19	--	▼	--	▼	↻ Reset
CYP2C9	*2	▼	*9	▼	↻ Reset
CYP2D6	--	▼	--	▼	↻ Reset
CYP3A4	--	▼	--	▼	↻ Reset

CYP2C19	--	▼	--	▼	or	<div style="border: 1px solid gray; padding: 5px;">           ✓ --            Ultrarapid Metabolizer            Rapid Metabolizer            Normal Metabolizer  <b>Intermediate Metabolizer</b>            ...         </div>	↻ Reset
---------	----	---	----	---	----	--	---------

# GSI: Genotype Selection Interface

PharmGKB's **Genotype Selection Interface (GSI)** allows users to access and compare pharmacogenomic guideline recommendations from the [Clinical Pharmacogenetics Implementation Consortium \(CPIC\)](#) and the [Dutch Pharmacogenetics Working Group \(DPWG\)](#), and wording from [FDA](#)'s approved drug labels and Table of Pharmacogenetic Associations, based on individual genotypes.

- Enter genotypes or phenotypes for one or more genes below and then click the "Make Report" button to see genotype-specific drug dosing recommendations from CPIC, DPWG and FDA.
- Two alleles are required for all genes except MT-RNR1, HLA-A/HLA-B, and G6PD.

Read the [GSI documentation](#) for more information. Major changes to the content are documented [here](#).

Pick genotypes to see specific information

Gene	Diplotype		or	Phenotype	Reset
ABCG2	--	▼	--	▼	↺ Reset
CACNA1S	--	▼	--	▼	↺ Reset
CFTR	--	▼	--	▼	↺ Reset
CYP2B6	--	▼	--	▼	↺ Reset
CYP2C19	--	▼	--	▼	↺ Reset
CYP2C9	*2	▼	*9	▼	↺ Reset
CYP2D6	--	▼	--	▼	↺ Reset
CYP3A4	--	▼	--	▼	↺ Reset

# GSI: Results



PharmGKB's **Genotype Selection Interface (GSI)** allows users to access and compare pharmacogenomic guideline recommendations from the [Clinical Pharmacogenetics Implementation Consortium \(CPIC\)](#), and the [Dutch Pharmacogenetics Working Group \(DPWG\)](#), and wording from [FDA's](#) approved drug labels and Table of Pharmacogenetic Associations, based on individual genotypes.

- The report has sections below for the "Selected Genotypes" and the corresponding "Annotated Drugs", with columns CPIC guidelines, DPWG guidelines, FDA-approved drug labels and FDA's Table of Pharmacogenetic Associations.
- [Read more](#) about what the yellow boxes in the report mean.

Read the [GSI documentation](#) for more information.

## Selected Genotypes

Gene	Genotype	Phenotype
CYP2C19	<i>unspecified</i>	Intermediate Metabolizer
CYP2C9	*2/*9	Intermediate Metabolizer



Change



Start Over

## Annotated Drugs (21 unique) i

Show 21 actionable drugs  Show all 69 drugs with guidance related to the genes selected

### ALIMENTARY TRACT AND METABOLISM DRUGS

[dexlansoprazole](#) [dronabinol](#) [lansoprazole](#) [omeprazole](#) [pantoprazole](#)

### ANTIINFECTIVES FOR SYSTEMIC USE

[voriconazole](#)

### ANTINEOPLASTIC AND IMMUNOMODULATING AGENTS

[siponimod](#)

### BLOOD AND BLOOD FORMING ORGAN DRUGS

[avatrombopag](#) [clopidogrel](#) [warfarin](#)

### CARDIOVASCULAR SYSTEM DRUGS

[fluvastatin](#) [mavacamten](#)

### MUSCULO-SKELETAL SYSTEM DRUGS

[piroxicam](#)

### NERVOUS SYSTEM DRUGS

[rivaracetam](#) [citalopram](#) [clonazepam](#) [doxepin](#) [escitalopram](#) [fosphenytoin](#)

**CPIC**

**DPWG**

**FDA Label Annotations**

**FDA Table of Pharmacogenetic Associations**

[avatrombopag](#)

CPIC has no guidance for avatrombopag and the genes selected.

DPWG has no guidance for avatrombopag and the genes selected.

FDA Label provides no wording for avatrombopag and CYP2C9 \*2/\*9, after evaluating the evidence. [see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (PK Section) wording for avatrombopag and CYP2C9 \*2/\*9.

Other Guidance

[brivaracetam](#)

CPIC has no guidance for brivaracetam and the genes selected.

DPWG has no guidance for brivaracetam and the genes selected.

FDA Label provides no wording for brivaracetam and CYP2C19 Intermediate Metabolizer, after evaluating the evidence. [see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (Recommendations Section) wording for brivaracetam and CYP2C19 Intermediate Metabolizer.

Dosing Info

[citalopram](#)

CPIC recommended clinical action for citalopram and CYP2C19 Intermediate Metabolizer.

DPWG recommended clinical action for citalopram and CYP2C19 Intermediate Metabolizer.

