

Pharmacogenetics Panel for Psychotropics

Last Literature Review: July 2022 Last Update: December 2025

Variation in genes affecting pharmacokinetics and/or pharmacodynamics (pharmacogenetics) may influence medication selection and dose planning.¹ For example, variants in genes that code for metabolizing enzymes (*CYP2B6*, *CYP2C19*, *CYP2C9*, *CYP2D6*, *CYP3A4*, *CYP3A5*, and *UGT2B15*) may be associated with altered (slower or faster) metabolism, which would affect the kinetics of medication activation, inactivation, and/or elimination. Other genes may predict the risk of side effects and/or the likelihood of response (*ANKK1*, *COMT*, *DRD2*, *GRIK4*, *HTR2A*, *HTR2C*, *MTHFR*, and *OPRM1*). These panels may be used to assess genetic variants that may inform selection and dosing for many prescription medications, including medications relevant to psychiatry such as psychostimulants (eg, ADHD medication), antidepressants, antipsychotics, and anxiolytics. These tests are appropriate for individuals with a personal or family history of therapeutic failure or adverse events related to psychotropic medications.

For more information on pharmacogenetic testing, refer to the ARUP Consult [Germline Pharmacogenetics - PGx](#) topic.

Featured ARUP Testing

[Pharmacogenetics Panel: Psychotropics 3004471](#)

Method: Polymerase Chain Reaction (PCR) / Fluorescence Monitoring / Sequencing

[Pharmacogenetics Panel: Psychotropics, with GeneDose Access 3006366](#)

Method: Polymerase Chain Reaction (PCR) / Fluorescence Monitoring / Sequencing

Report may include comprehensive medication guidance based on the genotypes detected and access to GeneDose Live, a cloud-based medication management and risk mitigation tool.

Refer to the ARUP [Laboratory Test Directory](#) for additional testing to assess genetic risk of abnormal drug metabolism for specific substrates, including *CYP2B6*, *CYP2C19*, *CYP2C8*, *CYP2C9*, *CYP2D6*, *CYP3A4*, *CYP3A5*, and the methylenetetrahydrofolate reductase variant.

Treatment Issues

- Actual metabolic phenotype is subject to interactions related to pharmacogenes, medications, supplements, food, lifestyle choices, clinical factors, as well as genetic and nongenetic factors not identified by this test. This test is intended only to identify targeted pharmacogenetic variants.
- Therapeutic and clinical monitoring are needed to guide pharmacotherapy for a particular patient.
- A table providing known clinical associations between the gene variants detected in this test and several common psychotropic medications can be found [here](#).
- Additional resources are available through the U.S. Food and Drug Administration (FDA),² the Clinical Pharmacogenetics Implementation Consortium (CPIC),³ and the ClinPGx.⁴

Testing Considerations

- This panel provides a comprehensive analysis for multiple genes that have strong pharmacogenomic associations with medications used in the treatment of psychiatric disorders, including depression. Specific gene-drug interactions are supported by guidelines from CPIC, Association for Molecular Pathology (AMP), or Dutch Pharmacogenetics Working Group (DPWG) (*CYP2B6*, *CYP2C19*, *CYP2C9*, *CYP2D6*, and *CYP3A4*).
- This panel also provides analysis for multiple genes that have level 3 evidence linking the genes to psychotropic medications according to ClinPGx (*MTHFR*, *CYP3A5*, *DRD2*, *ANKK1*, *GRIK4*, *HTR2A*, *HTR2C*, *UGT2B15*, *COMT*, and *OPRM1*). More details are available in the ClinPGx [Levels of Evidence for Selected Medication/Gene Associations](#) table or on the ClinPGx website.⁴
- Some of the genes tested in the psych panel are part of other panels offered by ARUP, and hence there might be instances where some of the alleles reported in the psych panel are not linked to psychotropic medications. Caution must be taken when clinically interpreting these findings relevant to the medications patients are using.
- The test result does not replace the need for therapeutic medication or clinical monitoring.

Genetics

Genes

ANKK1, *COMT*, *CYP2B6*, *CYP2C19*, *CYP2C9*, *CYP2D6*, *CYP3A4*, *CYP3A5*, *DRD2*, *GRIK4*, *HTR2A*, *HTR2C*, *MTHFR*, *OPRM1*, *UGT2B15*

Inheritance

Autosomal codominant

Variants Tested

Variant(s) from a reference sequence are detected by targeted genotyping (not full gene sequencing) and by copy number determinations (*CYP2D6* only). Conventional nomenclature for describing the variants and predicting phenotype is applied, when available. For many pharmacogenes, variants are grouped and classified as “star” (*) alleles that are associated with predicted enzyme or protein function, based on international consensus nomenclature. However, not all variants in a gene are tested, and assumptions about phase or diplotype assignments are made. More details about the nomenclature, allele frequencies, and phenotype predictions are available at the ClinPGx⁴ and PharmVar⁵ websites.

