

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 associations³

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 | Serotonin Transporter (SLC6A4) | 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 assess alternatives to SSRIs . SNRIs or other non-SSRI antidepressants may be considered if clinically indicated |
| | Serotonin Receptor 2A (HTR2A) | A serotonin receptor which is a target for several serotonergic drugs | Associated with response to certain antidepressants | May prompt consideration of citalopram or non-SSRIs |
| | Brain-derived Neurotrophic Factor (BDNF) | Important protein for proper neuronal development and neural plasticity | Impaired BDNF secretion, which may be associated with response to some antidepressants or exercise | Increased physical activity/exercise may be more beneficial for Met carriers if clinically indicated. Ethnicity dependent response to SSRIs vs SNRIs |
| | Major Histocompatibility Complex 1,A (HLA-A 31:01) | Human Leukocyte Antigen-A | Associated with risk of skin reactions to carbamazepine | Do not initiate carbamazepine |
| | Major Histocompatibility Complex 1,B (HLA-B 15:02) | Human Leukocyte Antigen-B | Associated with risk of skin reactions to carbamazepine, oxcarbazepine, phenytoin, fosphenytoin and possibly lamotrigine, phenobarbital and eslicarbazepine | Do not initiate carbamazepine, oxcarbazepine, phenytoin or fosphenytoin. Caution with lamotrigine, eslicarbazepine, or phenobarbital |
| | Calcium Channel (CACNA1C) | 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 |
| | Sodium Channel (ANKK3) | 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 |
| | Serotonin Receptor 2C (5HT2C) | Receptor involved in regulation of satiety | Associated with differential weight gain risk with 2nd generation antipsychotics | Assess weight gain risk with 2nd generation antipsychotics ; anti-obesity therapies may be considered to mitigate weight gain if clinically indicated |
| | Melanocortin 4 Receptor (MC4R) | Receptor that plays a role in the control of food intake | Associated with differential weight gain risk with 2nd generation antipsychotics | Assess weight gain risk with 2nd generation antipsychotics ; anti-obesity therapies may be considered to mitigate weight gain if clinically indicated |
| | Dopamine Receptor D2 (DRD2) | 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 | Assess dose, alternatives or adjuncts to antipsychotics . Assess non-genetic risk factors for opioid dependence in Asians |
| | Catechol-O-Methyltransferase (COMT) | 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 | Val/Val: Dopaminergic stimulants , COMT inhibitors and/or TMS/ECT may be considered if clinically indicated Decreased sensitivity to opioids Met/Met: Assess alternatives to dopaminergic stimulants. 2nd generation antipsychotics may be considered for psychotic-related disorders if clinically indicated Increased sensitivity to opioids |
| | Alpha-2A Adrenergic Receptor (ADRA2A) | Receptor involved in norepinephrine signaling | Associated with variable response to methylphenidate | Methylphenidate may be used if clinically indicated |
| | Methylenetetrahydrofolate Reductase (MTHFR, A1298C, C677T) | 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 L-methylfolate may be considered if clinically indicated |
| | Glutamate Receptor Kainate 1 (GRIK1) | An excitatory neurotransmitter receptor in the brain | Associated with response to topiramate for alcohol abuse | Topiramate may be considered for treatment of alcohol abuse if clinically indicated |
| | μ-Opioid Receptor (OPRM1) | Opioid receptor affected by endogenous and exogenous opioids | Associated with differential opioid sensitivity. Associated with response to naltrexone for alcohol use disorder | Monitor opioid dose response . Naltrexone consideration for alcohol use disorder |

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

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