FDA Label provides no wording for citalopram and CYP2C19 Intermediate Metabolizer, after evaluating the evidence. [see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (Recommendations Section) provides no wording for citalopram and CYP2C19 Intermediate Metabolizer. [see the full table](#)

Dosing Info

Dosing Info

[clobazam](#)

CPIC has no guidance for clobazam and the genes selected.

DPWG has no guidance for clobazam and the genes selected.

FDA Label provides no wording for clobazam and CYP2C19 Intermediate Metabolizer, after evaluating the evidence. [see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (Recommendations Section) wording for clobazam and CYP2C19 Intermediate Metabolizer.

Dosing Info





CPIC

DPWG

FDA Label Annotations

FDA Table of Pharmacogenetic Associations

avatrombopag

CPIC has no guidance for avatrombopag and the genes selected.

DPWG has no guidance for avatrombopag and the genes selected.

FDA Label provides no wording for avatrombopag and CYP2C9 \*2/\*9, after evaluating the evidence. [see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (PK Section) wording for avatrombopag and CYP2C9 \*2/\*9.

Other Guidance

brivaracetam

CPIC has no guidance for brivaracetam and the genes selected.

DPWG has no guidance for brivaracetam and the genes selected.

FDA Label provides no wording for brivaracetam and CYP2C19 Intermediate Metabolizer, after evaluating the evidence. [see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (Recommendations Section) wording for brivaracetam and CYP2C19 Intermediate Metabolizer.

Dosing Info

Yellow boxes indicates the recommended prescribing change at a high level

citalopram

CPIC recommended clinical action for citalopram and CYP2C19 Intermediate Metabolizer.

DPWG recommended clinical action for citalopram and CYP2C19 Intermediate Metabolizer.

FDA Label provides no wording for citalopram and CYP2C19 Intermediate Metabolizer, after evaluating the evidence. [see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (Recommendations Section) provides no wording for citalopram and CYP2C19 Intermediate Metabolizer. [see the full table](#)

Dosing Info

Dosing Info

clobazam

CPIC has no guidance for clobazam and the genes selected.

DPWG has no guidance for clobazam and the genes selected.

FDA Label provides no wording for clobazam and CYP2C19 Intermediate Metabolizer, after evaluating the evidence. [see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (Recommendations Section) wording for clobazam and CYP2C19 Intermediate Metabolizer.

Dosing Info



## [clopidogrel](#)

CPIC recommended clinical action for clopidogrel and CYP2C19 Intermediate Metabolizer.

- + Patients with cardiovascular indications of acute coronary syndrome (ACS) and/or percutaneous coronary intervention (PCI).

Alternate Drug ⓘ

- + Patients with neurovascular indications.

Alternate Drug ⓘ

- + Patients with non-acute coronary syndrome (non-ACS) and non-percutaneous coronary intervention (non-PCI) cardiovascular indications.

No Action

DPWG recommended clinical action for clopidogrel and CYP2C19 Intermediate Metabolizer.

+ Alternate Drug ⓘ

FDA Label provides no wording for clopidogrel and CYP2C19 Intermediate Metabolizer, after evaluating the evidence.

[see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (Recommendations Section) wording for clopidogrel and CYP2C19 Intermediate Metabolizer.

+ Alternate Drug ⓘ

**Click + to open, - to close more detailed annotation**

## [clopidogrel](#)

CPIC recommended clinical action for clopidogrel and CYP2C19 Intermediate Metabolizer.

 **Alternate Drug** ⓘ

### Matched Phenotype

CYP2C19: Intermediate Metabolizer

### Population

Patients with cardiovascular indications of acute coronary syndrome (ACS) and/or percutaneous coronary intervention (PCI).

### Implications

CYP2C19: Reduced clopidogrel active metabolite formation; increased on-treatment platelet reactivity; increased risk for adverse cardiac and cerebrovascular events

### Recommendation

Avoid standard dose (75 mg) clopidogrel if possible. Use prasugrel or ticagrelor at standard dose if no contraindication.

### Other Considerations

For cardiovascular indications of acute coronary syndrome (ACS) and/or percutaneous coronary intervention (PCI). ACS and/or PCI includes patients undergoing PCI for an ACS or non-ACS (elective) indication.

### Classification

Strong

[read full annotation...](#)

DPWG recommended clinical action for clopidogrel and CYP2C19 Intermediate Metabolizer.

 **Alternate Drug** ⓘ

### Matched Phenotype

CYP2C19: Intermediate Metabolizer

### Population

unspecified

### Implications

The risk of serious cardiovascular and cerebrovascular events is increased in patients undergoing balloon angioplasty or stent placement (percutaneous coronary intervention) and in patients with a stroke or TIA, as the genetic variation reduces the activation of clopidogrel. No negative clinical consequences have been observed in other patients.

### Recommendation

PERCUTANEOUS CORONARY INTERVENTION, STROKE or TIA: choose an alternative or double the dose to 150 mg/day (600 mg loading dose). Prasugrel, ticagrelor and acetylsalicylic acid/thyridamole are not metabolised by CYP2C19 (or to a lesser extent). OTHER INDICATIONS: no action required

### Classification

N/A

[read full annotation...](#)

FDA Label provides no wording for clopidogrel and CYP2C19 Intermediate Metabolizer, after evaluating the evidence.

