

RightMed[®] Comprehensive Test Report

The RightMed Comprehensive Test is a pharmacogenomic test that may aid healthcare providers in determining a therapeutic strategy for a patient. Providers may use this report, along with other clinical factors, to help them when selecting medications and dosages for this patient. This report is not intended to be used in isolation, and the provider needs to take into account all clinical considerations and FDA prescribing information before making any changes to treatment.





Patient and report summary

Patient name: **Sample Patient**
 Patient date of birth: **1977-09-02**
 OneOme report date: **2023-11-29**

Ordering provider: **Sample Doctor**
 Ordering facility: **Sample Clinic (US)**
 Product type: **Comprehensive**
 Report type: **Original**








Report legend

Based on this patient's genetic profile, medications are reported and classified according to the gene-drug interactions described below.

| | | |
|---|---------------------------------------|---|
|  | Major gene-drug interaction | Major genotype-drug interaction identified that affects the metabolism of the medication and/or indicates an elevated risk of adverse reaction or loss of efficacy. |
|  | Moderate gene-drug interaction | Moderate genotype-drug interaction identified that affects the metabolism of the medication and/or indicates an elevated risk of adverse reaction or loss of efficacy. |
|  | Minimal gene-drug interaction | Minimal genotype-drug interaction identified that does not significantly affect medication metabolism nor indicate an elevated risk of adverse reaction or loss of efficacy. |
|  | Limited pharmacogenetic impact | No pharmacogenetic variants demonstrate a significant impact on medication response. Other types of genetic tests that may guide prescribing (e.g., tumor marker testing, diagnostic, or indication-establishing testing) are not taken into account. |

Icon legend

Some medications are reported with icons to indicate that specific clinical annotations and/or dosing guidelines provided by FDA, CPIC, or other professional associations are available in Vantage.

| | | |
|---|-------------------------------|--|
|  | FDA evidence | This medication is listed on the FDA Table of Pharmacogenetic Associations. Not all genetic variants or genetic variant-inferred phenotypes may be accounted for. Further information can be found at: https://www.fda.gov/medical-devices/precision-medicine/table-pharmacogenetic-associations . |
|  | Increased exposure | Total exposure to active compound(s) may be increased. Monitor for adverse effects. |
|  | Decreased exposure | Total exposure to active compound(s) may be decreased. Monitor for lack of therapeutic response. |
|  | Difficult to predict | Total exposure to active compound(s) is difficult to predict. Monitor patient response. |
|  | Reduced response | Response to medication may be lowered due to genetic changes impacting mechanisms other than exposure (e.g. receptor function). |
|  | Additional testing | According to FDA labeling, additional laboratory testing may be indicated. |
|  | Professional guideline | Medication has professional guidelines associated with this patient's genetic test results. Avoidance, dose adjustment, or heightened monitoring may be indicated. |

Report and laboratory comments

Interpretive guidance

Hemizygous males and homozygous females are reported as HTR2C CC.

The reported NUDT15 genotype was inferred based on available variant data. The *2 and *3 alleles are detected by the rs116855232 variant; however, both alleles have no function and *2 is defaulted as *3. Please contact OneOme with any additional questions.

Genotype-predicted interactions for medications

Allergy/Pulmonology

⚠ Major gene-drug interaction

- Dextromethorphan + 1 (Delsym[®])

⚠ Moderate gene-drug interaction

✔ Minimal gene-drug interaction

ℹ Limited pharmacogenetic impact

- Desloratadine (Clarinet[®])
- Montelukast (Singulair[®])

Analgesic/Anti-inflammatory

⚠ Major gene-drug interaction

- Codeine * - 📖 1, 25, 33, 77, 85, 132
- Hydrocodone + 1, 23, 24 (Hysingla[®], Zohydro[®])
- Tramadol * - 📖 1, 25, 33, 85, 132 (Ultram[®])

⚠ Moderate gene-drug interaction

✔ Minimal gene-drug interaction

ℹ Limited pharmacogenetic impact

- Carisoprodol * 1, 42 (Soma[®])
- Celecoxib 📖 1, 128, 132 (Celebrex[®])
- Flurbiprofen 📖 1, 128, 132 (Ansaid[®])
- Ibuprofen * 📖 1, 65, 128 (Advil[®], Motrin[®])
- Meloxicam 📖 1, 128, 132 (Mobic[®])
- Methadone 1, 25, 34 (Dolophine[®], Methadose[®])
- Piroxicam * 📖 1, 105, 106, 128 (Feldene[®])

- Gabapentin (Neurontin[®])
- Naloxone (Evzio[®], Narcan[®])
- Pregabalin (Lyrica[®])

Anesthesiology

⚠ Major gene-drug interaction

⚠ Moderate gene-drug interaction

✔ Minimal gene-drug interaction

ℹ Limited pharmacogenetic impact

- Dexmedetomidine (Precedex[®])

