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Pharmacogenomics: Statements, Standards, and Competencies

Clinical implementation PGx Dissemination Materials competencies



CPIC Informatics Working Group

- Growing interest in informatics aspects of CPIC guidelines and clinical implementation of pharmacogenetics
- Goal: To support the adoption of the CPIC guidelines by identifying, and resolving where possible, potential technical barriers to the implementation of the guidelines within a clinical electronic environment.
- Working group leaders
 - Bob Freimuth (Mayo Clinic)
 - James Hoffman (St. Jude)
 - Michelle Carrillo (Stanford)



CPIC Informatics Working Group: Initial Focus

- <u>Create comprehensive translation tables</u> from genotype to phenotype to clinical recommendation for CPIC guidelines
 - Define structure and process to efficiently develop and maintain in the most useful format(s)
 - Publish as part of CPIC guidelines





Official journal of the American College of Medical Genetics and Genomics ORIGINAL RESEARCH ARTICLE in Medicine

Open

Standardizing terms for clinical pharmacogenetic test results: consensus terms from the Clinical Pharmacogenetics Implementation Consortium (CPIC)

Kelly E. Caudle, PharmD, PhD¹, Henry M. Dunnenberger, PharmD², Robert R. Freimuth, PhD³, Josh F. Peterson, MD^{4,5}, Jonathan D. Burlison, PhD¹, Michelle Whirl-Carrillo, PhD⁶, Stuart A. Scott, PhD⁷, Heidi L. Rehm, PhD⁸, Marc S. Williams, MD⁹, Teri E. Klein, PhD⁶, Mary V. Relling, PharmD¹, James M. Hoffman, PharmD, MS¹

Introduction: Reporting and sharing pharmacogenetic test results across clinical laboratories and electronic health records is a crucial step toward the implementation of clinical pharmacogenetics, but allele function and phenotype terms are not standardized. Our goal was to develop terms that can be broadly applied to characterize pharmacogenetic allele function and inferred phenotypes.

Materials and methods: Terms currently used by genetic testing laboratories and in the literature were identified. The Clinical Pharmacogenetics Implementation Consortium (CPIC) used the Delphi method to obtain a consensus and agree on uniform terms among pharmacogenetic experts.

Results: Experts with diverse involvement in at least one area of pharmacogenetics (clinicians, researchers, genetic testing laborato-

rians, pharmacogenetics implementers, and clinical informaticians; n = 58) participated. After completion of five surveys, a consensus (>70%) was reached with 90% of experts agreeing to the final sets of pharmacogenetic terms.

Discussion: The proposed standardized pharmacogenetic terms will improve the understanding and interpretation of pharmacogenetic tests and reduce confusion by maintaining consistent nomenclature. These standard terms can also facilitate pharmacogenetic data sharing across diverse electronic health care record systems with clinical decision support.

Genet Med advance online publication 21 July 2016

Key Words: CPIC; nomenclature; pharmacogenetics; pharmacogenomics; terminology SNOMED terms were not sufficiently granular to build precise CDS

CPIC has worked to submit standard terms per consensus process for addition to SNOMED and they are now available.

New SNOMED terms also enable:

- Interoperability (e.g. phenotype will travel with patient)
- Sharing CDS

Proof of concept –now scale up!



Developing knowledge resources to support precision medicine: principles from the Clinical Pharmacogenetics Implementation Consortium (CPIC) RECEIVED 1 September 2015 REVISED 7 December 2015 ACCEPTED 13 January 2016



James M. Hoffman,¹ Henry M Dunnenberger,² J Kevin Hicks,³ Kelly E Caudle,¹ Michelle Whirl Carrillo,⁴ Robert R Freimuth,⁵ Marc S Williams,⁶ Teri E Klein,⁴ and Josh F Peterson⁷

ABSTRACT

To move beyond a select few genes/drugs, the successful adoption of pharmacogenomics into routine clinical care requires a curated and machine-readable database of pharmacogenomic knowledge suitable for use in an electronic health record (EHR) with clinical decision support (CDS). Recognizing that EHR vendors do not yet provide a standard set of CDS functions for pharmacogenetics, the Clinical Pharmacogenetics Implementation Consortium (CPIC) Informatics Working Group is developing and systematically incorporating a set of EHR-agnostic implementation resources into all CPIC guidelines. These resources illustrate how to integrate pharmacogenomic test results in clinical information systems with CDS to facilitate the use of patient genomic data at the point of care. Based on our collective experience creating existing CPIC resources and implementing pharmacogenomics at our practice sites, we outline principles to define the key features of future knowledge bases and discuss the importance of these knowledge resources for pharmacogenomics and ultimately precision medicine.

