

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
SLCO1B3	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,	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	*2331/*2331	*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, *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

<u>Disclaimer: Laboratory-developed</u> 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:

- 1. Gene/Locus refers to gene or intergenic region of genetic marker location.
- 2. Marker refers to the tested marker's unique identifier.
- 3. 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.
- 4. 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 - Internal Medicine

Type: Antiemetic

Drug Class	Substance	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antie	emetic			
Antiemetic, 5-HT3 receptor	<u>Dolasetron</u>	CYP3A4	CYP2D6, CYP3A5			
antagonist Indole derivative	<u>Tropisetron</u>	CYP3A4	CYP2D6, CYP3A5		Ø	
Antiemetic, 5-HT3 receptor antagonist Isoquinoline derivative	<u>Palonosetron</u>	CYP1A2	CYP2D6, CYP3A4, CYP3A5		Ø	
Antiemetic, 5-HT3 receptor antagonist Indazole derivative	Granisetron	СҮРЗА4	СҮРЗА5		Ø	
Antiemetic, 5-HT3 receptor antagonist	<u>Ondansetron</u>	CYP2B6	CYP1A2, CYP2D6, CYP3A4, ABCB1			
_	<u>Domperidone</u>	CYP3A4	CYP3A5		Ø	
Antiemetic, dopamine-		CYP2D6	CYP3A4, CYP3A5			
receptor antagonist	<u>Metoclopramide</u>	CYP2D6	CYP1A2, CYB5R1, CYB5R2, CYB5R3, CYB5R4		Ø	
Antiemetic, NK1 receptor antagonist	<u>Aprepitant</u>	CYP3A4	CYP3A5, CYP1A2, CYP2C19		0	
	<u>Diphenhydramine</u>	CYP2D6	CYP3A4, CYP3A5, UGT1A3, UGT1A4			
Antiemetic, H1 histamine receptor antagonist	<u>Hydroxyzine</u>	ADHs	CYP3A4, CYP3A5			
Teceptor untagonist	Promethazine	CYP2D6	SULTs			
Cannabinoids	<u>Dronabinol</u>	CYP2C9	CYP2C19, CYP3A4, CYP3A5			
	Lorazepam	UGT2B15	UGT2B7			
Benzodiazepines	<u>Midazolam</u>	CYP3A4	СҮРЗА5			
Anticholinergics	Scopolamine	CYP3A4	СҮРЗА5			
Steroids	<u>Dexamethasone</u>	CYP3A4	CYP17A1, CYP3A5		0	
		Abbreviations: 5-HT, Ser	otonin; NK1, neurokinin 1.			'

PGx Report - Internal Medicine

Type: Drugs Prescribed for the Treatment of Peptic Ulcers and/or Gastro-Esophageal Reflux Disease

Drug Class	Substance	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
Histamine H2-receptor antagonists	<u>Ranitidine</u>	Renal Excretion	CYP1A2, CYP2C19, CYP3A4, CYP3A5			
	<u>Omeprazole</u>	CYP2C19	CYP3A4, CYP2C9, CYP3A5			
	<u>Dexiansoprazole</u>	CYP2C19	CYP3A4, CYP3A5		Ø	
	<u>Esomeprazole</u>	CYP2C19	CYP3A4, CYP3A5		Ø	
Proton-pump inhibitor	<u>Lansoprazole</u>	CYP3A4	CYP2C19, CYP3A5		Ø	
	<u>Rabeprazole</u>	Non Enz	CYP2C19, CYP3A4, CYP3A5		Ø	
	<u>Ilaprazole</u>	CYP3A4	CYP3A5		Ø	
	<u>Pantoprazole</u>	CYP2C19	CYP3A4, CYP2D6, CYP2C9, CYP3A5		Ø	
		Abbreviations: Non Enz, n	on-enzymatic metabolism.			