Anticoagulant/Antiplatelet

⚠ Major gene-drug interaction

⚠ Moderate gene-drug interaction

✔ Minimal gene-drug interaction

ℹ Limited pharmacogenetic impact

- Clopidogrel * + 📖 1, 2, 33, 121, 122 (Plavix[®])

- Cilostazol 1, 130 (Pletal[®])
- Ticagrelor 1 (Brilinta[®])
- Warfarin * 1, 18, 59, 60 (Coumadin[®], Jantoven[®])

- Prasugrel (Effient[®])

Cardiovascular

Major gene-drug interaction

- Atorvastatin * + 1, 22, 33, 132 (Lipitor®)
- Carvedilol * + 1 (Coreg®)
- Flecainide + 1, 2 (Tambocor®)
- Lovastatin + 1, 22 (Mevacor®)
- Metoprolol * + 1, 2, 33 (Lopressor®, Toprol XL®)
- Pitavastatin + 1, 22 (Livalo®)
- Propafenone * + 1, 2, 33 (Rythmol®)
- Rosuvastatin * + 1, 22, 132 (Crestor®)
- Simvastatin * + 1, 22, 33, 73, 132 (Zocor®)

Moderate gene-drug interaction

- Fluvastatin + 1, 22, 33 (Lescol®)
- Pravastatin + 1, 22 (Pravachol®)

Minimal gene-drug interaction

- Amiodarone 1 (Cordarone®, Pacerone®)
- Disopyramide 1 (Norpace®)
- Dofetilide 1 (Tikosyn®)
- Losartan 1, 8, 31, 72, 116 (Cozaar®)
- Quinidine 1 (Quin-G®)

Limited pharmacogenetic impact

- Digoxin (Digitek®, Digox®, Lanoxin®)
- Lisinopril (Prinivil®, Zestril®)
- Spironolactone (Aldactone®)

Endocrinology

Major gene-drug interaction

Moderate gene-drug interaction

Minimal gene-drug interaction

Limited pharmacogenetic impact

- Ethinyl estradiol 1, 2
- Nateglinide 1, 132

- Exenatide (Bydureon®, Byetta®)
- Metformin (Fortamet®, Glucophage®)
- Risedronate (Actonel®, Atelvia®)

Gastroenterology

Major gene-drug interaction

Moderate gene-drug interaction

Minimal gene-drug interaction

Limited pharmacogenetic impact

- Dexlansoprazole 1, 75, 132 (Dexilant®)
- Lansoprazole 1, 33, 75, 132 (Prevacid®)
- Metoclopramide * + 1, 9, 132 (Reglan®)
- Omeprazole 1, 33, 37, 75, 90, 132 (Prilosec®)
- Ondansetron + 1, 11, 63, 133 (Zofran®)
- Pantoprazole 1, 33, 75, 132 (Protonix®)

- Dronabinol 1, 132 (Marinol®, Syndros®)
- Fosaprepitant 1, 91 (Emend Injection®)

Genetic disease

Major gene-drug interaction

Moderate gene-drug interaction

Minimal gene-drug interaction

Limited pharmacogenetic impact

- Eliglustat * + 1, 33, 89, 132 (Cerdelga®)

Hematology/Oncology

Major gene-drug interaction

- **Gefitinib** * 1 (Iressa[®])
- **Mercaptopurine** * 1, 2, 3, 15, 111 (Purixan[®])
- **Tamoxifen** * 1, 2, 41 (Soltamox[®])
- **Thioguanine** * 1, 2, 3, 15, 111 (Tabloid[®])

Moderate gene-drug interaction

- **Irinotecan** * 1, 2, 33, 35, 40, 70 (Camptosar[®])
- **Nilotinib** * 1, 5 (Tasigna[®])

Minimal gene-drug interaction

- **Belinostat** 1, 132, 140 (Beleodaq[®])
- **Capecitabine** * 1, 6, 28, 33, 46, 87, 123 (Xeloda[®])
- **Dasatinib** 1 (Sprycel[®])
- **Docetaxel** 1 (Docefrez[®], Taxotere[®])
- **Fluorouracil** * 1, 6, 28, 33, 46, 87, 123 (Adrucil[®])
- **Lapatinib** * 1, 117 (Tykerb[®])
- **Pazopanib** * 1 (Votrient[®])
- **Ruxolitinib** 1 (Jakafi[®])
- **Temsirolimus** 1 (Torisel[®])

Limited pharmacogenetic impact

Immunosuppression

Major gene-drug interaction

- **Azathioprine** * 1, 2, 15, 111 (Imuran[®])

Moderate gene-drug interaction

Minimal gene-drug interaction

- **Cyclosporine** 1 (Gengraf[®], Neoral[®], Sandimmune[®])
- **Sirolimus** 1 (Rapamune[®])
- **Tacrolimus** * 1, 12 (Prograf[®])