Keywords: pharmacogenetics, knowledge bases, electronic health records, clinical decision support systems, precision medicine



Hicks JK et al. Am J Health-Syst Pharm 2016 Hoffman JM et al. J Am Med Inform Assoc. 2016

CPIC's Five Principles for Knowledge Resources



Pharmacogenomic knowledge resources must

- 1. support traceability between interrogated variants primary results, and clinical interpretations.
- 2. rate level of evidence for each variant as well as for the overall recommendation
- 3. use standards to facilitate information exchange and enable interoperability among disparate systems

CPIC's Five Principles for Knowledge Resources Pharmacogenomic knowledge resources must:

4. - support long-term reinterpretation of results

5. - be positioned to be integrated with other knowledge at the point of care

Hoffman et al. Developing knowledge resources to support precision medicine: principles from the Clinical Pharmacogenetics Implementation Consortium (CPIC) JAMIA 2016



	7				•				1
	Guideline	Sent to Rose	Rose review	allele defintion	functionality	diplotype to phenotype/consult	pre/post alerts	Sent to Rose	Rose review
2	CFTR/ivacaftor			?	n/a	mapping from Table 1	n/a	n/a	
3	CYP2C19/voriconazole	x	x	x	x	x	x	x	x
1	CYP2C19 and CYP2D6/TCA	x		x	х	x	x	x	
5	CYP2C19/SSRI	x	x	x	х	x	x	x	x
5	CYP2C19/clopidogrel	х	x	x	x	x	Does not have	Does not have	Does not have
7	CYP2B6/efavirenz	х	x	no change	x	x	x	x	
3	CYP2D6 and atomoxetine	x	x	finalizing with codier	finalizing with codien	finalizing with codiene authors	x	x	
9	CYP2D6 and codeine	x	х	finalizing with codier	finalizing with codien	finalizing with codiene authors	Does not have	Does not have	Does not have
0	CYP2D6 and ondansetron/tropisetron	x	х	finalizing with codier	finalizing with codien	finalizing with codiene authors	х	x	
1	CYP2D6 and tamoxifen	x	х	finalizing with codier	finalizing with codien	finalizing with codiene authors	х	х	
2	CYP2C9, HLA-b/Phenytoin	x	x	in progress	in progress	mapping from Table 1 for HLA	in progress		
3	warfarin guideline	??	??	??			Does not have		
4	CYP3A5/tacrolimus	x	x	x	x	x	x	x	
5	DPYD/fluoropyrimidines	x	x	x	x	x	x	x	
6	G6PD/rasburicase	x	x		need	need	need???		
7	HLA/carbamazepine/oxcarbazepine	x	x	x		mapping from Table 1	x	x	reviewing agai

Over 100 tables updated; 23 authors groups to review and approve

Phenotype	Implication	Therapeutic recommendation ^{a,b}	Classification of recommendation for amitriptyline and nortripyline ^c	Classification of recommendation for other TCAs ^{c,d}
CYP2D6 ultrarapid metabolizer	Increased metabolism of TCAs to less active compounds compared to normal metabolizers Lower plasma concentrations of active drug will increase probability of phar- macotherapy failure	Avoid tricyclic use due to potential lack of efficacy. Consider alternative drug not metabolized by CYP2D6. If a TCA is warranted, consider titrating to a higher target dose (compared to normal metabolizers). ^e Utilize thera- peutic drug monitoring to guide dose adjustments.	Strong	Optional
CYP2D6 normal metabolizer	Normal metabolism of TCAs	Initiate therapy with recommended starting dose. ^f	Strong	Strong
CYP2D6 intermediate metabolizer	Reduced metabolism of TCAs to less active compounds compared to normal metabolizers Higher plasma concentrations of active drug will increase the probability of side effects	Consider a 25% reduction of recom- mended starting dose. ^f Utilize thera- peutic drug monitoring to guide dose adjustments. ^e	Moderate	Optional
CYP2D6 poor metabolizer	Greatly reduced metabolism of TCAs to less active compounds compared to normal metabolizers Higher plasma concentrations of active drug will increase the probability of side effects	Avoid tricyclic use due to potential for side effects. Consider alternative drug not metabolized by CYP2D6. If a TCA is warranted, consider a 50% reduction of recommended starting dose. ¹ Utilize therapeutic drug monitor- ing to guide dose adjustments. ^e	Strong	Optional

