UPD Glucuronosyltransferase 1A1 (UGT1A1) and Dihydropyrimidine Dehydrogenase (DPYD) Genotyping

[redacted]

Client: ARUP Example Report Only
500 Chipeta Way
Salt Lake City, UT 84108
[redacted]

ARUP Test Code: 3019841

Collection Date: 07/24/2025
Received in lab: 07/24/2025
Completion Date: 08/05/2025

DPYD Genotyping Specimen Whole Blood

Patient Results

Gene	Genotype	Flag	Phenotype	Flag
DPYD Allele 1	* 1			
DPYD Allele 2	* 1			
DPYD			Normal	

DPYD Interpretation

Activity Score: 2

Interpretation: The following DPYD allele(s) were detected: *1/*1. This result predicts the normal metabolizer phenotype for dihydropyrimidine dehydrogenase (DPD) and normal risk for 5-FU toxicity.

Recommendations: Guidelines for genotype-based dosing are published by the Clinical Pharmacogenetics Implementation Consortium (CPIC) and can be found at: https://cpicpgx.org/ and https://www.pharmgkb.org/.

This result has been reviewed and approved by Sherin Shaaban, M.D., Ph.D.

Interpretive Comments

BACKGROUND INFORMATION: Dihydropyrimidine Dehydrogenase (DPYD)
CHARACTERISTICS: 5-fluorouracil (5-FU) is the most frequently used chemotherapeutic drug for the treatment of many types of cancer, particularly colorectal adenocarcinoma. Grade III-IV drug toxicity attributed to 5-FU occurs in approximately 16 percent of patients, and may include hematologic, gastrointestinal, and dermatologic complications. In some cases, this toxicity can cause death. When 5-FU is metabolized in the body, approximately 80 percent is catabolized by the dihydropyrimidine dehydrogenase (DPD) enzyme. Variants in the DPYD gene can lead to reduced 5-FU catabolism, resulting in the aforementioned toxicity complications.
INHERITANCE: Autosomal codominant.
CAUSE: DPYD gene mutations.
DPYD Variants Tested:
(Variants are numbered according to NM_000110 transcript)
Nonfunctional alleles and increased toxicity risk:
- c.1024G>A (rs183385770)
- c.1774C>T (rs59086055)
- *13 (c.1679T>G, rs55886062)
- *2A (c.1905+1G>A, rs3918290)

[redacted]

Decreased function alleles and increased toxicity risk:
c.557A>G (rs115232898)
c.868A>G (rs146356975)
c.2279C>T (rs112766203)
c.2846A>T (rs67376798)
c.1129-5923C>G (rs75017182)

Functional alleles and normal enzymatic activity:
*1 indicates no variants detected.

METHODOLOGY: Polymerase chain reaction (PCR) and fluorescence monitoring.
ANALYTICAL SENSITIVITY and SPECIFICITY: Greater than 99 percent.
LIMITATIONS: Only the targeted DPYD variants will be detected by this panel. Diagnostic errors can occur due to rare sequence variations. 5-FU drug metabolism, efficacy, and risk for toxicity may be affected by genetic and nongenetic factors that are not evaluated by this test. Genotyping does not replace the need for therapeutic drug monitoring or clinical observation.

Please note the information contained in this report does not contain medication recommendations, and should not be interpreted as recommending any specific medications. Any dosage adjustments or other changes to medications should be evaluated in consultation with a medical provider.

This test was developed and its performance characteristics determined by ARUP Laboratories. It has not been cleared or approved by the U.S. Food and Drug Administration. This test was performed in a CLIA-certified laboratory and is intended for clinical purposes.

Counseling and informed consent are recommended for genetic testing. Consent forms are available online.

[redacted]

UGT1A1 Genotyping Specimen Whole Blood

Patient Results

Gene	Genotype	Flag
UGT1A1 Genotyping Allele 1	(TA)6 or *1	
UGT1A1 Genotyping Allele 2	(TA)6 or *1	

UGT1A1 Genotyping Interpretation

Indications for ordering:
- Determine sensitivity to irinotecan or related compounds.
- Confirm a diagnosis of Gilbert Syndrome.