Limited pharmacogenetic impact

- **Mycophenolate sodium** (Myfortic[®])

Infectious disease

Major gene-drug interaction

- **Voriconazole** * 1, 2 (Vfend[®])

Moderate gene-drug interaction

- **Atovaquone/Proguanil** 1 (Malarone[®])

Minimal gene-drug interaction

- **Abacavir** * 1, 2, 36, 78, 79, 82, 83, 114, 127 (Ziagen[®])
- **Atazanavir** 38, 58 (Reyataz[®])
- **Efavirenz** * 1, 2, 29, 33, 103 (Sustiva[®])
- **Isavuconazole** 1 (Cresemba[®])
- **Itraconazole** 1 (Onmel[®], Sporanox[®])
- **Peginterferon alfa-2a-containing regimens** 1, 92 (Pegasys[®])
- **Peginterferon alfa-2b-containing regimens** 1, 92 (Pegintron[®])
- **Quinidine** 1 (Quin-G[®])

Limited pharmacogenetic impact

- **Fluconazole** (Diflucan[®])
- **Levofloxacin** (Levaquin[®])
- **Moxifloxacin** (Avelox[®])

Neurology

Major gene-drug interaction

- Amitriptyline * + 1, 2, 33, 49, 50, 141 (Elavil®)
- Metoprolol * + 1, 2, 33 (Lopressor®, Toprol XL®)
- Tetrabenazine * + 1 (Xenazine®)

Moderate gene-drug interaction

- Diazepam * 1, 55 (Valium®)

Minimal gene-drug interaction

- Brivaracetam 1, 132 (Briviact®)
- Carbamazepine 1, 7, 33, 108, 132 (Carbatrol®, Tegretol®)
- Clobazam 1, 132, 142 (Onfi®)
- Eletriptan 1 (Relpax®)
- Eslicarbazepine 1, 7, 64, 108 (Aptiom®)
- Fosphenytoin 1, 2, 7, 17, 19, 80, 97 (Cerebyx®)
- Lamotrigine 1, 7, 80, 108 (Lamictal®)
- Oxcarbazepine * 1, 7, 108 (Trileptal®)
- Phenytoin 1, 2, 7, 17, 19, 80, 97 (Dilantin®)

Limited pharmacogenetic impact

- Gabapentin (Neurontin®)
- Levetiracetam (Keppra®)
- Pramipexole (Mirapex®)
- Pregabalin (Lyrica®)

Psychiatry

Major gene-drug interaction

- Amitriptyline * + 1, 2, 33, 49, 50, 141 (Elavil®)
- Aripiprazole * + 1, 2, 62, 88 (Abilify®)
- Brexpiprazole * + 1, 56 (Rexulti®)
- Chlorpromazine + 1, 104, 126 (Thorazine®)
- Clomipramine * 1, 33, 49, 132 (Anafranil®)
- Desipramine * + 1, 49, 132 (Norpramin®)
- Doxepin * 1, 49 (Silenor®)
- Haloperidol + 1, 2, 102, 124, 135 (Haldol®)
- Iloperidone * + 1 (Fanapt®)
- Imipramine * 1, 33, 49, 118, 119 (Tofranil®)
- Nortriptyline * + 1, 33, 50, 101, 136 (Pamelor®)
- Paroxetine * + 1, 13, 14, 33, 132 (Paxil®)
- Perphenazine * + 1, 100 (Etrafon®)
- Risperidone * + 1, 33, 62, 88, 144 (Risperdal®)
- Thioridazine * + 1
- Trimipramine * 1, 49, 68, 69 (Surmontil®)

Moderate gene-drug interaction

- Amphetamine/
Dextroamphetamine mixed salts 1, 45, 52, 86 (Adderall®)
- Citalopram 1, 13, 14, 33, 132 (Celexa®)
- Clozapine * + 1, 33, 71, 132 (Clozaril®)
- Dextroamphetamine 1, 45, 52, 86 (Dexedrine®)
- Diazepam * 1, 55 (Valium®)
- Escitalopram 1, 13, 14, 33, 132 (Lexapro®)
- Fluoxetine + 1, 33, 39, 43, 48, 53, 57, 74, 76, 81, 107, 109, 120, 125, 143 (Prozac®, Sarafem®)
- Fluvoxamine * + 1, 13, 14, 33, 132
- Lisdexamfetamine 1, 45, 52, 86 (Vyvanse®)
- Sertraline 1, 30, 32, 33, 48, 74, 94, 96, 98, 110, 113, 131, 139 (Zoloft®)
- Venlafaxine * + 1, 2, 33, 138 (Effexor®)