Table 2 Dosing recommendations for tricyclic antidepressants based on cyp2d6 phenotype

......

CYP2C19 Diplotype	Activity Score	CYP2C19 phenotype	EHR Priority Result Notation
*24/*26	n/a	Likely CYP2C19 Poor Metabolizer	Abnormal/Priority/High Risk
*25/*35	n/a	Likely CYP2C19 Poor Metabolizer	Abnormal/Priority/High Risk
*25/*36	n/a	Likely CYP2C19 Poor Metabolizer	Abnormal/Priority/High Risk
*25/*37	n/a	Likely CYP2C19 Poor Metabolizer	Abnormal/Priority/High Risk
*26/*35	n/a	Likely CYP2C19 Poor Metabolizer	Abnormal/Priority/High Risk
*26/*36	n/a	Likely CYP2C19 Poor Metabolizer	Abnormal/Priority/High Risk
*26/*37	n/a	Likely CYP2C19 Poor Metabolizer	Abnormal/Priority/High Risk
*1/*17	n/a	CYP2C19 Rapid Metabolizer	Abnormal/Priority/High Risk
*11/*17	n/a	CYP2C19 Rapid Metabolizer	Abnormal/Priority/High Risk
*13/*17	n/a	CYP2C19 Rapid Metabolizer	Abnormal/Priority/High Risk
*15/*17	n/a	CYP2C19 Rapid Metabolizer	Abnormal/Priority/High Risk
*17/*18	n/a	CYP2C19 Rapid Metabolizer	Abnormal/Priority/High Risk
*17/*28	n/a	CYP2C19 Rapid Metabolizer	Abnormal/Priority/High Risk
*17/*17	n/a	CYP2C19 Ultrarapid Metabolizer	Abnormal/Priority/High Risk
*1/*12	n/a	Indeterminate	none
*1/*14	n/a	Indeterminate	none
*1/*23	n/a	Indeterminate	none

CYP2D6 Phenotype	Activity Score	Implications for Phenotypic Measures	Therapeutic Recommendation	Classification of Reco	Comments
CYP2D6 ultrarapid metabolizer	>2.25	Increased metabolism of tricyclic antidepre	Avoid tricyclic use due to potential lack of efficacy. Con	Strong	n/a
CYP2D6 normal metabolizer	2.25	Normal metabolism of tricyclic antidepress	Initiate therapy with recommended starting dose	Strong	Patients may receive ar
CYP2D6 normal metabolizer	2	Normal metabolism of tricyclic antidepress	Initiate therapy with recommended starting dose	Strong	Patients may receive ar
CYP2D6 normal metabolizer	1.5	Normal metabolism of tricyclic antidepress	Initiate therapy with recommended starting dose	Strong	Patients may receive an
CYP2D6 normal metabolizer	1.25	Normal metabolism of tricyclic antidepress	Initiate therapy with recommended starting dose	Strong	Patients may receive ar
CYP2D6 intermediate metabolizer	1	Reduced metabolism of tricyclic antidepres	Consider a 25% reduction of recommended starting dos	Optional	Patients may receive ar
CYP2D6 intermediate metabolizer	0.75	Reduced metabolism of tricyclic antidepres	Consider a 25% reduction of recommended starting dos	Optional	Patients may receive ar
CYP2D6 intermediate metabolizer	0.5	Reduced metabolism of tricyclic antidepres	Consider a 25% reduction of recommended starting dos	Moderate	Patients may receive ar
CYP2D6 intermediate metabolizer	0.25	Reduced metabolism of tricyclic antidepres	Consider a 25% reduction of recommended starting dos	Moderate	Patients may receive ar
CYP2D6 poor metabolizer	0	Greatly reduced metabolism of tricyclic ant	Avoid tricyclic use due to potential for side effects. Con	Strong	Patients may receive ar
Indeterminate	n/a	n/a	No recommendation	No recommendation	n/a

