

Patient	XXX	Requesting physician	
Date of birth	01/06/1956 Sex F		
Sample type	DNA	Report generated	29/08/2022
Collection date	10/06/2022	Laboratory director	Dr C. Lapucci
Received date	28/06/2022	Contact email	cristina.lapucci@synlab.it
Sample number	xxx		

MyPGx[®] - Pharmacogenetic short screening panel (method: PCR, MassArray and MLPA)

Provided clinical information:

Clinical information	
Known problematic medication	NKDA
Relevant medical history	None

Summary of key pharmacogenetic results (predicted Poor or Ultrarapid activity):

Gene	Prediction
CYP2C19	Rapid metabolizer
VKORC1	Warfarin resistance
SLC22A1	Low function
SLC01B3	Low function
SULT1A1	Poor metabolizer
NAT2	Poor acetylator

The detailed pharmacogenetic results are presented on the following pages.

Technical comments and limitations:

Coverage 100%. Haplotypes not determined (failed SNPs): None

PGx is a rapidly-evolving field primarily providing evidence-based predictions of how the tested individual's genetic profile may affect reaction to certain drugs. Factors such as drug-drug interaction and also age, diet, ethnicity, family and personal health history, can also impact the likelihood of exhibiting certain drug reactions, independently of genotype-based predictions.

This report is intended for use by a healthcare professional. Based on PGx results, **patients should make no changes to medical care without the prior advice of and consultation with a healthcare professional** [including, but not limited to, changes in dosage or frequency of medication, diet and/or exercise regimens, or pregnancy planning].

ELECTRONIC SIGNATURE	Dott.ssa Cristina Lapucci SPECIALISTA IN GENETICA MEDICA SYNLAB ITALIA
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GENOTYPE/HAPLOTYPE/PHENOTYPE DETAIL

Gene	Genotype-Haplotype	Allele Tested	Predicted Phenotype
CYP1A1	*1/*1	*1, *2, *3, *4, *5, *6, *7, *8	Normal metabolizer
CYP1A2	*1A/*1F	*1A, *1F, *1K, *7	Normal metabolizer
CYP2A6	*1A/*9	*1A, *1B, *2, *4, *5, *6, *7, *8, *9, *11, *17, *20	Intermediate metabolizer
CYP2B6	*6/*6	*1, *6, *8, *10, *18, *28	Intermediate metabolizer
CYP2C8	*1/*1	*1, *2, *3, *4, *5, *7, *8	Normal metabolizer
CYP2C9	*1/*3	*1, *2, *3, *4, *5, *6, *8, *9, *10, *11, *12, *13, *15, *25, *27	Intermediate metabolizer
CYP2C19	*1A/*17	*1A, *1B, *2A, *3, *4, *5A, *5B, *6, *7, *8, *12, *17	Rapid metabolizer
CYP2D6	*2A/*41	*1, *2A, *3, *4A, *4M, *5, *6A, *7, *8, *9, *10, *11, *12, *14A, *14B, *17, *18, *19, *20, *21, *36, *38, *40, *41, *42, *44, *56A, *56B, and CNVs	Normal metabolizer
CYP2E1	*1/*7	*1, *2, *7	Normal metabolizer
CYP3A4	*1/*1	*1, *2, *6, *20, *22	Normal metabolizer
CYP3A5	*1A/*1A	*1A, *3A, *3K, *5, *6, *7	Normal metabolizer
VKORC1	H7/H7	H1, H3, H7, H9	Warfarin resistance
SLC15A2	*1/*1	*1, *509K, *284A, *350F, *409S	Normal function
SLC22A1	*420Del/*420Del	*1, *2, *3, *4, *5, *6, *220V, *283L, *287G, *341L, *408V, *420Del	Low function
SLC22A2	*1/*270A	*1, *54S, *165V, *270A, *400C, *432N	Normal function
SLC22A6	*1/*1	*1, *50H	Normal function
SLCO1B1	*1A/*1A	*1A, *1B, *2, *3, *5, *6, *9, *10, *11, *12, *13, *15	Normal function
SLCO1B3	*233I/*233I	*1, *112A, *233I	Low function
SLCO2B1	*1/*1	*1, *3	Normal function
ABCB1	*1/*2	*1, *2	Intermediate function
ABCC2	*1/*1324I	*1, *417I, *789F, *768W, *1324I, *1450S	Intermediate function
ABCG2	*1/*1	*1, *141K, *126Ter	Normal function
SULT1A1	*3/*3	*1, *2, *3, *4	Poor metabolizer
NAT1	*4/*11	*1, *5, *11, *14, *15, *17, *19, *22	Normal acetylator
NAT2	*5B/*6A or *5A/*6C or *6B/*5G or *12C/*5J	*4, *5A, *5B, *5C, *5D, *5E, *5G, *5J, *6A, *6B, *6C, *6E, *7A, *7B, *11A, *12A, *12B, *12C, *13, *14A, *14B, *14C, *14D, *14E, *14F, *14G, *19	Poor acetylator
TPMT	*1/*1	*1, *2, *3A, *3B, *3C, *4, *8	Normal metabolizer
GSTM1	*1/*1	*1, *173Asn	Normal metabolizer
GSTP1	*1A/*1A	*1A, *1B, *1D, *1C	Normal metabolizer
UGT1A1	*28(*60)/*28(*60)	*1, *6, *7, *27, *29, *60	Intermediate metabolizer
UGT2B7	*1a/*1a	*1a, 2b	Normal metabolizer
UGT2B15	*2/*2	*1, *2	Intermediate metabolizer
DPYD	*1/*2A	*1, *2A, *7, *8, *9A, *9B, *10	Intermediate metabolizer