[see the annotation for more details](#)

FDA Table of Pharmacogenetic Associations (Recommendations Section) wording for clopidogrel and CYP2C19 Intermediate Metabolizer.

 **Alternate Drug** ⓘ

### Matched Phenotype

CYP2C19: Intermediate Metabolizer

### Population

Affected subgroup: CYP2C19 intermediate or poor metabolizers

### Recommendation

"Results in lower systemic active metabolite concentrations, lower antiplatelet response, and may result in higher cardiovascular risk. Consider use of another platelet P2Y12 inhibitor."

[read full annotation...](#)

**Links to read the full guideline, label or table annotations**







This tool provides genotype-based drug prescribing guidance from [CPIC](#), [DPWG](#) and [FDA](#) curated by [PharmGKB](#).

Search by Genotype

Search by Drug Name

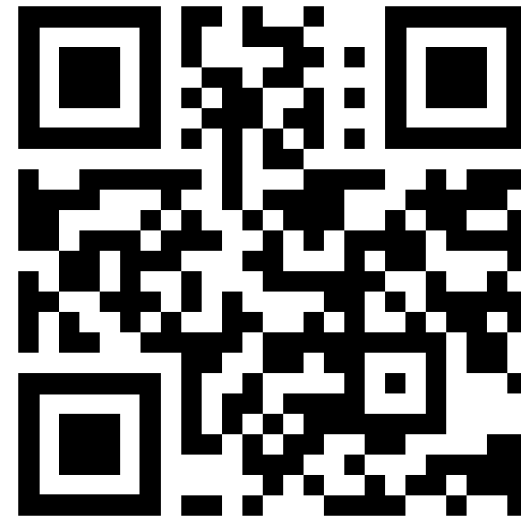


# Introducing the DDR<sub>x</sub> Mobile App!

Powered by  PHARMGKB

(Pre-release)

Funded by the Center of Excellence for Precision Health and Pharmacogenomics in the Department of Biomedical Data Science at Stanford University via Scott Penberthy (Google)



<https://ddrx.pharmgkb.org>

DDR<sub>x</sub>



more information



This tool provides genotype-based drug prescribing guidance from [CPIC](#), [DPWG](#) and [FDA](#) curated by [PharmGKB](#).

Search by Genotype

Search by Drug Name



DDR<sub>x</sub>



dark mode

### What is this?

DNA-Driven Rx (DDR<sub>x</sub>) is a tool, mostly for mobile devices, that will find drug prescribing information for particular genotypes that you specify.

You can search the prescribing guidance in two different ways.

### Search By Genotype

In this view, you will see a variety of genes and selectors for their alleles. Enter the genotyping information that you have and then all relevant drug prescribing information annotated by



## Installing DDR<sub>x</sub>

Read the [instructions for installing DDR<sub>x</sub> on your device.](#)

the genes that have been annotated in association with that drug. Enter that information and then you will be able to read genotype-specific guidance for that particular drug.

### Guidance Sources

[PharmGKB](#) has annotated prescribing guidance from the following sources:

back to home screen



Powered by PHARMGKB

## Genotype Selection

Enter a genotype to see genotype-specific drug prescribing guidance from [CPIC](#), [DPWG](#) and [FDA](#).

For more detailed information about included genes read our documentation.

## ABCG2



## CACNA1S



## CFTR



## CYP2B6

0 genes selected

[Reset](#)

View recommendations

## Genotype Selection

Enter a genotype to see genotype-specific drug prescribing guidance from [CPIC](#), [DPWG](#) and [FDA](#).

✓ -- t included

rs2231142 reference (G)

rs2231142 variant (T)



## CACNA1S



## CFTR



## CYP2B6

0 genes selected

[Reset](#)

View recommendations

## Genotype Selection

Enter a genotype to see genotype-specific drug prescribing guidance from [CPIC](#), [DPWG](#) and [FDA](#).

For more detailed information about included genes read our documentation.

## ABCG2

rs2231142 variant (T)

This gene requires 2 alleles

## CACNA1S



## CFTR



0 genes selected

[Reset](#)

View recommendations

# Genotype Selection

Enter a genotype to see genotype-specific drug prescribing guidance from [CPIC](#), [DPWG](#) and [FDA](#).

For more detailed information about included genes read our documentation.

## ABCG2

rs2231142 variant (T)

rs2231142 variant (T)

## CACNA1S

--

--

## CFTR

--

--

## CYP2B6

1 gene selected [Reset](#)

[View recommendations](#)



# Search Results

Here are drugs with actionable guidance for the submitted genotypes. For more information, including descriptions of the yellow tags, [read our documentation](#).

[Change Genotypes](#) [Start Over](#)

## ANNOTATED DRUGS (2)

### MUSCULO-SKELETAL SYSTEM DRUGS

#### Antigout Preparations

[allopurinol](#)

### CARDIOVASCULAR SYSTEM DRUGS

#### Lipid Modifying Agents

[rosuvastatin](#)

[allopurinol](#)

**DPWG**

DPWG recommended clinical action for ABCG2 rs2231142 variant (T)/rs2231142 variant (T)



# [rosuvastatin](#)



CPIC recommended clinical action for ABCG2 rs2231142 variant (T)/rs2231142 variant (T) and SLCO1B1 undefined.