Minimal gene-drug interaction

- Asenapine 1 (Saphris®)
- Bupropion 1, 129, 145 (Wellbutrin®)
- Carbamazepine 1, 7, 33, 108, 132 (Carbatrol®, Tegretol®)
- Cariprazine 1, 4, 16, 20, 95 (Vraylar®)
- Flibanserin 1, 132 (Addyi®)
- Guanfacine 1, 84 (Intuniv®, Tenex®)
- Lamotrigine 1, 7, 80, 108 (Lamictal®)
- Lurasidone 1 (Latuda®)
- Methadone 1, 25, 34 (Dolophine®, Methadose®)
- Nefazodone 1, 112, 137 (Serzone®)
- Nicotine 21, 26, 61, 93 (Nicoderm C-Q®, Nicorette®, Nicotrol®)
- Quetiapine 1, 10, 67, 130, 134 (Seroquel®)
- Trazodone 1 (Desyrel®)





Limited pharmacogenetic impact

- Desvenlafaxine (Pristiq®)
- Lithium (Lithobid®)
- Naloxone (Evzio®, Narcan®)
- Paliperidone (Invega®)
- Temazepam (Restoril®)
- Varenicline (Chantix®)

Psychiatry (cont.)

| | | | |
|---|--|---|--|
|  Major gene-drug interaction |  Moderate gene-drug interaction |  Minimal gene-drug interaction |  Limited pharmacogenetic impact |
| <ul style="list-style-type: none"> ■ Vortioxetine * + 1 (Trintellix®) | | | |





Rheumatology

| | | | |
|---|--|--|--|
|  Major gene-drug interaction |  Moderate gene-drug interaction |  Minimal gene-drug interaction |  Limited pharmacogenetic impact |
| | <ul style="list-style-type: none"> ■ Cevimeline * + 1 (Evoxac®) | <ul style="list-style-type: none"> ■ Allopurinol * 27, 47, 51, 66, 115 (Aloprim®, Zyloprim®) ■ Lesinurad 1 (Zurampic®) ■ Tofacitinib 1 (Xeljanz®) | |

Sleep medicine

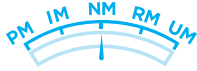
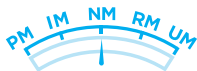
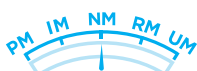





| | | | |
|---|--|---|--|
|  Major gene-drug interaction |  Moderate gene-drug interaction |  Minimal gene-drug interaction |  Limited pharmacogenetic impact |
| <ul style="list-style-type: none"> ■ Doxepin * + 1, 49 (Silenor®) | <ul style="list-style-type: none"> ■ Dextroamphetamine 1, 45, 52, 86 (Dexedrine®) | | <ul style="list-style-type: none"> ■ Temazepam (Restoril®) |

Urology









| | | | |
|--|--|---|--|
|  Major gene-drug interaction |  Moderate gene-drug interaction |  Minimal gene-drug interaction |  Limited pharmacogenetic impact |
| <ul style="list-style-type: none"> ■ Fesoterodine * + 1 (Toviaz®) ■ Tamsulosin * + 1 (Flomax®) | | | |

Genotype-derived classification of medications is provided as a service by OneOme and is intended solely for use by a medical professional who has reviewed and understands all sections within this report, including possible limitations of the services provided by OneOme. The relationships between the drugs and pharmacogenes annotated in this report are supported by scientific evidence that meets OneOme's criteria for inclusion. The order in which drugs are listed does not have any clinical or medical implications. Commonly used trade names for medications are listed for reference only. The list may not be inclusive of all trade names available and does not indicate preference or recommendation by OneOme of one medication product over another. For more information on these medications, for a list of additional medications curated but not annotated by OneOme, or to evaluate possible drug-to-drug interactions, please consult Vantage, which is accessible through the provider portal at portal.oneome.com.


Gene and phenotype summary

| Gene | Genotype | | Phenotype summary / Metabolic status |
|---------------|---------------|---|--|
| CYP1A2 | *1A/*1A |  | Normal metabolizer Fully functional enzyme activity is likely based on the genotype results. Normal metabolism of the medication affected by this gene is predicted. |
| CYP2B6 | *1/*1 |  | Normal metabolizer Fully functional enzyme activity is likely based on the genotype results. Normal metabolism of the medication affected by this gene is predicted. |
| CYP2C9 | *1/*1 |  | Normal metabolizer Fully functional enzyme activity is likely based on the genotype results. Normal metabolism of the medication affected by this gene is predicted. |
| CYP2C19 | *1/*17 |  | Rapid metabolizer Increased enzyme activity is likely based on the genotype results. This activity is more than a normal metabolizer, but less than an ultrarapid metabolizer. The metabolism of the medication affected by this gene is predicted to be increased. |
| CYP2C Cluster | rs12777823 GG |  | Normal CYP2C rs12777823 homozygous wild-type genotype consistent with normal clearance of a certain medication, independent of the impact of CYP2C9*2 and *3. CYP2C rs12777823, together with CYP4F2, CYP2C9, and VKORC1, may affect treatment management of a certain medication. |
| CYP2D6 | *3/*3 |  | Poor metabolizer Little to no enzyme activity is likely based on the genotype results. Little to no metabolism of the medication affected by this gene is predicted. |
| CYP3A4 | *1/*1 |  | Normal metabolizer Fully functional enzyme activity is likely based on the genotype results. Normal metabolism of the medication affected by this gene is predicted. |
| CYP3A5 | *3/*3 |  | Poor metabolizer This CYP3A5 genotype is associated with the phenotype most prevalent in studies used to define standard dosing guidelines. |