PGx Report - Internal Medicine

Type: Drugs Prescribed for the Treatment of Functional Gastrointestinal Disorders, Obesity

Drug Class	Substance	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Drugs for functional ga	astrointestinal disorders			
Acting on serotonin	Alosetron	CYP2C9	CYP3A4, CYP1A2			
receptors 5-HT3 antagonists	Cilansetron	CYP3A4	CYP2D6, CYP1A2, CYP2C19, CYP3A5			
Acting on serotonin	<u>Mosapride</u>	CYP3A4	CYP3A5			
receptors 5-HT4 agonists	<u>Prucalopride</u>	Renal Excretion	CYP3A4, CYP3A5			
·		Gastrop	rokinetic			
Serotonin 5-HT ₄ receptor	<u>Cisapride</u>	CYP3A4	CYP3A5		Ø	
agonist	<u>Cinitapride</u>	CYP3A4	CYP2C8, CYP3A5			
Parasympatho mimetic	<u>Itropride</u>	FMO3				
	<u>Metoclopramide</u>	CYP2D6	CYP1A2, CYB5R1, CYB5R2, CYB5R3, CYB5R4		Ø	
Dopamine antagonists	<u>Clebopride</u>	CYP3A4	CYP3A5		Ø	
	<u>Domperidone</u>	CYP3A4	CYP3A5			
·		Antipro	pulsives			
Opioids	<u>Loperamide</u>	CYP3A4	CYP2C8, CYP3A5		Ø	
Opiolas	<u>Morphine</u>	UGT2B7	ABCB1, UGT1A1, COMT		Ø	
		Centrally acting a	anti-obesity drugs			
Stimulant/ Amphetamine/	<u>Sibutramine</u>	CYP3A4	СҮРЗА5			
Appetite suppressant agent	<u>Phentermine</u>	Renal Excretion	CYP3A4, CYP3A5		Ø	
Anorectic	<u>Lorcaserin</u>	CYP2D6	CYP3A4, CYP3A5			

PGx Report - Infectology

Type: Antibiotics

Drug Class	Substance	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antibacterials: protein	synthesis inhibitors 50S			
Amphenicols	Chloramphenicol	CYP2C9	UGT2B7			
Lincosamides	Clindamycin	CYP3A4	CYP3A5			
		Anti	biotic			
	Clarithromycin	CYP3A4	CYP3A5			
Macrolides	<u>Erythromycin</u>	CYP3A4			Ø	
	Telithromycin	CYP3A4	CYP3A5			
		Antibacterials: nu	cleic acid inhibitors			
DHPS inhibitor Short-acting	<u>Sulfadimidine</u>	NAT2	Renal Excretion			
sulfonamides	<u>Sulfapyridine</u>	NAT2	Renal Excretion			
DHPS inhibitor Intermediate- acting sulfonamides	Sulfamethoxazole	Renal Excretion	NAT2, CYP2C9			Ø
Anaerobic DNA inhibitors/	<u>Tinidazole</u>	CYP3A4	CYP3A5			
Nitroimidazole	<u>Ornidazole</u>	CYP3A4	CYP3A5			
DNA-dependent RNA	Rifampicin	CYP3A4	CYP2C8, CYP3A5, CYP2C19, CYP2A6, RE		Ø	
polymerase inhibitors	<u>Rifabutin</u>	CYP3A4	CYP1A2, CYP3A5			
	<u>Dapsone</u>	CYP2E1	NAT2, CYP3A4, CYP2C9, CYP3A5, CYP2D6, G6PD		Ø	
Other drugs against	<u>Bedaquiline</u>	CYP3A4	CYP2C8, CYP2C19, CYP3A5		Ø	
mycobacteria	<u>Isoniazid</u>	NAT2	CYP2E1, Renal Excretion			9
	Pyrazinamide	AOX1, XDH	CYP1A2, CYP3A4, CYP3A5, RE			
		Abbreviations: DHPS, D	ihydropteroate synthase.			