## POPULATION

**General patient population.**

[Alternate Drug](#) [Dosing Info](#)

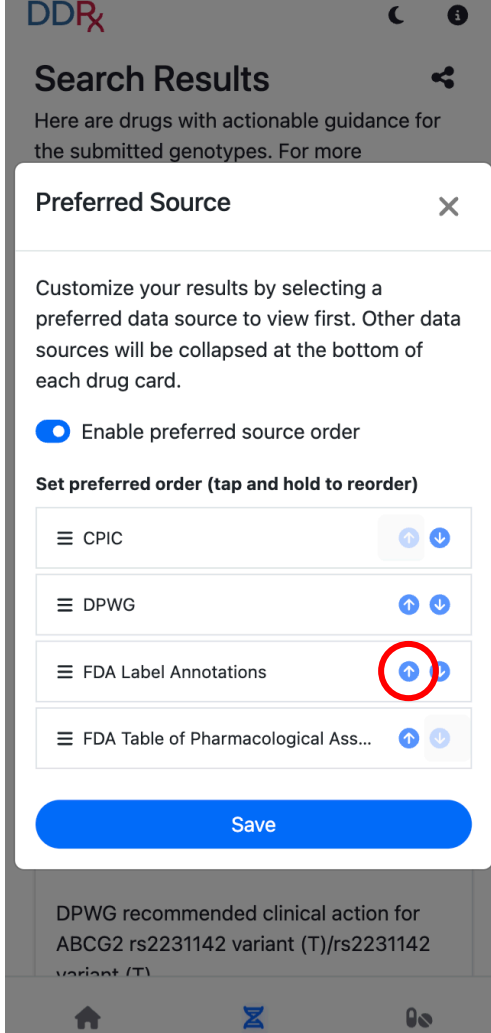
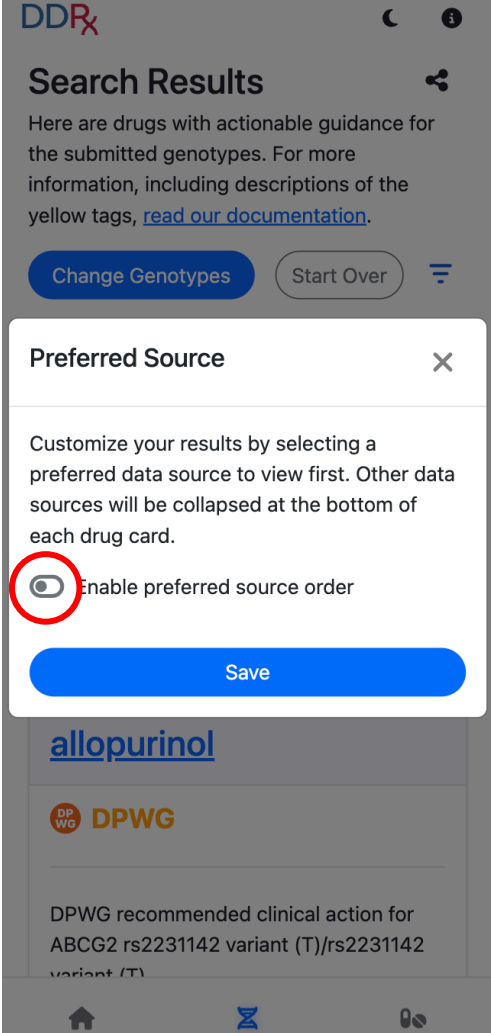
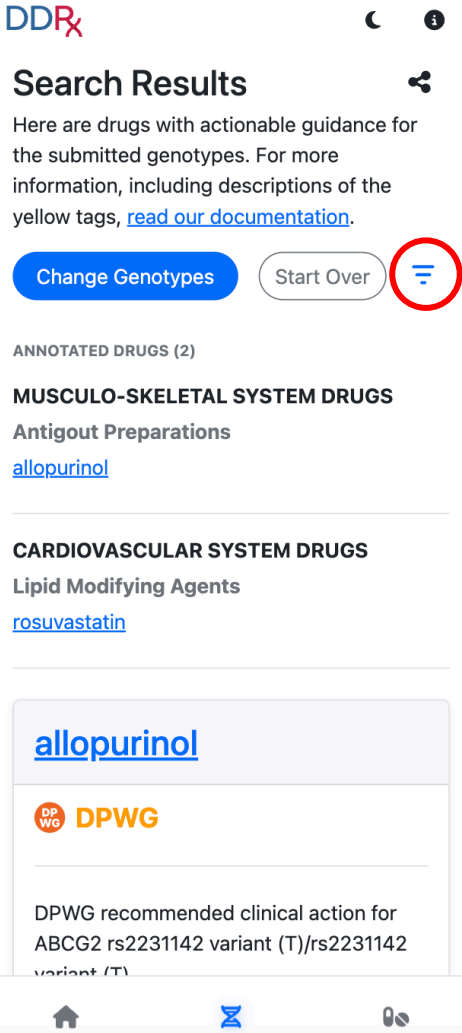
## IMPLICATIONS