Disclaimer: Laboratory-developed screening test and interpretation protocols, employing research-use only (RUO) materials. The result of the "Phenotype" shown in the table is to be considered as a generic parameter and not specific to a single drug. For a reference to a specific drug check the tables below in the report. Additional Disclaimer: MyPGx® is a registered trade mark of SYNLAB International GmbH. **Patients should not initiate or modify any treatment or otherwise use the information in this report without the prior advice, consultation and supervision of a licensed healthcare professional such as a pharmacist or medical doctor.**

Methodology: PCR-based RUO assays detect listed alleles, including all common and most rare variants with known clinical significance at analytical sensitivity and specificity >99%. Phenotypic predictions based on the current state of the scientific literature and PharmGKB. MLPA assay to identify CYP2D6 CNVs (deletions, duplications).

Limitations: Testing cannot detect all genetic variants, inactive or altered genes. The absence of a finding of a detectable gene or variant does not necessarily indicate patient possesses intermediate- or high-sensitivity phenotypes or that patient has an undetected variant. Drug-drug interactions may significantly modify phenotypes, especially in polymedicated patients. The lack of data in the literature and the absence of clinical reports do not allow any ambiguous variants and related phenotypic consequences to be adequately characterised. For these reasons, in the absence of this information, the result is given on the basis of haplotype frequencies.

PHARMACOGENOMICS

Genetic Markers Tested for Pharmacogenomics:

Results are arranged by drug response. Each individual report contains six sections, including: Patient's current medication (if any), Medication history, genotype/haplotype/phenotype detail, PGx report, Genomic Test Results, and Patient Information Card. Inclusion of the PGx Report indicates that the tested individual: displays decreased efficacy to the drug (yellow dots), should use the drug as directed (green dots), or exhibits increased toxicity to the drug (orange dots). Inclusion of Genomic Test Results indicates genotype, haplotype, phenotype, or presence of mutation.

Organization of Table:

- Gene/Locus refers to gene or intergenic region of genetic marker location.
- Marker refers to the tested marker's unique identifier.
- Genotype/Haplotype refers to the particular marker's combination of nucleotides. The letter(s) on either side of the slash refer(s) to the two (2) copies of the patient DNA. Del and dashes denotes nucleotide indels in patient DNA. Empty cells indicate an absence of genotyping results.
- Phenotype refers to the CYP specific drug metabolizing capabilities of an individual.