Homozygous UGT1A1 (TA)6: Two copies of the UGT1A1 *1 (TA)6 were detected predicting a normal metabolizer status This is associated with normal UGT1A1 enzyme levels. This genotype has not been associated with Gilberts syndrome (benign familial hyperbilirubinemia).

This result has been reviewed and approved by [redacted]

Interpretive Comments

BACKGROUND INFORMATION: UDP Glucuronosyltransferase 1A1 (UGT1A1) Genotyping
CHARACTERISTICS: UGT1A1 is responsible for the clearance of drugs (e.g., irinotecan) and endobiotic compounds (e.g., bilirubin). Irinotecan's major active and toxic metabolite (SN-38) is inactivated by the UGT1A1 enzyme and then eliminated via the bile. UGT1A1 gene mutations cause accumulation of SN-38, which may lead to irinotecan-related toxicities (neutropenia, diarrhea).
CAUSE: Variations in TA repeat number in the TATAAA element of the 5'UGT1A1-promoter affects transcription efficiency. The common number of repeats is six [(TA)6, *1 allele], while seven repeats [(TA)7, *28 allele] is associated with reduced transcription activity. Homozygosity for the (TA)7 allele is also associated with Gilbert Syndrome (benign familial hyperbilirubinemia).
ALLELES TESTED: *36 allele, (TA)5; *1 allele, (TA)6; *28 allele, (TA)7 and *37 allele, (TA)8.
CLINICAL SENSITIVITY/SPECIFICITY: Risk of irinotecan toxicity by genotype (Br J Cancer (2004) 91:678-82).
6/6 (*1/*1): diarrhea 17 percent; neutropenia 15 percent
6/7 (*1/*28): diarrhea 33 percent; neutropenia 27 percent
7/7 (*28/*28): diarrhea 70 percent; neutropenia 40 percent

ALLELIC FREQUENCY:
*1(TA)6: Caucasians 0.61, Asians 0.84, African Americans 0.47
*28(TA)7: Caucasians 0.39, Asians 0.16, African Americans 0.43

METHODOLOGY: Polymerase chain reaction followed by size analysis using capillary electrophoresis.

[redacted]

ANALYTICAL SENSITIVITY AND SPECIFICITY: Greater than 99 percent.
LIMITATIONS: Variations in the UGT1A1 gene, other than those targeted, will not be detected. Clinical significance of the rare *36, (TA)5 and *37, (TA)8 alleles in predicting irinotecan toxicities is not well established. Genetic and non-genetic factors other than UGT1A1, may contribute to irinotecan toxicity and efficacy. Diagnostic errors can occur due to rare sequence variations.

This test was developed and its performance characteristics determined by ARUP Laboratories. It has not been cleared or approved by the US Food and Drug Administration. This test was performed in a CLIA certified laboratory and is intended for clinical purposes.

Counseling and informed consent are recommended for genetic testing. Consent forms are available online.

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Test Information

The following tables list the available gene-drug pairs and genotype-based dosing published by the Clinical Pharmacogenetics Implementation Consortium (CPIC) and the FDA table of pharmacogenomic biomarkers in drug labeling.

Published CPIC guidelines

GUIDELINES	DRUGS	GENES
CYP2B6 and efavirenz	efavirenz	CYP2B6
CYP2C19 and Clopidogrel	clopidogrel	CYP2C19
CYP2C19 and Proton Pump Inhibitors	dexlansoprazole	CYP2C19
	esomeprazole	
	lansoprazole	
	omeprazole	
	pantoprazole	
	rabeprazole	
CYP2C19 and Voriconazole	voriconazole	CYP2C19
CYP2C9 and NSAIDs	aceclofenac	CYP2C9
	celecoxib	
	diclofenac	
	flurbiprofen	
	ibuprofen	
	indomethacin	
	lornoxicam	
	lumiracoxib	
	meloxicam	
	metamizole	
	nabumetone	
	naproxen	
	piroxicam	
	tenoxicam	
CYP2C9, VKORC1, CYP4F2 and Warfarin	warfarin	CYP2C9, CYP4F2, VKORC1
CYP2D6 and Atomoxetine	atomoxetine	CYP2D6
CYP2D6 and Ondansetron and Tropisetron	ondansetron	CYP2D6
	tropisetron	
CYP2D6 and Tamoxifen	tamoxifen	CYP2D6
	citalopram	