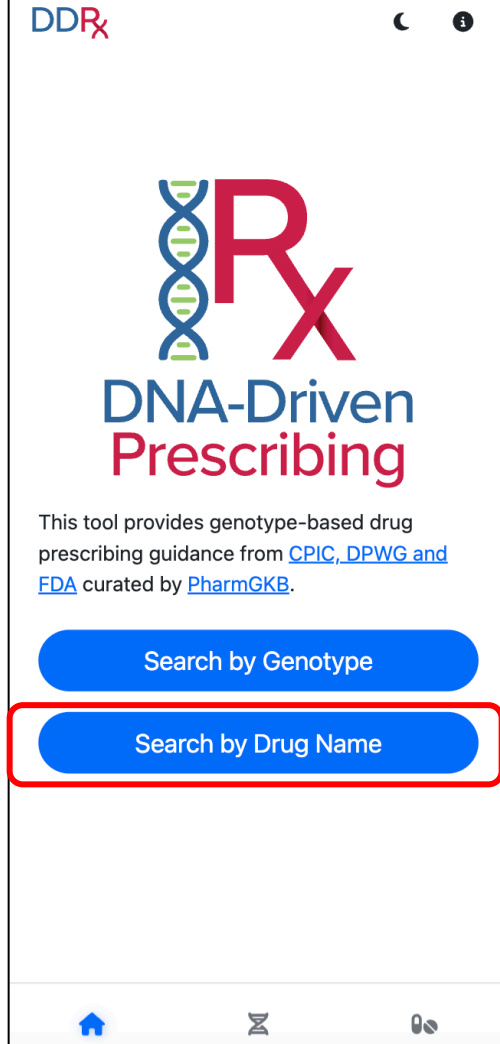
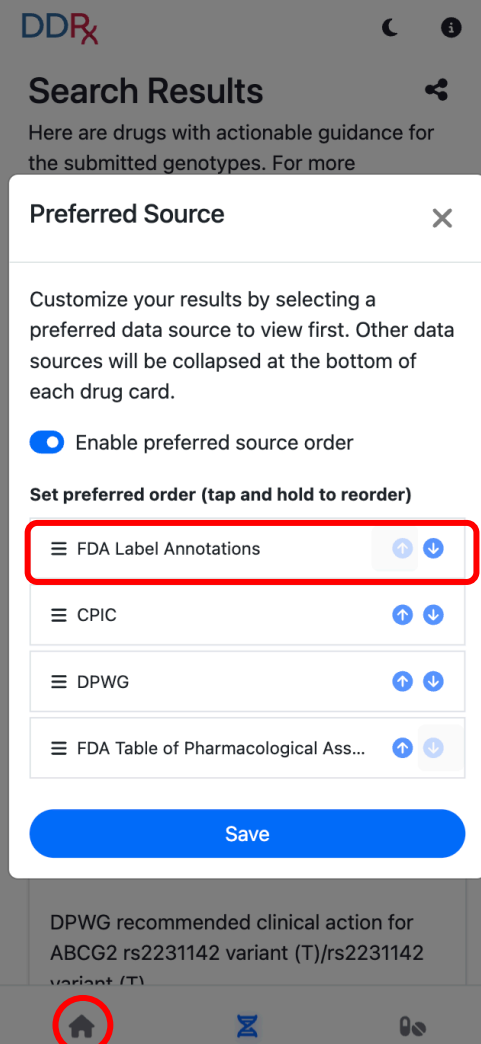
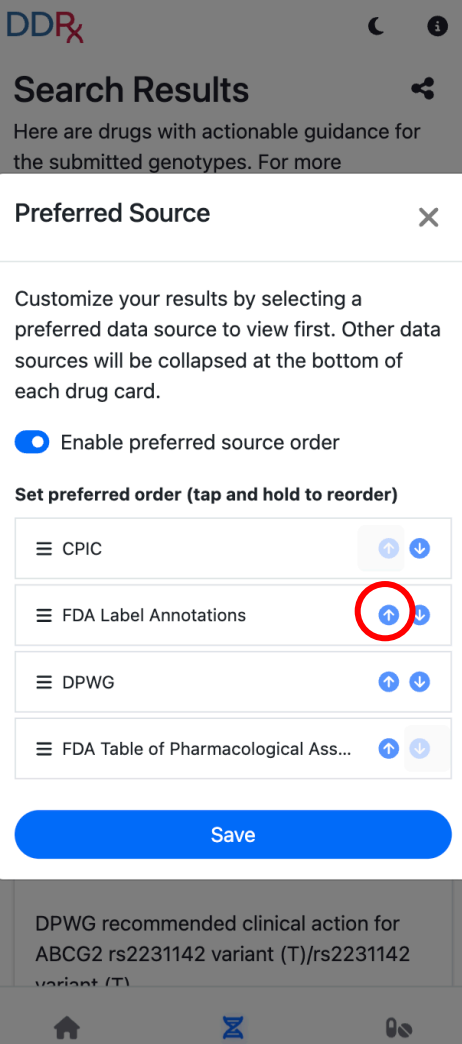
ABCG2: Increased rosuvastatin exposure compared to normal and decreased function; unknown myopathy risk; increased lipid-lowering effects  
SLCO1B1: n/a

## RECOMMENDATION

Based on ABCG2 status, prescribe  $\leq 20\text{mg}$  as a starting dose and adjust doses of rosuvastatin based on disease-specific and specific population guidelines. If dose  $> 20\text{mg}$  needed for desired efficacy, consider an alternative statin or combination therapy (i.e., rosuvastatin plus non-statin guideline directed medical therapy) (PMID: 30423391). SLCO1B1 genotype result is not available.