Gene and phenotype summary (cont.)

| | | | |
|--------|--------------|---|--|
| CYP4F2 | *1/*1 |  | <p>Normal activity</p> <p>Genotype consistent with normal activity of the CYP4F2 enzyme, which catalyzes the metabolism of vitamin K, in counterpoint to the activity of VKORC1. CYP4F2, together with CYP2C9, VKORC1, and a variant in CYP2C Cluster, may affect treatment management of a certain medication.</p> |
| COMT | rs4680 AA |  | <p>Low activity</p> <p>The AA (Met/Met) genotype is associated with lower COMT activity than the GG (Val/Val) or GA (Val/Met) genotypes.</p> |
| DPYD | *1/*1 |  | <p>Normal metabolizer</p> <p>DPD activity score= 2. This genotype and activity score is consistent with a normal metabolizer phenotype.</p> |
| DRD2 | rs1799978 AA |  | <p>Normal receptor expression</p> <p>Homozygous wild-type dopamine receptor D2 (DRD2) rs1799978 AA genotype is consistent with normal receptor expression.</p> |
| F2 | rs1799963 GG |  | <p>Normal risk</p> <p>Normal risk of thrombosis associated with Factor II (prothrombin). Other genetic and clinical factors contribute to the risk for thrombosis.</p> |
| F5 | rs6025 GG |  | <p>Normal risk</p> <p>Normal risk of thrombosis associated with Factor V. Other genetic and clinical factors contribute to the risk for thrombosis.</p> |
| GRIK4 | rs1954787 TT |  | <p>Altered receptor function</p> <p>Homozygous variant glutamate ionotropic receptor kainate type subunit 4 (GRIK4) genotype is consistent with altered receptor function.</p> |
| HLA-A | Negative |  | <p>Normal risk</p> <p>Negative for the presence of the HLA-A*31:01 allele. Normal risk of hypersensitivity induced by certain medications, and possibly others of structural similarity. Hypersensitivity and severe cutaneous reactions may occur regardless of the presence of the HLA-A*31:01 allele, in particular the presence of the HLA-B*15:02 allele is associated with severe cutaneous reactions induced by certain medications.</p> |

Gene and phenotype summary (cont.)

| | | | |
|---------|---------------|---|---|
| HLA-B | Negative |  | <p>Normal risk</p> <p>Negative for presence of the HLA-B*15:02, HLA-B*57:01, and HLA-B*58:01 alleles. Normal risk of hypersensitivity, severe cutaneous reactions, and severe hepatotoxicity induced by certain medications. Hypersensitivity, severe cutaneous reactions, and severe hepatotoxicity may occur regardless of the presence of HLA-B*15:02, HLA-B*57:01, or HLA-B*58:01 alleles. In particular, the presence of the HLA-A*31:01 allele is associated with hypersensitivity reactions induced by a certain medication, and possibly other medications of structural similarity.</p> |
| HTR2A | rs7997012 AA |  | <p>Intron 2 genotype AA</p> <p>Homozygous wild-type HTR2A (5-hydroxytryptamine receptor 2A) genotype is consistent with normal HTR2A receptor function.</p> |
| HTR2C | rs3813929 CC |  | <p>Normal receptor expression</p> <p>This genotype is associated with normal transcriptional activity of the HTR2C gene.</p> |
| IFNL4 | rs12979860 CC |  | <p>Normal</p> <p>Genotype consistent with a normal likelihood of hepatitis C sustained virologic response (SVR) with certain treatment options.</p> |
| NUDT15 | *1/*3 |  | <p>Intermediate metabolizer</p> <p>NUDT15 genotype is consistent with impaired enzyme activity and is associated with an increased risk of thiopurine-induced toxicities.</p> |
| OPRM1 | rs1799971 AA |  | <p>Variant absent</p> <p>The AA genotype (or Asn/Asn isoform) is the wildtype genotype associated with the mu-1 opioid receptor.</p> |
| SLC6A4 | L/L (La/La) |  | <p>Typical to increased expression</p> <p>Genotype consistent with a typical to increased expression of the SLC6A4 transporter compared to other genotypes. This genotype was shown to exhibit different phenotypes in East Asian populations, as opposite outcomes were observed for this genotype in East Asian populations when compared to Caucasian populations.</p> |
| SLCO1B1 | *5/*5 |  | <p>Poor function</p> <p>SLCO1B1 genotype consistent with poor function of the OATP1B1 transporter.</p> |
| TPMT | *1/*1 |  | <p>Normal metabolizer</p> <p>TPMT genotype is consistent with a normal metabolizer phenotype and is not associated with an increased risk of thiopurine-induced toxicities. Toxicities with thiopurines can occur due to impaired NUDT15 activity independently from the TPMT activity.</p> |