CYP2C19 Phenotype	CYP2C19	CYP2D6 Phenotype	CYP2D6	CYP2C19	CYP2D	Therapeutic	Classification of	Comments
	Activity		Activity	Implication	n 6	Recommendation	Recommendation	
	Score		Score	s for	Impli 💌	· · · · · · · · · · · · · · · · · · ·	· ·	
CYP2C19 intermediate	n/a	CYP2D6 intermediate metabolizer	1	Reduced m	e Reduce	Consider a 25% redu	Optional	Patients may receive an initial I
CYP2C19 intermediate	n/a	CYP2D6 intermediate metabolizer	0.75	Reduced m	e Reduce	Consider a 25% redu	Optional	Patients may receive an initial I
CYP2C19 intermediate	n/a	CYP2D6 intermediate metabolizer	0.5	Reduced m	e Reduce	Consider a 25% redu	Optional	Patients may receive an initial I
CYP2C19 intermediate	n/a	CYP2D6 intermediate metabolizer	0.25	Reduced m	e Reduce	Consider a 25% redu	Optional	Patients may receive an initial I
CYP2C19 normal metabolizer	n/a	CYP2D6 intermediate metabolizer	1	Normal met	ta Reduce	Consider a 25% redu	u Moderate	Patients may receive an initial I
CYP2C19 normal metabolizer	n/a	CYP2D6 intermediate metabolizer	0.75	Normal met	ta Reduce	Consider a 25% redu	u Moderate	Patients may receive an initial I
CYP2C19 normal metabolizer	n/a	CYP2D6 intermediate metabolizer	0.5	Normal met	ta Reduce	Consider a 25% redu	u Moderate	Patients may receive an initial I
CYP2C19 normal metabolizer	n/a	CYP2D6 intermediate metabolizer	0.25	Normal met	ta Reduce	Consider a 25% redu	u Moderate	Patients may receive an initial I
CYP2C19 poor metabolizer	n/a	CYP2D6 intermediate metabolizer	1	Greatly red	u Reduce	Avoid amitriptyline	Optional	Dosing recommendations only a
CYP2C19 poor metabolizer	n/a	CYP2D6 intermediate metabolizer	0.75	Greatly red	u Reduce	Avoid amitriptyline	Optional	Dosing recommendations only a
CYP2C19 poor metabolizer	n/a	CYP2D6 intermediate metabolizer	0.5	Greatly red	u Reduce	Avoid amitriptyline	Optional	Dosing recommendations only a
CYP2C19 poor metabolizer	n/a	CYP2D6 intermediate metabolizer	0.25	Greatly red	u Reduce	Avoid amitriptyline	Optional	Dosing recommendations only a
CYP2C19 rapid metabolizer	n/a	CYP2D6 intermediate metabolizer	1	Increased n	n Reduce	Consider alternative	Optional	Dosing recommendations only a
CYP2C19 rapid metabolizer	n/a	CYP2D6 intermediate metabolizer	0.75	Increased n	n Reduce	Consider alternative	Optional	Dosing recommendations only a
CYP2C19 ranid metabolizer	n/a	CYP2D6 intermediate metabolizer	0.5	Increased n	n Reduce	Consider alternative	Ontional	Dosing recommendations only a

The referenc	ed substrates and cited artic	les are examples and may not represent a	all information that may be avail	able for an allele.	
GENE: CYP2C19	5/7/2019			Dru	ıg substrate
Allele	Allele Functional Status	References	PMID	in vitro	in vivo
*1	Normal function	Romkes 1991	2009263		
		Richardson 1995	7487078	S-mephenytoin, tolbutamide	
		Blaisdell 2002	12464799	S-mephenytoin	
		Hanioka 2007	17455109	S-mephenytoin	
		Hanioka 2008	18312490	omeprazole	
		Wang 2011	21325430	S-mephenytoin, omeprazole	
		Takahashi 2015	25001882	clopidogrel, S-mephenytoin	
		Drögemöller 2010	20712527		
		Dodgen 2015	26244421		omeprazole
*2	No function	de Morais 1994	8195181	S-mephenytoin	
		Ibeanu 1998	9732415		S-mephenytoin
		Lee 2009	19661214		S-mephenytoin, omeprazole
		Xiao 1997	9103550		S-mephenytoin
*3	No function	de Morais 1994	7969038		S-mephenytoin
		Xiao 1997	9103550		S-mephenytoin
*4	No function	Ferguson 1998	9435198		S-mephenytoin
		Scott 2012	21358751		clopidogrel