PGx Report - Infectology

Type: Antimalarial, Anthelmintic, Antifungal

Drug Class	Substance	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
		Antim	nalarial			
	<u>Chloroquine</u>	CYP2C8	CYP3A4, CYP3A5, G6PD			
Aminoquinolines	<u>Hydroxychloroquine</u>	CYP2D6	CYP2C8, CYP3A4, CYP3A5			
Ammoquinomies	<u>Amodiaquine</u>	CYP2C8				
	<u>Primaquine</u>	CYP2D6	G6PD			
	<u>Quinine</u>	CYP3A4, CYP2D6	CYP2C19, CYP3A5, G6PD			
Methanolquinolines	<u>Mefloquine</u>	CYP3A4	CYP3A5			
	<u>Artemisinin</u>	CYP3A4	CYP2B6, CYP3A5			
	<u>Artemether</u>	CYP3A4	CYP3A5			
Artemisinin and derivatives	<u>Artesunate</u>	CYP2A6				
	<u>Arteether</u>	CYP3A4	CYP2B6, CYP3A5			
Biguanides	<u>Proguanil</u>	CYP2C19				
011 11 1 1 1	<u>Halofantrine</u>	CYP3A4	CYP3A5			
Other antimalarials	<u>Pentamidine</u>	CYP2C19	CYP1A2, CYP2D6			
		Anthe	lmintic			
Benzimidazoles	<u>Albendazole</u>	CYP3A4	CYP1A2, CYP3A5			
		Antifo	ungals			
Imidazoles	<u>Ketoconazole</u>	CYP3A4	UGT1A1, CYP26A1			
	<u>Itraconazole</u>	CYP3A4				
Triazoles	<u>Voriconazole</u>	CYP2C19	CYP2C9, CYP3A4, CYP3A5			
	<u>Fluconazole</u>	Renal Excretion			Ø	
Allylamines	<u>Terbinafine</u>	CYP2C9	CYP1A2, CYP3A4, CYP2C8, CYP2C19			

PGx Report - Infectology

Type: Antiretroviral, Antiviral

Drug Class	Substance	Primary Mechanism Involved	Other Mechanisms Involved	May Have Decreased Efficacy	Used As Directed	May Have Increased Toxicity
	<u>Lopinavir</u>	СҮРЗА4	SLCO1B1, CYP3A5, ABCC1, ABCC2		Ø	
Protease inhibitor 1st	Ritonavir	СҮРЗА4	CYP2D6, CYP3A5, ABCC1		Ø	
	<u>Saquinavir</u>	CYP3A4	CYP3A5		Ø	
generation	<u>Indinavir</u>	CYP3A4	CYP2D6, CYP3A5, ABCC4		Ø	
	<u>Nelfinavir</u>	CYP2C19	CYP3A4, CYP3A5		Ø	
	<u>Fosamprenavir</u>	CYP3A4	CYP3A5		Ø	
	<u>Atazanavir</u>	CYP3A4	CYP3A5, ABCB1			
Protease inhibitor 2nd generation	<u>Darunavir</u>	CYP3A4	CYP3A5, SLCO3A1			
	<u>Tipranavir</u>	CYP3A4	CYP3A5			
	<u>Delavirdine</u>	CYP3A4	CYP2D6, CYP3A5			
NNRTI 1st generation	<u>Efavirenz</u>	CYP2B6	CYP2A6, ABCB1, SLCO3A1, ABCG2			
	<u>Nevirapine</u>	CYP3A4	CYP2B6, CYP3A5, ABCB1, SLCO3A1			
NNRTI 2nd generation	<u>Etravirine</u>	CYP3A4	CYP2C9, CYP2C19, CYP3A5		2	
	Rilpivirine	CYP3A4	CYP3A5			
Nucleoside reverse	<u>Zidovudine</u>	UGT2B7	Renal Excretion, SLCO3A1, ABCC1, ABCC4		Ø	
anscriptase inhibitor (NRTI)	<u>Abacavir</u>	ADH6	UGT1A1, ADK, HLA-B*5701		Ø	
Neuraminidase	Zanamivir	Renal Excretion				
inhibitors/release phase	<u>Peramivir</u>	Renal Excretion				
CCR5 Co-receptor Antagonist	Maraviroc	CYP3A4	CYP3A5		Ø	
	<u>Boceprevir</u>	CYP3A4	IFNL3, CYP3A5		Ø	
Hepatitis C Virus NS3/4A	<u>Telaprevir</u>	CYP3A4	CYP3A5, IFNL3		Ø	
Protease Inhibitor		CYP3A5		Ø		
	Simeprevir	CYP3A4	CYP2C8, CYP2C19, CYP3A5, IFNL3		Ø	
	<u>Enfuvirtide</u>	CYP2C19	CYP2E1, CYP1A2		6	
Other and in inch	Raltegravir	UGT1A1	SLCO1A2			Ø
Other antivirals	Elvitegravir	CYP3A4	CYP3A5			_
	Dolutegravir	UGT1A1, CYP3A4	CYP3A5			

Patient Information Card

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

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	*2331/*2331	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