Gene and phenotype summary (cont.)

| | | | |
|--------|--------------|---|--|
| UGT1A1 | *1/*28 |  | <p>Intermediate metabolizer (Heterozygous *28)</p> <p>Genotype consistent with decreased UGT1A1 enzyme activity, or an intermediate metabolizer phenotype, and is associated with an increased risk of certain drug-induced toxicities.</p> |
| VKORC1 | rs9923231 GG |  | <p>Normal activity</p> <p>Genotype consistent with normal activity of the vitamin K epoxide reductase enzyme, associated with c.-1639GG (rs9923231). VKORC1, together with CYP2C9, CYP4F2, and a variant in CYP2C Cluster, may affect treatment management of a certain medication.</p> |

CYP phenotype abbreviations

| | |
|-----------|--------------------------|
| PM | Poor metabolizer |
| IM | Intermediate metabolizer |
| NM | Normal metabolizer |
| RM | Rapid metabolizer |
| UM | Ultrarapid metabolizer |

Test information

| | | |
|------------------------------------|---|---|
| Specimen ID: 2350706821927 | Clinical testing performed by: OneOme | Reported by: Cathryn Jennissen in None |
| Specimen type: Buccal swab | 807 Broadway St. NE Suite 100 | CLIA: 24D2109855 |
| Collection date: 2023-11-27 | Minneapolis, MN 55412, United States | CAP: 9432670 |
| Receive date: 2023-11-29 | | NY PFI: 9226 |

Test results

The following analytical results were interpreted by OneOme to produce the pharmacogenomic interpretations and annotations described in the *Gene and phenotype summary*. Method-specific analytical limitations or inferred haplotypes may limit the ability to produce a definitive phenotype interpretation. See *Methodology and limitations* and/or the *Report and laboratory comments* sections for additional information.

| | | | | |
|------------------------------------|-------------------------------------|-----------------------------|------------------------------|----------|
| CYP1A2 *1A/*1A | rs5030656 | NM_000106.5:c.841_843delAAG | AAGAAG | |
| rs762551 | NM_000761.4:c.-9-154C>A | rs35742686 | NM_000106.5:c.775delA | -- |
| rs2069514 | NG_008431.2:g.28338G>A | rs72549353 | NM_000106.5:c.765_768delAACT | AACTAACT |
| rs2069526 | NM_000761.4:c.-10+103T>G | rs5030655 | NM_000106.5:c.454delT | TT |
| rs12720461 | NM_000761.4:c.-10+113C>T | rs774671100 | NM_000106.5:c.137_138insT | -- |
| rs35694136 | NM_000761.4:c.-1635delT | rs1080985 | NM_000106.5:c.-1584C>G | CC |
| | | rs59421388 | NM_000106.5:c.1012G>A | GG |
| | | rs28371725 | NM_000106.5:c.985+39G>A | GG |
| | | rs72549346 | NM_000106.5:c.1088_1089insGT | -- |
| | | rs5030865 | NM_000106.5:c.505G>[A,T] | GG |
| CYP2B6 *1/*1 | | CYP3A4 *1/*1 | | |
| rs3745274 | NM_000767.4:c.516G>T | rs2740574 | NM_017460.5:c.-392G>A | AA |
| rs2279343 | NM_000767.4:c.785A>G | rs35599367 | NM_017460.5:c.522-191C>T | CC |
| rs34223104 | NM_000767.5:c.-82T>C | | | |
| rs3211371 | NM_000767.4:c.1459C>T | | | |
| rs36079186 | NM_000767.5:c.593T>C | | | |
| rs28399499 | NM_000767.4:c.983T>C | | | |
| CYP2C9 *1/*1 | | CYP3A5 *3/*3 | | |
| rs7900194 | NM_000771.3:c.449G>A | rs776746 | NM_000777.4:c.219-237G>A | GG |
| rs1799853 | NM_000771.3:c.430C>T | rs10264272 | NM_000777.4:c.624G>A | GG |
| rs1057910 | NM_000771.3:c.1075A>C | rs41303343 | NM_000777.4:c.1035_1036insT | -- |
| rs28371686 | NM_000771.3:c.1080C>G | | | |
| rs56165452 | NM_000771.3:c.1076T>C | | | |
| rs28371685 | NM_000771.3:c.1003C>T | | | |
| rs9332131 | NM_000771.3:c.817delA | | | |
| CYP2C19 *1/*17 | | CYP4F2 *1/*1 | | |
| rs12248560 | NM_000769.2:c.-806C>T | rs2108622 | NM_001082.4:c.1297G>A | GG |
| rs4244285 | NM_000769.2:c.681G>A | | | |
| rs4986893 | NM_000769.2:c.636G>A | | | |
| rs6413438 | NM_000769.2:c.680C>T | | | |
| rs28399504 | NM_000769.2:c.1A>G | | | |
| CYP2C Cluster rs12777823 GG | | COMT rs4680 AA | | |
| rs12777823 | NC_000010.10:g.96405502G>A | rs4680 | NM_000754.3:c.472G>A | AA |
| CYP2D6 *3/*3 | | DPYD *1/*1 | | |
| rs28371706 | NM_000106.5:c.320C>T | rs75017182 | NM_000110.4:c.1129-5923C>G | CC |
| rs267608319 | NM_000106.5:c.1319G>A | rs55886062 | NM_000110.3:c.1679T>G | TT |
| rs16947 | NM_000106.5:c.886C>T | rs67376798 | NM_000110.3:c.2846A>T | TT |
| rs79292917 | NM_000106.5:c.975G>A | rs3918290 | NM_000110.3:c.1905+1G>A | GG |
| rs1065852 | NM_000106.5:c.100C>T | rs115232898 | NM_000110.4:c.557A>G | AA |
| rs1135840 | NM_000106.5:c.1457G>C | | | |
| rs3892097 | NM_000106.5:c.506-1G>A | | | |
| rs769258 | NM_000106.5:c.31G>A | | | |
| rs5030862 | NM_000106.5:c.124G>A | | | |
| rs201377835 | NM_000106.5:c.181-1G>C | | | |
| rs5030867 | NM_000106.5:c.971A>C | | | |
| rs765776661 | NM_000106.5:c.1411_1412insTGCCCACTG | | | |
| | | DRD2 rs1799978 AA | | |
| | | rs1799978 | NM_000795.3:c.-585A>G | AA |
| | | F2 rs1799963 GG | | |
| | | rs1799963 | NM_000506.4:c.*97G>A | GG |
| | | F5 rs6025 GG | | |
| | | rs6025 | NM_000130.4:c.1601G>A | GG |
| | | GRIK4 rs1954787 TT | | |
| | | rs1954787 | NM_001282470.2:c.83-10039T>C | TT |
| | | HLA-A Negative | | |