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CYP2D6, CYP2C19 and Selective Serotonin Reuptake Inhibitors	escitalopram	CYP2C19, CYP2D6
	fluvoxamine	
	paroxetine	
	sertraline	
CYP2D6, CYP2C19 and Tricyclic Antidepressants	amitriptyline	CYP2C19, CYP2D6
	clomipramine	
	desipramine	
	doxepin	
	imipramine	
	nortriptyline	
	trimipramine	
CYP2D6, OPRM1, COMT, and Opioids	alfentanil	COMT, CYP2D6, OPRM1
	buprenorphine	
	codeine	
	fentanyl	
	hydrocodone	
	hydromorphone	
	levomethadone	
	methadone	
	morphine	
	naltrexone	
	oxycodone	
	remifentanil	
	sufentanil	
	tramadol	
CYP3A5 and Tacrolimus	tacrolimus	CYP3A5
DPYD and Fluoropyrimidines	capecitabine	DPYD
	fluorouracil	
	tegafur	
SLCO1B1, ABCG2, CYP2C9, and Statins	atorvastatin	ABCG2, CYP2C9, SLCO1B1
	fluvastatin	
	lovastatin	
	pitavastatin	
	pravastatin	
	rosuvastatin	

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	simvastatin	
TPMT, NUDT15 and Thiopurines	azathioprine	NUDT15, TPMT
	mercaptopurine	
	thioguanine	
UGT1A1 and Atazanavir	atazanavir	UGT1A1

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FDA Table of Pharmacogenomic Biomarkers in Drug Labeling

Drug	Therapeutic Area*	Biomarker†	Labeling Sections
Abrocitinib	Dermatology	CYP2C19	Dosage and Administration, Use in Specific Populations, Clinical Pharmacology
Amitriptyline	Psychiatry	CYP2D6	Precautions
Amoxapine	Psychiatry	CYP2D6	Precautions
Amphetamine	Psychiatry	CYP2D6	Clinical Pharmacology
Arformoterol	Pulmonary	UGT1A1	Clinical Pharmacology
Arformoterol	Pulmonary	CYP2D6	Clinical Pharmacology
Aripiprazole	Psychiatry	CYP2D6	Dosage and Administration, Use in Specific Populations, Clinical Pharmacology
Aripiprazole Lauroxil	Psychiatry	CYP2D6	Dosage and Administration, Use in Specific Populations, Clinical Pharmacology
Atomoxetine	Psychiatry	CYP2D6	Dosage and Administration, Warnings and Precautions, Adverse Reactions, Drug Interactions, Use in Specific Populations, Clinical Pharmacology
Avatrombopag	Hematology	CYP2C9	Clinical Pharmacology
Azathioprine	Rheumatology	TPMT	Dosage and Administration, Warnings, Precautions, Drug Interactions, Adverse Reactions, Clinical Pharmacology
Azathioprine	Rheumatology	NUDT15	Dosage and Administration, Warnings, Precautions, Adverse Reactions, Clinical Pharmacology
Belinostat	Oncology	UGT1A1	Dosage and Administration, Clinical Pharmacology
Belzutifan	Oncology	CYP2C19	Warnings and Precautions, Drug Interactions, Use in Specific Populations, Clinical Pharmacology
Binimetinib	Oncology	UGT1A1	Clinical Pharmacology
Brexpiprazole	Psychiatry	CYP2D6	Dosage and Administration, Use in Specific Populations, Clinical Pharmacology
Brivaracetam	Neurology	CYP2C19	Clinical Pharmacology
Bupropion	Psychiatry	CYP2D6	Clinical Pharmacology
Cabotegravir and Rilpivirine	Infectious Diseases	UGT1A1	Clinical Pharmacology
Capecitabine	Oncology	DPYD	Warnings and Precautions, Clinical Pharmacology, Patient Counseling Information
Cariprazine	Psychiatry	CYP2D6	Clinical Pharmacology
Carisoprodol	Rheumatology	CYP2C19	Use in Specific Populations, Clinical Pharmacology
Carvedilol	Cardiology	CYP2D6	Drug Interactions, Clinical Pharmacology

