

# GENOMIND® PROFESSIONAL PGx™ ANALYZES TWO TYPES OF GENES TO INFORM OPTIMAL TREATMENT.

The FDA currently includes pharmacogenetic biomarker labeling on over 200 medications due to specific actionable gene-drug associations<sup>3</sup>

**Pharmacodynamic genes indicate the effect a drug has on the body and may inform drug candidate selection.**

	Gene	Physiological Role	Impact of Mutation	Treatment Impact
Pharmacodynamic	<b>Serotonin Transporter (SLC6A4)</b>	Protein responsible for reuptake of serotonin from the synapse	Associated with increased side effects or poorer response to SSRIs	Monitor for adverse events with SSRIs, or <b>assess alternatives to SSRIs</b> . SNRIs or other non-SSRI antidepressants may be considered if clinically indicated
	<b>Serotonin Receptor 2A (HTR2A)</b>	A serotonin receptor which is a target for several serotonergic drugs	Associated with response to certain antidepressants	May prompt consideration of <b>citalopram or non-SSRIs</b>
	<b>Brain-derived Neurotrophic Factor (BDNF)</b>	Important protein for proper neuronal development and neural plasticity	Impaired BDNF secretion, which may be associated with response to some antidepressants or exercise	<b>Increased physical activity/exercise</b> may be more beneficial for <b>Met carriers</b> if clinically indicated. Ethnicity dependent <b>response to SSRIs vs SNRIs</b>
	<b>Major Histocompatibility Complex 1,A (HLA-A 31:01)</b>	Human Leukocyte Antigen-A	Associated with risk of skin reactions to carbamazepine	<b>Do not initiate carbamazepine</b>
	<b>Major Histocompatibility Complex 1,B (HLA-B 15:02)</b>	Human Leukocyte Antigen-B	Associated with risk of skin reactions to carbamazepine, oxcarbazepine, phenytoin, fosphenytoin and possibly lamotrigine, phenobarbital and eslicarbazepine	<b>Do not initiate carbamazepine, oxcarbazepine, phenytoin or fosphenytoin. Caution with lamotrigine, eslicarbazepine, or phenobarbital</b>
	<b>Calcium Channel (CACNA1C)</b>	A subunit of the calcium channel which mediates excitatory signaling	Associated with conditions characterized by mood instability/lability	2nd generation antipsychotics and/or mood stabilizers may be considered if clinically indicated
	<b>Sodium Channel (ANKK3)</b>	Protein that plays a role in sodium channel function and regulation of excitatory signaling	Associated with conditions characterized by mood instability/lability	2nd generation antipsychotics and/or mood stabilizers and/or may be considered if clinically indicated
	<b>Serotonin Receptor 2C (5HT2C)</b>	Receptor involved in regulation of satiety	Associated with differential weight gain risk with 2nd generation antipsychotics	Assess <b>weight gain risk with 2nd generation antipsychotics</b> ; anti-obesity therapies may be considered to mitigate weight gain if clinically indicated
	<b>Melanocortin 4 Receptor (MC4R)</b>	Receptor that plays a role in the control of food intake	Associated with differential weight gain risk with 2nd generation antipsychotics	Assess <b>weight gain risk with 2nd generation antipsychotics</b> ; anti-obesity therapies may be considered to mitigate weight gain if clinically indicated
	<b>Dopamine Receptor D2 (DRD2)</b>	Receptor affected by dopamine in the brain	Associated with slower or poorer response to antipsychotics. Associated with small increased risk of opioid dependence in Asians	<b>Assess dose, alternatives or adjuncts to antipsychotics.</b> Assess non-genetic risk factors for <b>opioid dependence</b> in Asians
	<b>Catechol-O-Methyltransferase (COMT)</b>	Enzyme primarily responsible for the degradation of dopamine in the frontal lobes of the brain	Altered dopamine states can have emotional/behavioral effects and impact response to dopaminergic agents or opioids	<b>Val/Val: Dopaminergic stimulants, COMT inhibitors and/or TMS/ECT</b> may be considered if clinically indicated <b>Decreased sensitivity to opioids</b> <b>Met/Met:</b> Assess alternatives to dopaminergic stimulants. <b>2nd generation antipsychotics</b> may be considered for psychotic-related disorders if clinically indicated <b>Increased sensitivity to opioids</b>
	<b>Alpha-2A Adrenergic Receptor (ADRA2A)</b>	Receptor involved in norepinephrine signaling	Associated with variable response to methylphenidate	<b>Methylphenidate</b> may be used if clinically indicated
	<b>Methylenetetrahydrofolate Reductase (MTHFR, A1298C, C677T)</b>	Predominant enzyme that converts folic acid/folate to its active form (methylfolate) needed for synthesis of serotonin, dopamine, and norepinephrine	Associated with variable activity and conversion of folic acid/folate to methylfolate	Supplementation with <b>L-methylfolate</b> may be considered if clinically indicated
	<b>Glutamate Receptor Kainate 1 (GRIK1)</b>	An excitatory neurotransmitter receptor in the brain	Associated with response to topiramate for alcohol abuse	<b>Topiramate</b> may be considered for treatment of <b>alcohol abuse</b> if clinically indicated
<b>μ-Opioid Receptor (OPRM1)</b>	Opioid receptor affected by endogenous and exogenous opioids	Associated with differential opioid sensitivity. Associated with response to naltrexone for alcohol use disorder	Monitor <b>opioid dose response</b> . <b>Naltrexone</b> consideration for alcohol use disorder	

**Pharmacokinetic genes indicate the effect the body has on the drug and may inform drug dosage.**

Pharmacokinetic	<b>Cytochrome P450 (CYP450: 1A2, 2B6, 2C9, 2C19, 2D6, 3A4/5)</b>	Most psychiatric medications are metabolized by CYP450s	May influence exposure to certain psychotropic medications	<b>Dose adjustment (an increase or decrease) may be considered</b>
	<b>UDP Glucuronosyltransferase (UGT: 1A4, 2B15)</b>	Several psychiatric medications are metabolized by CYP450s	May influence exposure to certain psychotropic medications	<b>Dose adjustment (an increase or decrease) may be considered</b>
	<b>ATP Binding Cassette B1 (ABCB1)</b>	Proteins that impact absorption or brain penetration of certain drugs	Associated with response or sensitivity to select opioids, antipsychotics or antidepressants	<b>Increased exposure possible for select opioids &amp; antipsychotics, as well as citalopram, escitalopram, paroxetine, venlafaxine, amitriptyline, nortriptyline and trimipramine.</b>