Gene (Transcript) and Brief Summary of Associated Protein Function	Alleles	Predicted Allele Function ^a
<i>ANKK1</i> (NM_178510) (DRD2 associated) This variant in ANKK1 represents the Taq1A polymorphism that affects the expression of binding sites for dopamine on the D2 receptor and can influence response to psychotropic medications that target the dopaminergic system	rs1800497, c.2137G>A	Drug dependent ^b
<i>COMT</i> (NM_000754) Codes for catechol-o-methyltransferase, an enzyme that metabolizes catecholamines such as dopamine and is involved in response to several medications	rs4680, c.472G>A	Drug dependent ^b
<i>CYP2B6</i> (NM_000767) Codes for the cytochrome P450 2B6 enzyme, which is involved in metabolism of many medications; impact of the phenotype depends on whether a medication is activated or inactivated by this enzyme	<i>CYP2B6</i> *4: rs2279343, c.785A>G	Increased function
	<i>CYP2B6</i> *6: rs3745274, c.516G>T; rs2279343, c.785A>G	Decreased function
	<i>CYP2B6</i> *7: rs3745274, c.516G>T; rs2279343, c.785A>G; rs3211371, c.1459C>T	Decreased function
	<i>CYP2B6</i> *9: rs3745274, c.516G>T	Decreased function
	<i>CYP2B6</i> *18: rs28399499, c.983T>C	No function
	<i>CYP2B6</i> *22: rs34223104, c.-82T>C	Increased function
	<i>CYP2B6</i> *36: rs34223104, c.-82T>C; rs3745274, c.516G>T; rs2279343, c.785A>G	Decreased function
<i>CYP2C19</i> (NM_000769) Codes for the cytochrome P450 2C19 enzyme, which is involved in metabolism of many medications; impact of the phenotype depends on whether a medication is activated or inactivated by this enzyme	<i>CYP2C19</i> *2: rs4244285, c.681G>A; rs12769205, c.332-23A>G	No function
	<i>CYP2C19</i> *3: rs4986893, c.636G>A	No function
	<i>CYP2C19</i> *4A: rs28399504, c.1A>G	No function
	<i>CYP2C19</i> *4B: rs28399504, c.1A>G, rs12248560, c.-806C>T;	No function
	<i>CYP2C19</i> *5: rs56337013, c.1297C>T	No function
	<i>CYP2C19</i> *6: rs72552267, c.395G>A	No function
	<i>CYP2C19</i> *7: rs72558186, c.819+2T>A	No function
	<i>CYP2C19</i> *8: rs41291556, c.358T>C	No function
	<i>CYP2C19</i> *9: rs17884712, c.431G>A	Decreased function

Gene (Transcript) and Brief Summary of Associated Protein Function	Alleles	Predicted Allele Function ^a
<p><i>CYP2C9</i> (NM_000771)</p> <p>Codes for the cytochrome P450 2C9 enzyme, which is involved in metabolism of many medications; impact of the phenotype depends on whether a medication is activated or inactivated by this enzyme</p>	<i>CYP2C9</i> *17: rs12248560, c.-806C>T	Increased function
	<i>CYP2C9</i> *35: rs12769205, c.332-23A>G	No function
	<i>CYP2C9</i> *2: rs1799853, c.430C>T	Decreased function
	<i>CYP2C9</i> *3: rs1057910, c.1075A>C	No function
	<i>CYP2C9</i> *4: rs56165452, c.1076T>C	Decreased function
	<i>CYP2C9</i> *5: rs28371686, c.1080C>G	Decreased function
	<i>CYP2C9</i> *6: rs9332131, c.818del	No function
	<i>CYP2C9</i> *8: rs7900194, c.449G>A	Decreased function
	<i>CYP2C9</i> *11: rs28371685, c.1003C>T	Decreased function
	<i>CYP2C9</i> *12: rs9332239, c.1465C>T	Decreased function
<p><i>CYP2D6</i>^c (M33388 sequence)</p> <p>Codes for the cytochrome P450 2D6 enzyme, which is involved in metabolism of many medications; impact of the phenotype depends on whether a medication is activated or inactivated by this enzyme. The test also evaluates copy number variations and can discriminate which allele is affected in most cases that exhibit gene duplication.</p>	<i>CYP2D6</i> *2: rs16947, g.2850C>T; rs1135840, g.4180G>C	Functional
	<i>CYP2D6</i> *2A: rs1080985, g.-1584C>G; rs16947, g.2850C>T; rs1135840, g.4180G>C	Functional
	<i>CYP2D6</i> *3: rs35742686, g.2549delA	No function
	<i>CYP2D6</i> *4: rs1065852, g.100C>T; rs3892097, g.1846G>A; rs1135840, g.4180G>C	No function
	<i>CYP2D6</i> *5: gene deletion	No function
	<i>CYP2D6</i> *6: rs5030655, g.1707delT	No function
	<i>CYP2D6</i> *7: rs5030867, g.2935A>C	No function
	<i>CYP2D6</i> *8: rs5030865, g.1758G>T; rs16947, g.2850C>T; rs1135840, g.4180G>C	No function
	<i>CYP2D6</i> *9: rs5030656, g.2615_2617delAAG	Decreased function
	<i>CYP2D6</i> *10: rs1065852, g.100C>T; rs1135840, g.4180G>C	Decreased function
	<i>CYP2D6</i> *11: rs1080985, g.-1584C>G; rs201377835, g.883G>C; rs16947, g.2850C>T; rs1135840, g.4180G>C	No function
	<i>CYP2D6</i> *13: a CYP2D7-derived exon 1 conversion	No function
	<i>CYP2D6</i> *14: rs5030865, g.1758G>A; rs16947, g.2850C>T; rs1135840, g.4180G>C	Decreased function
	<i>CYP2D6</i> *15: rs774671100, g.137_138insT	No function
	<i>CYP2D6</i> *17: rs28371706, g.1023C>T; rs16947, g.2850C>T; rs1135840, g.4180G>C	Decreased function
	<i>CYP2D6</i> *29: rs16947, g.2850C>T; rs59421388, g.3183G>A; rs1135840, g.4180G>C	Decreased function
	<i>CYP2D6</i> *31: rs267608319, g.4042G>A; rs16947, g.2850C>T; rs1135840, g.4180G>C	No function