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Celecoxib	Rheumatology	CYP2C9	Dosage and Administration, Use in Specific Populations, Clinical Pharmacology
Cevimeline	Dental	CYP2D6	Precautions
Cisplatin	Oncology	TPMT	Adverse Reactions
Citalopram	Psychiatry	CYP2C19	Dosage and Administration, Warnings, Clinical Pharmacology
Citalopram	Psychiatry	CYP2D6	Clinical Pharmacology
Clobazam	Neurology	CYP2C19	Dosage and Administration, Use in Specific Populations, Clinical Pharmacology
Clomipramine	Psychiatry	CYP2D6	Precautions
Clopidogrel	Cardiology	CYP2C19	Boxed Warning, Warnings and Precautions, Clinical Pharmacology
Clozapine	Psychiatry	CYP2D6	Dosage and Administration, Use in Specific Populations, Clinical Pharmacology
Codeine	Anesthesiology	CYP2D6	Boxed Warning, Warnings and Precautions, Use in Specific Populations, Patient Counseling Information
Darifenacin	Urology	CYP2D6	Clinical Pharmacology
Desipramine	Psychiatry	CYP2D6	Precautions
Desvenlafaxine	Psychiatry	CYP2D6	Clinical Pharmacology
Deutetrabenazine	Neurology	CYP2D6	Dosage and Administration, Warnings and Precautions, Use in Specific Populations, Clinical Pharmacology
Dexlansoprazole	Gastroenterology	CYP2C19	Drug Interactions, Clinical Pharmacology
Dextromethorphan and Quinidine	Neurology	CYP2D6	Warnings and Precautions, Clinical Pharmacology
Diazepam	Neurology	CYP2C19	Clinical Pharmacology
Dolutegravir	Infectious Diseases	UGT1A1	Clinical Pharmacology
Donepezil	Neurology	CYP2D6	Clinical Pharmacology
Doxepin	Psychiatry	CYP2D6	Clinical Pharmacology
Doxepin	Psychiatry	CYP2C19	Clinical Pharmacology
Dronabinol	Gastroenterology	CYP2C9	Use in Specific Populations, Clinical Pharmacology
Drospirenone and Ethinyl Estradiol	Gynecology	CYP2C19	Clinical Pharmacology
Duloxetine	Psychiatry	CYP2D6	Drug Interactions
Efavirenz	Infectious Diseases	CYP2B6	Clinical Pharmacology
Elagolix	Gynecology	SLCO1B1	Clinical Pharmacology
Eliglustat	Inborn Errors of Metabolism	CYP2D6	Indications and Usage, Dosage and Administration, Contraindications, Warnings and Precautions, Drug Interactions, Use in Specific Populations, Clinical Pharmacology, Clinical Studies