See RISKS AND LIMITATIONS on the last pages of this Report.

PGx Report - Pain Management

Type: Anti-inflammatory Agent, Analgesic, Antipyretic

Drug Class	Substance	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
The Nonsteroidal Antiinflammatory Drugs (NSAIDs)						
Acetic acid derivatives	Diclofenac	UGT2B7	CYP2C9, CYP2E1, CYP3A4		✔	
	Nabumetone	CYP1A2	CYP2C19, CYP3A4		✔	
	Indomethacin	CYP2C9	CYP2C19			⚠
Enolic acid (Oxicam) derivatives	Meloxicam	CYP2C9	CYP1A2, CYP3A4, CYP3A5			⚠
	Piroxicam	CYP2C9	CYP3A4, CYP3A5			⚠
	Tenoxicam	CYP2C9				⚠
	Lornoxicam	CYP2C9				⚠
Selective COX-2 inhibitors (Coxibs)	Etoricoxib	CYP3A4	CYP3A5, CYP2C9, CYP2D6, CYP1A2		✔	
	Parecoxib	CYP2C9	CYP3A4, CYP3A5			⚠
	Celecoxib	CYP2C9	CYP2C19			⚠
Propionic acid derivatives	Ibuprofen	CYP2C9	CYP2C19, CYP2C8, UGT2B7		✔	
	Flurbiprofen	CYP2C9				⚠
	Ketoprofen	CYP3A4	CYP2C9, CYP3A5, UGT2B7		✔	
	Fenoprofen	CYP2C9	UGT2B7			⚠
	Vicoprofen	CYP2D6	CYP3A4		✔	
	Naproxen	CYP2C9	CYP1A2, CYP2C8, UGT2B7, SULT1A1			⚠
Anthranilic acid derivatives (Fenamates)	Mefenamic acid	CYP2C9				⚠
The Non-NSAIDs Analgesic	Acetaminophen	UGT1A1, SULT1A1, GSHs	CYP2E1, CYP3A4, CYP3A5, CYP2D6, CYP1A2, ABCG2		✔	

PGx Report - Pain Management

Type: Opioid

Drug Class	Substance	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
Opioid Analgesics						
Opium alkaloids	Morphine	UGT2B7	ABCB1, UGT1A1, COMT		✔	
	Codeine	CYP2D6	CYP3A4, UGT2B7, CYP3A5		✔	
Ethers of morphine	Dihydrocodeine	CYP3A4	CYP2D6, CYP3A5		✔	
	Ethylmorphine	CYP2D6	CYP3A4, CYP3A5		✔	
Semi-synthetic alkaloid derivatives	Hydrocodone	CYP2D6	CYP3A4, CYP3A5		✔	
	Hydromorphone	UGT2B7			✔	
	Oxycodone	CYP3A4	CYP3A5, CYP2D6, ABCB1, UGT2B7, COMT		✔	
	Oxymorphone	UGT2B7			✔	
Synthetic opioids						
Anilidopiperidine derivatives	Alfentanil	CYP3A4	CYP3A5, ABCB1		✔	
	Fentanyl	CYP3A4	CYP3A5, ABCB1		✔	
	Sufentanil	CYP3A4	CYP3A5		✔	
Phenylpiperidine derivatives	Meperidine	CYP2B6	CYP3A4, CYP2C19, CYP3A5			⚠
	Ketobemidone	CYP2C9	CYP3A4, CYP3A5			⚠
Diphenylpropylamine derivatives	Dextropropoxyphene	CYP3A4	CYP3A5, Renal Excretion		✔	
	Levacetylmethadol	CYP3A4	CYP3A5		✔	
	Methadone	CYP3A4	CYP2B6, CYP2D6, CYP3A5, ABCB1, UGT2B7, COMT		✔	
Oripavine derivatives	Buprenorphine	CYP3A4	CYP3A5, CYP2C8, UGT1A1, UGT2B7		✔	
Morphinan derivatives	Dextromethorphan	CYP2D6	CYP3A4, CYP3A5		✔	
Others	Tramadol	CYP2D6	CYP3A4, CYP2B6, CYP3A5, SLC22A1, COMT	⚠		
	Tapentadol	CYP2C9	CYP2C19, CYP2D6		✔	
	Tilidine	CYP3A4	CYP2C19, CYP3A5		✔	
Anti-opioid	Methylnaltrexone	CYP2D6	CYP3A4, CYP3A5		✔	
	Naltrexone	UGT2B7	UGT1A1		✔	