Test results (cont.)

| | | | |
|----------------------------|---|----------|---|
| HLA00097 | NM_002116 (interrogated at exon 2) | Negative | SLC6A4 L/L (La/La) |
| HLA-B Negative | | | rs25531 NM_001045.5:c.-1936A>G AA |
| rs144012689 | NM_005514.7:c.1012+104A>T | TT | rs774676466 NM_001045.5:c.-1917_-1875del43 LL |
| HLA00386 | NM_005514 (interrogated at exon 2 and intron 2) | Negative | SLCO1B1 *5/*5 |
| HLA00381 | NM_005514 (interrogated at exon 3) | Negative | rs2306283 NM_006446.4:c.388A>G AA |
| HTR2A rs7997012 AA | | | rs4149056 NM_006446.4:c.521T>C CC |
| rs7997012 | NM_000621.4:c.614-2211T>C | TT | rs4149015 NM_006446.4:c.-910G>A GG |
| HTR2C rs3813929 CC | | | TPMT *1/*1 |
| rs3813929 | NM_000868.3:c.-759C>T | CC | rs1142345 NM_000367.3:c.719A>G AA |
| IFNL4 rs12979860 CC | | | rs1800584 NM_000367.3:c.626-1G>A CC |
| rs12979860 | NM_001276254.2:c.151-152G>A | CC | rs1800462 NM_000367.3:c.238G>C GG |
| NUDT15 *1/*3 | | | rs1800460 NM_000367.3:c.460G>A GG |
| rs116855232 | NM_018283.3:c.415C>T | CT | UGT1A1 *1/*28 |
| OPRM1 rs1799971 AA | | | rs4148323 NM_001072.3:c.862-6536G>A GG |
| rs1799971 | NM_000914.4:c.118A>G | AA | rs1976391 NM_001072.3:c.862-9697A>G AG |
| | | | VKORC1 rs9923231 GG |
| | | | rs7200749 NM_024006.5:c.358C>T GG |
| | | | rs9923231 NM_001311311.1:c.-1639G>A GG |

Electronically signed by:
Cathryn Jennissen in None
2023-11-29

Methodology and limitations

This test was developed and its performance characteristics determined by OneOme, LLC, a clinical laboratory located at 807 Broadway Street NE Suite 100, Minneapolis, MN 55413. These tests have not been cleared or approved by the U.S. Food and Drug Administration. The FDA does not require this test to go through premarket FDA review. OneOme is certified under CLIA-88 and accredited by the College of American Pathologists as qualified to perform high-complexity testing. This test is used for clinical purposes and should not be regarded as investigational or for research.