Gene (Transcript) and Brief Summary of Associated Protein Function	Alleles	Predicted Allele Function ^a
	<i>CYP2D6*35</i> : rs769258, g.31G>A; rs16947, g.2850C>T; rs1135840, g.4180G>C; rs1080985, g.-1584C>G	Functional
	<i>CYP2D6*36</i> : a <i>CYP2D6</i> *10 carrying a <i>CYP2D7</i> -derived exon 9 conversion	No function
	<i>CYP2D6*36*10</i> : a <i>CYP2D6*36</i> and a <i>CYP2D6*10</i> in tandem	Decreased function
	<i>CYP2D6*40</i> : rs28371706, g.1023C>T; rs72549356, g.1863_1864ins TTTCGCCCCTTTCGCCCC; rs16947, g.2850C>T; rs1135840, g.4180G>C	No function
	<i>CYP2D6*41</i> : rs16947, g.2850C>T; rs28371725, g.2988G>A; rs1135840, g.4180G>C	Decreased function
	<i>CYP2D6*42</i> : rs16947, g.2850C>T; rs72549346, g.3260_3261insTG; rs1135840, g.4180G>C	No function
	<i>CYP2D6*49</i> : rs1065852, g.100C>T; rs1135822, g.1611T>A; rs1135840, g.4180G>C	Decreased function
	<i>CYP2D6*56</i> : rs72549347, g.3201C>T; rs1135840, g.4180G>C	No function
	<i>CYP2D6*59</i> : rs79292917, g.2939G>A; rs16947, g.2850C>T; rs1135840, g.4180G>C	Decreased function
	<i>CYP2D6*69</i> : rs1065852, g.100C>T; rs16947, g.2850C>T; rs28371725, g.2988G>A; rs1135840, g.4180G>C	No function
	<i>CYP2D6*114</i> : rs1065852, g.100C>T; rs5030865, g.1758G>A; rs16947, g.2850C>T; rs1135840, g.4180G>C	No function
	DUP: complete gene duplication	Varies based on the allele that is duplicated
<i>CYP3A4</i> (NM_017460) Codes for the cytochrome P450 3A4 enzyme, which is involved in metabolism of many medications; impact of the phenotype depends on whether a medication is activated or inactivated by this enzyme	<i>CYP3A4*22</i> : rs35599367, c.522-191C>T	Decreased function
<i>CYP3A5</i> (NM_000777) Codes for the cytochrome P450 3A5 enzyme, which is involved in metabolism of many medications; impact of the phenotype depends on whether a medication is activated or inactivated by this enzyme	<i>CYP3A5*3</i> : rs776746, c.219-237A>G	No function
	<i>CYP3A5*6</i> : rs10264272, c.624G>A	No function
	<i>CYP3A5*7</i> : rs41303343, c.1035dupT	No function
<i>DRD2</i> (NM_000795) Codes for the D2 dopamine receptor that is involved in medication dependence and response to psychotropic medications, particularly antipsychotics	rs1799978, c.-585A>G	Drug dependent ^b
<i>GRIK4</i> (NM_014619) Codes for the ionotropic glutamate receptor also known as AMPA 4, kainate type subunit 4. This receptor is involved in response to psychotropic medications, particularly antidepressants.	rs1954787, c.83-10039T>C	Drug dependent ^b
<i>HTR2A</i> (NM_000621) Codes for a serotonin receptor that is involved in response to many psychotropic medications, including antidepressants and antipsychotics	rs6311, c.-998G>A	Drug dependent ^b
	rs7997012, c.614-2211T>C	Drug dependent ^b
<i>HTR2C</i> (NM_001256760)	rs3813929, c.-850C>T	Drug dependent ^b