# Drug search

PharmGKB PGx Prescribing Info encompasses annotations of Genotype-based Drug Dosing Guidelines and Rx Annotations. Search for drugs or browse the list below.

## Search by drug name

### ALIMENTARY TRACT AND METABOLISM DRUGS

ANTIDIARRHEALS, INTESTINAL ANTIINFLAMMATORY/ANTIINFECTIVE AGENTS

- [kanamycin](#) CPIC
- [paromomycin](#) CPIC
- [streptomycin](#) CPIC FDA
- [sulfasalazine](#) FDA

### Antiemetics And Antinauseants

- [dronabinol](#) FDA
- [ondansetron](#) CPIC
- [tropisetron](#) CPIC

### Drugs For Acid Related Disorders

- [dexlansoprazole](#) CPIC FDA



# Drug search

PharmGKB PGx Prescribing Info encompasses annotations of Genotype-based Drug Dosing Guidelines and Rx Annotations. Search for drugs or browse the list below.

## Search by drug name

- [clopidogrel](#)
- [Clopidogrel \[Ban:Inn\] \(clopidogrel\)](#)
- [Clopidogrel sulfate \(clopidogrel\)](#)
- [Clopixol \(zuclopenthixol\)](#)

- [paromomycin](#) CPIC
- [streptomycin](#) CPIC FDA
- [sulfasalazine](#) FDA

### Antiemetics And Antinauseants

- [dronabinol](#) FDA
- [ondansetron](#) CPIC
- [tropisetron](#) CPIC

### Drugs For Acid Related Disorders

- [dexlansoprazole](#) CPIC FDA



[Back to search](#)

**clopidogrel** 🔗

CPIC DPWG FDA Label

FDA PGx Association Table

### Select genotypes for specific annotations

**CYP2C19**

▼

▼

0 genes selected [Reset](#)

[View recommendations](#)

### CLASS:

Antithrombotic Agents  
Blood And Blood Forming Organs  
Platelet Aggregation Inhibitors Excl. Heparin

### TRADE NAMES:

Clopidogrel [Ban:Inn]  
Clopidogrel sulfate

### Select genotypes for specific annotations

#### CYP2C19

\*2

\*2

1 gene selected

[Reset](#)

[View recommendations](#)

#### CLASS:

Antithrombotic Agents  
Blood And Blood Forming Organs  
Platelet Aggregation Inhibitors Excl. Heparin

#### TRADE NAMES:

Clopidogrel [Ban:Inn]  
Clopidogrel sulfate  
Plavix

### Select genotypes for specific annotations

#### CYP2C19

\*2

\*2

1 gene selected

[Reset](#)

[View recommendations](#)

#### FDA FDA Label Annotations

FDA Label wording for CYP2C19 \*2/\*2.

##### RECOMMENDATION

"Consider use of another platelet P2Y12 inhibitor in patients identified as CYP2C19 poor metabolizers." See label for more information.

[Read full annotation](#)

### Select genotypes for specific annotations

#### CYP2C19

\*1

\*2

1 gene selected

[Reset](#)

[View recommendations](#)

#### CPIC

CPIC recommended clinical action for CYP2C19 \*1/\*2.

##### POPULATION

**Patients with non-acute coronary syndrome (non-ACS) and non-percutaneous coronary intervention (non-PCI) cardiovascular indications.**

##### IMPLICATIONS

CYP2C19: Reduced clopidogrel active



# Variant and Star Allele Frequencies

- CPIC
- UKBB/PharmCAT
- All of Us/PharmCAT

PHARMGKB CYP2C19\*3

Overview >

- Prescribing Info
- Drug Label Annotations
- Clinical Annotations
- Variant Annotations
- Literature
- Related To
- Automated Annotations
- Links & Downloads

PRESCRIBING INFO 24

CLINICAL ANNOTATIONS 34

CPIC Gene PharmVar Gene AMP Tier 1

**PharmGKB ID**  
PA165980636

**Gene**  
[CYP2C19](#)

**HGVS Representation**  
NG\_008384.3:g.[22973G>A;85186A>G]

**CPIC Function Assignment**  
No function

**DPWG Function Assignment**  
No function

**Definition**

Any chromosomal positions listed below are assumed to be on the GRCh38 assembly. Be aware, the assembly may differ for variants elsewhere on the PharmGKB site.