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Erdafitinib	Oncology	CYP2C9	Use in Specific Populations, Clinical Pharmacology
Escitalopram	Psychiatry	CYP2D6	Drug Interactions
Escitalopram	Psychiatry	CYP2C19	Adverse Reactions
Esomeprazole	Gastroenterology	CYP2C19	Drug Interactions, Clinical Pharmacology
Fesoterodine	Urology	CYP2D6	Drug Interactions, Clinical Pharmacology
Fosphenytoin	Neurology	CYP2C9	Warnings and Precautions, Use in Specific Populations, Clinical Pharmacology
Flibanserin	Gynecology	CYP2C9	Clinical Pharmacology
Flibanserin	Gynecology	CYP2C19	Adverse Reactions, Use in Specific Populations, Clinical Pharmacology
Flibanserin	Gynecology	CYP2D6	Clinical Pharmacology
Fluorouracil	Dermatology	DPYD	Contraindications, Warnings
Fluorouracil	Oncology	DPYD	Warnings and Precautions, Patient Counseling Information
Fluoxetine	Psychiatry	CYP2D6	Precautions, Clinical Pharmacology
Flurbiprofen	Rheumatology	CYP2C9	Clinical Pharmacology
Fluvoxamine	Psychiatry	CYP2D6	Drug Interactions
Formoterol	Pulmonary	CYP2D6	Clinical Pharmacology
Formoterol	Pulmonary	CYP2C19	Clinical Pharmacology
Galantamine	Neurology	CYP2D6	Clinical Pharmacology
Gefitinib	Oncology	CYP2D6	Clinical Pharmacology
Iloperidone	Psychiatry	CYP2D6	Dosage and Administration, Warnings and Precautions, Drug Interactions, Clinical Pharmacology
Imipramine	Psychiatry	CYP2D6	Precautions
Indacaterol	Pulmonary	UGT1A1	Clinical Pharmacology
Irinotecan	Oncology	UGT1A1	Dosage and Administration, Warnings and Precautions, Clinical Pharmacology
Lacosamide	Neurology	CYP2C19	Clinical Pharmacology
Lansoprazole	Gastroenterology	CYP2C19	Drug Interactions, Clinical Pharmacology
Lesinurad	Rheumatology	CYP2C9	Drug Interactions, Clinical Pharmacology
Lofexidine	Anesthesiology	CYP2D6	Use in Specific Populations
Mavacamten	Cardiology	CYP2C19	Dosage and Administration, Clinical Pharmacology
Meclizine	Neurology	CYP2D6	Warnings and Precautions
Meloxicam	Anesthesiology	CYP2C9	Use in Specific Populations, Clinical Pharmacology
Mercaptopurine	Oncology	TPMT	Dosage and Administration, Warnings and Precautions, Adverse Reactions, Clinical Pharmacology
Mercaptopurine	Oncology	NUDT15	Dosage and Administration, Warnings and Precautions, Clinical Pharmacology

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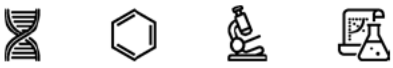
Metoclopramide	Gastroenterology	CYP2D6	Dosage and Administration, Use in Specific Populations, Clinical Pharmacology
Metoprolol	Cardiology	CYP2D6	Clinical Pharmacology
Mirabegron	Urology	CYP2D6	Clinical Pharmacology
Modafinil	Psychiatry	CYP2D6	Clinical Pharmacology
Nateglinide	Endocrinology	CYP2C9	Drug Interactions
Nebivolol	Cardiology	CYP2D6	Dosage and Administration, Clinical Pharmacology
Nefazodone	Psychiatry	CYP2D6	Precautions
Nilotinib	Oncology	UGT1A1	Clinical Pharmacology
Nortriptyline	Psychiatry	CYP2D6	Precautions
Oliceridine	Anesthesiology	CYP2D6	Warnings and Precautions, Drug Interactions, Use in Specific Populations, Clinical Pharmacology
Omeprazole	Gastroenterology	CYP2C19	Drug Interactions, Clinical Pharmacology
Ondansetron	Gastroenterology	CYP2D6	Clinical Pharmacology
Ospemifene	Gynecology	CYP2C9	Clinical Pharmacology
Ospemifene	Gynecology	CYP2B6	Clinical Pharmacology
Paliperidone	Psychiatry	CYP2D6	Clinical Pharmacology
Palonosetron	Gastroenterology	CYP2D6	Clinical Pharmacology
Pantoprazole	Gastroenterology	CYP2C19	Clinical Pharmacology
Paroxetine	Psychiatry	CYP2D6	Drug Interactions, Clinical Pharmacology
Pazopanib	Oncology	UGT1A1	Clinical Pharmacology
Perphenazine	Psychiatry	CYP2D6	Precautions, Clinical Pharmacology
Phenytoin	Neurology	CYP2C9	Warnings and Precautions, Use in Specific Populations, Clinical Pharmacology
Phenytoin	Neurology	CYP2C19	Clinical Pharmacology
Pimozide	Psychiatry	CYP2D6	Dosage and Administration, Precautions
Piroxicam	Rheumatology	CYP2C9	Clinical Pharmacology
Pitolisant	Psychiatry	CYP2D6	Dosage and Administration, Use in Specific Populations, Clinical Pharmacology
Prasugrel	Cardiology	CYP2C19	Use in Specific Populations, Clinical Pharmacology, Clinical Studies
Prasugrel	Cardiology	CYP2C9	Use in Specific Populations, Clinical Pharmacology, Clinical Studies
Prasugrel	Cardiology	CYP3A5	Use in Specific Populations, Clinical Pharmacology, Clinical Studies
Prasugrel	Cardiology	CYP2B6	Use in Specific Populations, Clinical Pharmacology, Clinical Studies