Gene (Transcript) and Brief Summary of Associated Protein Function	Alleles	Predicted Allele Function ^a
Codes for a serotonin receptor that is involved in response to psychotropic medications, particularly antipsychotics		
<i>MTHFR</i> (NM_005957) Codes for methylenetetrahydrofolate reductase, an enzyme that is involved in folate metabolism and is implicated in response to L-methylfolate supplementation and response to several psychotropic medications, particularly antipsychotics	rs1801131, c.1286A>C rs1801133, c.665C>T	Drug dependent ^b Drug dependent ^b
<i>OPRM1</i> (NM_000914) Codes for the mu 1 opioid receptor that is involved in response to opioid agonists and antagonists, as well as several other medications	rs1799971, c.118A>G	Drug dependent ^b
<i>UGT2B15</i> (NM_001076) Codes for UDP glucuronosyltransferase family 2 member B15 that is involved in metabolism of many medications, such as the anxiolytics oxazepam and lorazepam	rs1902023, c.253T>G	Drug dependent ^b

^aPredicted allele function based on PharmVar unless otherwise indicated.

^bRefer to ClinPGx.⁴

^cGenomic coordinates for *CYP2D6* variants are based on reference sequence M33388.

Sources: ClinPGx,⁴ PharmVar⁵

Test Interpretation

Analytic sensitivity/specificity is greater than 99%.

Results

- Genetic variant(s) detected: alleles detected are reported. The combination of alleles detected is used to predict phenotype and activity scores, as applicable (eg, *CYP2D6*).
- Samples where *CYP2D6* copy number testing reveals a duplication and where there are two alleles of different activity scores, will be reflexed to LR-PCR for additional testing to reveal which allele is duplicated.
- Phenotype predictions, allele definitions, and associations with specific medications are subject to change as the scientific and clinical evidence evolves.
- When appropriate, results are reported as “star” (*) alleles based on consensus nomenclature. No variants detected is reported as “Negative” and is predictive of *1 functional allele.
- For more information about the association between certain genes and medications, please refer to the ClinPGx [Levels of Evidence for Selected Medication/Gene Associations](#) table.

Limitations

- Only the targeted variants will be detected by this panel. Assumptions about phase and content are made to assign alleles.
- Diagnostic errors can occur due to rare sequence variations.
- A combination of the *CYP2D6**5 (gene deletion) and a *CYP2D6* gene duplication cannot be specifically identified; however, this combination is not expected to adversely affect the phenotype prediction.
- The assay used to detect the *CYP2D6**40 allele cannot distinguish between insertions of one or two copies; it also cannot distinguish between heterozygous and homozygous mutant samples due to unavoidable cross reactivity with the wild type sequence. Additional assays will be used to help differentiate the *CYP2D6**40 allele from other *CYP2D6* star alleles.
- Risk of therapeutic failure or adverse reactions with gene substrates may be affected by genetic and nongenetic factors that are not detected by this test.
- HLA-B*15:02 (an allele that is more common in individuals of Asian descent) and other HLA alleles with pharmacogenetic relevance are not included due to analytical limitations. For additional test options, including testing for HLA-B*15:02 status, refer to the ARUP [Laboratory Test Directory](#).

References

1. Hahn M, Müller DJ, Roll SC. [Frequencies of genetic polymorphisms of clinically relevant gene-drug pairs in a German psychiatric inpatient population.](#) *Pharmacopsychiatry*. 2021;54(2):81-89.
2. U.S. Department of Health and Human Services, Food and Drug Administration. [Table of pharmacogenetic associations](#). Updated May 2022; accessed Aug 2022.
3. CPIC. [Guidelines](#). Stanford University, St. Jude Children's Research Hospital. Accessed Jun 2025.
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ARUP Laboratories is a nonprofit enterprise of the University of Utah and its Department of Pathology. 500 Chipeta Way, Salt Lake City, UT 84108
(800) 522-2787 | (801) 583-2787 | aruplab.com | arupconsult.com