GENE: CYP2C19							
Allele/cDNA/rsID	Activity Score (Optional)	Allele Functional Status (Optional)	Allele Clinical Functional Status (Required)	Allele Clinical Function Substrate Specificity (Optional)	PMID (Optional)	Strength of Evidence (Optional)	Findings (Optional)
*1			Normal function		2009263, 7487078, 12464799, 17455109, 18312490, 21325430, 25001882, 20712527, 26244421		2009263; 7484048: S-mephenytoin (in vitro), tolbutamide (in vitro); 124647 vitro); 17455109: S-mephenytoin (in vitro); 18312490: omeprazole (in vitro mephenytoin (in vitro), omeprazole (in vitro); 25001882: clopidogrel (in vitro vitro); 20712527; 26244421: omeprazole (in vivo)
*2			No function		8195181, 9732415, 19661214, 9103550		8195181: S-mephenytoin (in vitro); 9732415: S-mephenytoin (in vivo); 196 (in vivo), omeprazole (in vivo); 9103550: S-mephenytoin (in vivo)
*3			No function		7969038, 9103550		7969038: S-mephenytoin (in vivo); 9103550: S-mephenytoin (in vivo)
*4			No function		9435198, 21358751		9435198: S-mephenytoin (in vivo); 21358751: clopidogrel (in vivo)
*5			No function		9103550, 10022751, 21325430, 25001882		9103550: S-mephenytoin (in vivo); 10022751: S-mephenytoin (in vitro), toll 21325430: S-mephenytoin (in vitro), omeprazole (in vitro); 25001882: clopi mephenytoin (in vitro)
***					9732415, 21325430,		9732415: S-mephenytoin (in vivo); 21325430: S-mephenytoin (in vitro), om

Phenotype Modifiers

Likely phenotype ^a	Genotypes	Examples of diplotypes		
Assignment of likely TPMT phenotypes ba	ised on genotypes			
Normal metabolizer	An individual carrying two normal function alleles	*1/*1		
Intermediate metabolizer	An individual carrying one normal function allele PLUS one no function allele	*1/*2, *1/*3A, *1/*3B, *1/*3C, *1/*4		
Possible intermediate metabolizer	An individual carrying one uncertain/unknown function allele PLUS one no function allele	*2/*8, *3A/*7		
Poor metabolizer	An individual carrying two no function alleles	*3A/*3A, *2/*3A, *3A/*3C, *3C/*4, *2/*3C, *3A/*4		
Indeterminate	An individual carrying two uncertain/unknown function alleles OR one normal function allele plus one uncertain allele function allele	*6/*8 *1/*8		

Table 1 Assignment of likely TPMT and NUDT15 phenotypes based on genotypes

GENE: SLCO1B1	2/8/2017			D)rug substrate
Allele	Allele Functional Status	References	PMID	in vitro	in vivo
*1a	Normal function	Mwinyi 2004	16568260		pravastatin
*1b	Normal function	Mwinyi 2004	16568260		pravastatin
		Kameyama 2005	15970799	pravastatin, atorvastatir	, cerivastatin
		Lee 2005	16198652		rosuvastatin
		Katz 2006	16513443	atrasentan	atrasentan
		Tirona 2001	11477075	estrone sulfate, estradio	ol 17beta-d-glucuronide
	Possible decreased				
*2	function	Tirona 2001	11477075	estrone sulfate, estradio	ol 17beta-d-glucuronide
		Tirona 2003	12490595	rifampin	
	Possible decreased				
*3	function	Tirona 2001	11477075	estrone sulfate, estradio	ol 17beta-d-glucuronide

Any combination with a "possible" function translated into a "possible" phenotype