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Propafenone	Cardiology	CYP2D6	Dosage and Administration, Warnings and Precautions, Drug Interactions, Clinical Pharmacology
Propranolol	Cardiology	CYP2D6	Clinical Pharmacology
Protriptyline	Psychiatry	CYP2D6	Precautions
Quinidine	Cardiology	CYP2D6	Precautions
Quinine Sulfate	Infectious Diseases	CYP2D6	Drug Interactions
Rabeprazole	Gastroenterology	CYP2C19	Drug Interactions, Clinical Pharmacology
Raltegravir	Infectious Diseases	UGT1A1	Clinical Pharmacology
Rimegepant	Neurology	CYP2C9	Clinical Pharmacology
Risperidone	Psychiatry	CYP2D6	Clinical Pharmacology
Rosuvastatin	Endocrinology	SLCO1B1	Clinical Pharmacology
Rucaparib (2)	Oncology	CYP2D6	Clinical Pharmacology
Sacituzumab Govitecan-hziy	Oncology	UGT1A1	Warnings and Precautions, Clinical Pharmacology
Siponimod	Neurology	CYP2C9	Dosage and Administration, Contraindications, Drug Interactions, Use in Specific Populations, Clinical Pharmacology
Tamoxifen	Oncology	CYP2D6	Clinical Pharmacology
Tamsulosin	Urology	CYP2D6	Warnings and Precautions, Adverse Interactions, Clinical Pharmacology
Tetrabenazine	Neurology	CYP2D6	Dosage and Administration, Warnings and Precautions, Use in Specific Populations, Clinical Pharmacology
Thioguanine	Oncology	TPMT	Dosage and Administration, Warnings, Precautions, Clinical Pharmacology
Thioguanine	Oncology	NUDT15	Dosage and Administration, Warnings, Precautions, Clinical Pharmacology
Thioridazine	Psychiatry	CYP2D6	Contraindications, Warnings, Precautions
Ticagrelor	Cardiology	CYP2C19	Clinical Pharmacology
Tolterodine	Urology	CYP2D6	Warnings and Precautions, Drug Interactions, Clinical Pharmacology
Tramadol	Anesthesiology	CYP2D6	Boxed Warning, Warnings and Precautions, Use in Specific Populations, Clinical Pharmacology, Patient Counseling Information
Trimipramine	Psychiatry	CYP2D6	Precautions
Umeclidinium	Pulmonary	CYP2D6	Clinical Pharmacology
Upadacitinib	Rheumatology	CYP2D6	Clinical Pharmacology

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Valbenazine	Neurology	CYP2D6	Dosage and Administration, Warnings and Precautions, Use in Specific Populations, Clinical Pharmacology
Venlafaxine	Psychiatry	CYP2D6	Drug Interactions, Use in Specific Populations, Clinical Pharmacology
Viloxazine	Psychiatry	CYP2D6	Clinical Pharmacology
Viloxazine	Psychiatry	SLCO1B1	Clinical Pharmacology
Voriconazole	Infectious Diseases	CYP2C19	Clinical Pharmacology
Vortioxetine	Psychiatry	CYP2D6	Dosage and Administration, Clinical Pharmacology
Warfarin	Hematology	CYP2C9	Dosage and Administration, Drug Interactions, Clinical Pharmacology
Warfarin	Hematology	VKORC1	Dosage and Administration, Clinical Pharmacology

* Therapeutic areas do not necessarily reflect the CDER review division.
† Representative biomarkers are listed based on standard nomenclature as per the Human Genome Organization (HUGO) symbol and/or simplified descriptors using other common conventions. Listed biomarkers do not necessarily reflect the terminology used in labeling. The term "Nonspecific" is provided when labeling does not explicitly identify the specific biomarker(s) or when the biomarker is represented by a molecular phenotype or gene signature, and in some cases the biomarker was inferred based on the labeling language.