[Download Allele Definition Table](#)

CHROMOSOME CHANGE	GENE CHANGE	PROTEIN CHANGE	VARIANT	REFERENCE	DBSNP
<a href="#">NC_000010.11:g.94780653 G&gt;A</a>	NG_008384.3:g.22973G>A	NP_000760.1:p.W212X	A	G	<a href="#">rs4986893</a>
<a href="#">NC_000010.11:g.94842866 A&gt;G</a>	NG_008384.3:g.85186A>G	NP_000760.1:p.I331V	G	A	<a href="#">rs3758581</a>

# Variant and Star Allele Frequencies

- CPIC
- UKBB/PharmCAT
- All of Us/PharmCAT

PHARMGBK CYP2C19\*3 Add a term to make a combination... Q Menu Focus Help

## CYP2C19\*3

### Frequencies

All of Us UK Biobank CPIC

Based on All of Us Research Program short-read whole genome sequencing data using PharmCAT v2.4.0. [More information.](#)

POPULATION	CYP2C19*3 OBSERVED	ALLELES TOTAL	FREQUENCY
AFR	40	107888	0.04%
AMR	34	81676	0.04%
EAS	683	10762	6.35%
EUR	42	246144	0.02%
MID	0	1056	0.00%
OTH	120	38578	0.31%
SAS	16	4684	0.34%

# Variant and Star Allele Frequencies

- CPIC
- UKBB/PharmCAT
- All of Us/PharmCAT

PHARMGBK CYP2C19\*3 Add a term to make a combination... Q Menu Focus Help

## CYP2C19\*3

### Frequencies

Overview > Prescribing Info • Drug Label Annotations • Clinical Annotations • Variant Annotations • Literature • Related To • Automated Annotations • Links

All of Us UK Biobank CPIC

Based on UK Biobank integrated WES + SNP array + imputed data using PharmCAT v2.2.1 in Spring 2023. [More information.](#)

POPULATION ⓘ	CYP2C19*3 OBSERVED ↕	ALLELES TOTAL ↕	FREQUENCY ↕
All populations	179	400088	0.04%
⋮ African American/Afro-Caribbean	2	3852	0.05%
⋮ Central/South Asian	14	6920	0.20%
⋮ East Asian	56	1274	4.40%
⋮ European	19	375320	0.01%
⋮ Sub-Saharan African	3	2470	0.12%
⋮ Other	85	10252	0.83%

# Variant and Star Allele Frequencies

- CPIC
- UKBB/PharmCAT
- All of Us/PharmCAT

PHARMGKB CYP2C19\*3 Add a term to make a combination... Q Menu Focus Help

## CYP2C19\*3

### Frequencies

Overview > All of Us UK Biobank **CPIC**

Prescribing Info Based on [CPIC Frequency Tables](#).

Drug Label Annotations POPULATION ⓘ CYP2C19\*3 OBSERVED ⬆ ALLELES TOTAL ⬆ FREQUENCY ⬆

Clinical Annotations	African American/Afro-Caribbean	7	2442	0.28%
Variant Annotations	American	1	1706	0.04%
Literature	Central/South Asian	126	8020	1.57%
Related To	East Asian	4675	64512	7.25%
Automated Annotations	European	109	67158	0.16%
Links	Latino	8	9698	0.08%
	Near Eastern	167	10126	1.65%
	Oceanian	2015	13768	14.64%
	Sub-Saharan African	16	6006	0.27%

# Variant and Star Allele Frequencies

- CPIC downloads by gene

## PGx Gene-specific Information Tables

- [ABCG2](#)
- [CACNA1S](#)
- [CYP2B6](#)
- [CYP2C19](#)
- [CYP2C9](#)
- [CYP2D6](#)
- [CYP3A5](#)
- [CYP4F2](#)
- [CFTR](#)
- [DPYD](#)
- [G6PD](#)
- [HLA-A/B](#)
- [IFNL3](#)
- [MT-RNR1](#)
- [NUDT15](#)
- [RYR1](#)
- [SLCO1B1](#)
- [TPMT](#)
- [UGT1A1](#)
- [VKORC1](#)

The above gene links lead to information tables created by PharmGKB and CPIC. The files support CPIC guidelines, but are also general resources for these PGx genes. The following types of files are provided by gene, as available:

- **Allele Definition Table**
  - Information about what variants define star (\*) or named alleles
  - Mapping of variants to the human genome GRCh38, the RefSeq Gene sequence and protein sequence, and provides rsIDs, if available
- **Allele Functionality Table**
  - Allele function assignments using [CPIC standardized terms](#)
  - References for the allele function
- **Phenotype Table**
  - Mapping allele function combinations to phenotypes
- **Diplotype-Phenotype Table**
  - Mapping of each diplotype to phenotype
- **Example CDS Table**
  - Mapping of possible phenotype to EHR priority result notation and consultation text
- **Workflow Diagram**
  - Possible implementation workflow diagram
- **Frequency Table**
  - Calculated allele frequency by PharmGKB biogeographical groups based on frequencies reported by references. Further details about the biogeographical grouping system can be found [here](#) or in [Article:[30506572](#)]
- **Gene Resource Mappings**
  - Mapping of gene to ID or code for HGNC, NCBI, Ensembl and PharmGKB

# Downloads

## From PharmGKB

In addition to the PharmGKB website, we are pleased to make PharmGKB data and knowledge available as downloads. We have found that it is often critical to check with our curators at [feedback@pharmgkb.org](mailto:feedback@pharmgkb.org) before embarking on a large project using these data, to be sure that the files and data we make available are being interpreted correctly. PharmGKB generally does NOT need to be a co-author on such analyses; we just want to make sure that there is a correct understanding of our data before lots of resources are spent.