Genomic DNA was analyzed by PCR using Thermo Fisher TaqMan® and/or LGC Biosearch BHQ® probe-based methods to interrogate the variant locations listed in the *Test results* table above. In addition, CYP2D6 copy number status was assessed at sites within the promoter, intron 2, intron 6, and exon 9. The test detects CYP2D6 deletions, duplications/multiplications, and hybrid alleles, but cannot differentiate duplications in the presence of a deletion.

Haplotypes, or combinations of inherited variants on a chromosome, are annotated according to legacy nomenclature for the genes and alleles in the table below.

| | |
|---------|--|
| CYP1A2 | *1C, *1D, *1E, *1F, *1J, *1K, *1L, *1V, *1W |
| CYP2B6 | *4, *5, *6, *7, *9, *18, *22, *35, *36 |
| CYP2C9 | *2, *3, *4, *5, *6, *8, *11 |
| CYP2C19 | *2, *3, *4, *4B, *10, *17 |
| CYP2D6 | *2, *2A, *3, *4, *4M, *4N, *5, *6, *6C, *7, *8, *9, *10, *11, *12, *13, *14, *15, *17, *18, *19, *29, *31, *34, *35, *36, *39, *41, *42, *59, *63, *64, *68, *69, *70, *91, *109, *114 |
| CYP3A4 | *1B, *22 |
| CYP3A5 | *3, *6, *7 |
| CYP4F2 | *3 |
| DPYD | *2A, Asp949Val, *13, HapB3, Tyr186Cys |
| SLCO1B1 | *1B, *5, *15, *17, *21 |
| TPMT | *2, *3A, *3B, *3C, *4 |
| UGT1A1 | *6, *28 |

The test does not detect all known and unknown variations in the gene(s) tested, nor does absence of a detectable variant (designated as *1 for genes encoding drug metabolizing enzymes) rule out the presence of other, non-detected variants.

As with other common SNP genotyping techniques, these assays cannot differentiate between the maternal and paternal chromosomes. In cases where observed variants are associated with more than one haplotype, OneOme infers and reports the most likely diplotype based on published allele frequency and/or ethnicity data. Inferences with potential clinical impact are reported in the *Report and laboratory comments* section.

PCR may be subject to general interference by factors such as reaction inhibitors and low quality or quantity of extracted DNA. When present, these interferences typically yield no result rather than an inaccurate one. Very infrequent variants or polymorphisms occurring in primer- or probe-binding regions may affect testing and could produce an erroneous result or assay failure. Variant locations tested by the assay but not assigned a genotype call are reported as “No Call.”

The variant detection methods validated by OneOme provide >99.9% accuracy for the adult population; however, clinical interpretation may be inaccurate for patients who have undergone or are receiving non-autologous blood transfusions, tissue, and/or organ transplant therapies. Although extremely rare, results could also be impacted by other factors not addressed above, such as laboratory error.

Pharmacogenetic correlation is largely based on studies of adult populations. Gene-drug guidance may not be informative in pediatric patients. For patients that may carry a probabilistic risk of disease, patients and providers should consider the benefits of consulting with a trained genetic counseling professional, physician, or pharmacogenomic specialist. For additional support, contact OneOme

Methodology and limitations (cont.)

through the website or by calling 844-663-6635.

OneOme liability disclaimer

The interpretations and clinical annotations provided by OneOme are intended solely for use by a medical professional in the treatment of adult patients and do not constitute medical advice by OneOme. The treating provider remains ultimately responsible for all diagnosis and treatment decisions for the patient. OneOme disclaims liability for any errors, omissions or ambiguities in any translation or interpretation of a report by a third party, including without limitation direct, indirect, incidental, special, consequential or exemplary damages, whether such damages arise in contract, negligence, tort, under statute, in equity, at law or otherwise. Information included in this report is based upon scientific literature, including information from and guidelines published by professional associations (e.g., CPIC, FDA, DPWG), and does not take into account other genetic variants and environmental or social factors that may affect a patient's response. Other factors not included in this report include, but are not limited to, environmental factors (e.g., smoking), health factors (e.g., diet), social and familial factors, various medical conditions, and drug-to-drug interactions. Administration of any medication, including the ones listed in the OneOme reports, requires careful therapeutic monitoring regardless of the phenotype or genotype-predicted interaction reported. As a matter of practice, OneOme will routinely update its pharmacogenomic database as new information becomes available to the scientific community. Genotype-predicted interactions and annotations found on the patient's RightMed Comprehensive Test Report, Vantage Reports, or RightMed specialty reports are therefore dependent on the date of generation and/or the database version used to generate that report. Providers may access these reports with updated annotations using OneOme's latest released version through the provider portal at portal.oneome.com.

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