PGx Report - Pain Management

Type: Drugs Prescribed for the Treatment of Gout, Antirheumatic

Drug Class	Substance	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
Drugs Prescribed for Gout						
Uricosurics	Sulfinpyrazone	CYP2C9	CYP3A4, CYP3A5		✔	
Mitotic inhibitors	Colchicine	CYP3A4	CYP3A5		✔	
Xanthine oxidase inhibitors	Febuxostat	CYP1A2, CYP2C8	CYP2C9, UGT1A1, UGT2B7		✔	
	Allopurinol	AOX1	Renal Excretion, HLA-B*5801		✔	
	Oxypurinol	Renal Excretion			✔	
Recombinant urate oxidase	Rasburicase		G6PD, CYB5R1, CYB5R2, CYB5R3, CYB5R4		✔	
Antimetabolites	Azathioprine	XO	TPMT, AOX1		✔	
	Methotrexate	Renal Excretion	AOX1, SLC01B1, SLC19A1, ABCC1, ABCC2, ABCC3, ABCG2		✔	
DMARDs	Leflunomide	CYP1A2			✔	
Anti-inflammatory	Tofacitinib	CYP3A4	CYP2C19, CYP3A5		✔	

Abbreviations: DMARDs, Disease-modifying antirheumatic drugs; RE, renal excretion (unchanged drug).

Additional SNPs of Importance for Pain Management

Gene	Marker	Genotype	Drug	Level of Evidence	Results
COMT	rs4680	G/G	Paroxetine	3	Patients may require a higher dose

Patient Information Card

An easily portable summary of the report patients can share with their medical professionals. (Please cut along dotted line.)



Pharmacogenomic Test Summary

CYP1A1	*1/*1	Normal metabolizer
CYP1A2	*1A/*1F	Normal metabolizer
CYP2A6	*1A/*9	Intermediate metabolizer
CYP2B6	*6/*6	Intermediate metabolizer
CYP2C8	*1/*1	Normal metabolizer
CYP2C9	*1/*3	Intermediate metabolizer
CYP2C19	*1A/*17	Rapid metabolizer
CYP2D6	*2A/*41	Normal metabolizer
CYP2E1	*1/*7	Normal metabolizer
CYP3A4	*1/*1	Normal metabolizer
CYP3A5	*1A/*1A	Normal metabolizer
VKORC1	H7/H7	Warfarin resistance
SLC15A2	*1/*1	Normal function
SLC22A1	*420Del/*420Del	Low function
SLC22A2	*1/*270A	Normal function
SLC22A6	*1/*1	Normal function
SLCO1B1	*1A/*1A	Normal function
SLCO1B3	*233I/*233I	Low function
SLCO2B1	*1/*1	Normal function
ABCB1	*1/*2	Intermediate function
ABCC2	*1/*1324I	Intermediate function
ABCG2	*1/*1	Normal function
SULT1A1	*3/*3	Poor metabolizer
NAT1	*4/*11	Normal acetylator
NAT2	*5B/*6A or *5A/*6C or *6B/*5G or *12C/*5J	Poor acetylator
TPMT	*1/*1	Normal metabolizer
GSTM1	*1/*1	Normal metabolizer
GSTP1	*1A/*1A	Normal metabolizer
UGT1A1	*28(*60)/*28(*60)	Intermediate metabolizer
UGT2B7	*1a/*1a	Normal metabolizer
UGT2B15	*2/*2	Intermediate metabolizer
DPYD	*1/*2A	Intermediate metabolizer

For a complete report contact Synlab.com