[Examples of papers that have been written by others using PharmGKB](#)



PharmGKB data is licensed under a [Creative Commons Attribution-ShareAlike](#) license.

# Variant and Star Allele Frequencies

- UKBB bulk download

## Other Datasets



### PharmGKB Branding

PharmGKB has a collection of logos, emblems, and other graphics that can be used when referring to PharmGKB on other sites and in publications.

[PharmGKB branding repository](#)



### PharmGKB Training Exercises

These exercises are intended to help new users familiarize themselves with the PharmGKB website. Be aware that this download includes the answers to the exercises. As a result, these exercises are not recommended to be used for credit/CME.

[PharmGKB Training Exercises](#)



### Papers of Interest Archive

This is an archive of all papers of interest (aka Curators Favorite Papers) from May 2006 to April 2017. All future papers of interest will be integrated into the [PharmGKB Blog](#).

[Papers of Interest Archive](#)



### UK Biobank Frequencies

Haplotype frequencies from an analysis of the UK Biobank dataset using PharmCAT. Data from *Frequencies of pharmacogenomic alleles across biogeographic groups in a large-scale biobank* (PMID: [37757824](#))

[pharmgkb\\_haplotype\\_frequencies\\_UKBB.zip](#)

31.4 KB



Search PharmCAT

Home

Methods



Using PharmCAT



Genes & Drugs

Examples

Changelog

FAQs

Disclaimers & Other  
Information

About Us

# PharmCAT:

## Pharmacogenomics Clinical Annotation Tool



Download v2.13.0

View on GitHub

An active area of genomic medicine implementation at many healthcare organizations and academic medical centers includes development of decision support and return of results around pharmacogenomics. One of the challenges in implementing pharmacogenomics is the representation of the information in clinical dosing guidelines, including star-allele haplotypes, and extracting these variants and haplotypes from genetic datasets. In a collaboration between the [Pharmacogenomics Research International \(PGRN\)](#) and the former [PGRN Statistical Analysis Resource \(P-STAR\)](#), with input from [PharmGKB](#) and other groups, we are developing a software tool to extract guideline variants from a genetic

<https://pharmcat.org>



# Pharmacogenomics Clinical Annotation Tool

## New Content & Features

- CPIC and PharmVar updates
  - E.g. DPYD HapB3 and RYR1 updates
- DPWG guidance and updates
  - E.g. F5 was removed
- FDA PGx guidance
  - Table of PGx Associations
  - PharmGKB label annotations
- Multiple pre-processor updates for improved performance





# Pharmacogenomics Clinical Annotation Tool

## New Content & Features

- 175 drugs with 18 genes
- Accommodate 4 additional genes
  - HLA-A, HLA-B, CYP2D6 & MT-RNR1
- Outreach
  - Increased website documentation
  - “How-to” videos on YouTube channel
  - Published tutorial paper (2022)
  - PharmCAT mailing list for releases/news/updates
  - Address user feedback through email and GitHub issues

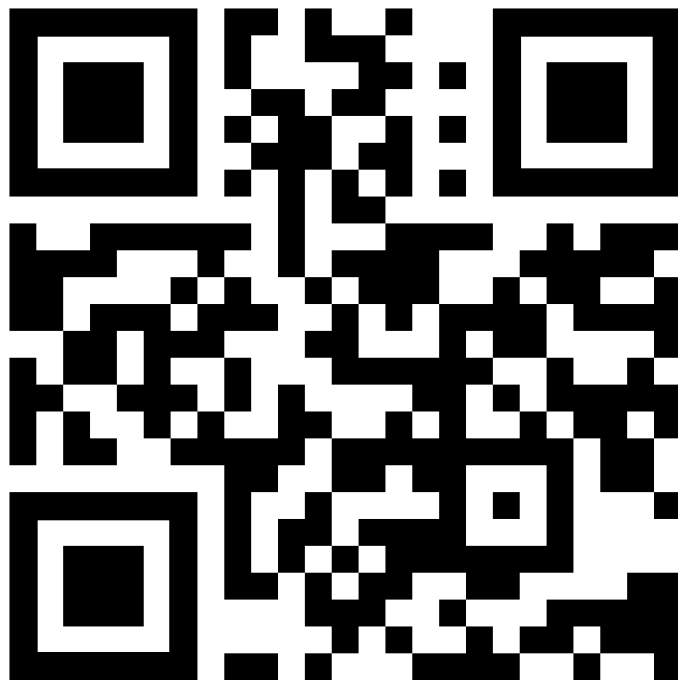


# Pharmacogenomics Clinical Annotation Tool

## Recent & Upcoming Projects

- UKBB allele frequencies
  - CYP4F2 allele frequencies
  - HapB3 observations
  - PMID: 37757824
- Stanford Health Care implementation
- Collaborations for inclusion of diverse populations
  - China Medical University Hospital, Taiwan
  - Galatea Bio
  - gnomAD/Broad
  - To be available on PharmGKB

# DDRx Mobile App



Funded by the Center of Excellence for Precision Health and Pharmacogenomics in the Department of Biomedical Data Science at Stanford University via Google/Scott Penberthy