CYP2C19 phenotype	Activity Score	Genotypes	Examples of CYP2C19 diplotypes
CYP2C19 ultrarapid metabolizer	n/a	An individual carrying two increased function alleles	*17/*17
CYP2C19 rapid metabolizer	n/a	An individual carrying one normal function allele and one increased function allele	*1/*17
CYP2C19 normal metabolizer	n/a	An individual carrying two normal function alleles	*1/*1
Likely CYP2C19 intermediate metabolizer	n/a	An individual carrying one normal function allele and one decreased function allele or one increased function allele and one decreased function allele or two decreased function alleles	*1/*9, *9/*17, *9/*9
CYP2C19 intermediate metabolizer	n/a	An individual carrying one normal function allele and one no function allele or one increased function allele and one no function allele	*1/*2, *1/*3, *2/*17, *3/*17
Likely CYP2C19 poor metabolizer	n/a	An individual carrying one decreased function allele and one no function allele	*2/*9, *3/*9
CYP2C19 poor metabolizer	n/a	An individual carrying two no function alleles	*2/*2, *3/*3, *2/*3
Indeterminate	n/a	An individual carrying one or two uncertain function alleles	*1/*12, *2/*12, *12/*14

Questions

- How does your institution implement "possible" phenotypes into the EHR?
- Do the terms we use for these modifiers make sense?
- If not, what are other terms we should/could consider?

"Possible" phenotype assignment using SNOMED-CT terms

Cyrine-Eliana Haidar, PharmD., BCPS, BCOP Clinical Pharmacogenetics Coordinator St. Jude Children's Research Hospital <u>cyrine.haidar@stjude.org</u>

Problem List View in EHR

- Pro	blems						
4	Add	🛒 Modify 🛸 Convert 🔣 No Chronic Problems	s Displa	ay: Active	.		
	Annotated	Display 🔺	Ranking	Vocabulary	Code	Classification	<u>i</u>
6	OPYD poor metabolizer			SNOMED CT	3532686015	Medical	
0	G6PD - G	lucose-6-phosphate dehydrogenase deficiency		SNOMED CT	1220687019	Medical	
0	NUDT15	POOR METABOLIZER		SNODO Codes	NUDT15 POOR	Medical	
0	Opioid co	ontract exists		IMO	24288042	Medical	
0	Opioid us	se agreement exists		IMO	24288036	Medical	
0	PID 12345	56 SID 789456				Medical	
6	UGT1A1 p	poor metabolizer		SNOMED CT	3532666019	Medical	
6						Medical	

Prob	lens				
FIOD	iems				
	Annotated Display 🔺	Ranking	Vocabulary	Code	Classification
6	DPYD poor metabolizer		SNOMED CT	3532686015	Medical
6	G6PD - Glucose-6-phosphate dehydrogenase deficiency		SNOMED CT	1220687019	Medical
6	NUDT15 POOR METABOLIZER		SNODO Codes	NUDT15 POOR	Medical
6	Opioid contract exists		IMO	24288042	Medical
6	Opioid use agreement exists		IMO	24288036	Medical
a	PID 123456_SID 789456				Medical

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CYP2C19 poor metabolizer options (SNOMED CT and IMO terms)

Search: Search by Name		Search by Code			
Terminology: IMO, SNODO Codes, SNOMED CT Terminology Axis: <all axes="" terminology=""> </all>					
View Synonym 🔚 Concept Family 📲 Multi Axial	Cross Mapping				
Term	Code	Terminology 💌	Terminology Axis		
CYP2C19 poor metaboliser	3717119016	SNOMED CT	Clinical Finding		
CYP2C19 poor metabolizer	3533642015	SNOMED CT	Clinical Finding		
CYP2C19 poor metabolizer	1493690520	IMO			
	140000020				

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*Confirmation Confirmed •	*Classification Medical	*Status ↓ Active	Cancel Reason	~			
Resolved At: Age Resolved: Date Resolved: Date <t< td=""></t<>							
Qualifier	Severity Cla	385	Severity	Course	•		
Status date:	Day • 02/06/2020	•					

Confirmation status options

*Problem			Laterality	Responsible Provider	
CYP2C19 poor metaboliz	er	👫 🗌 Free Text	-		
Display As		At:Age	Onset: Date	Comments	
CYP2C19 possible poor i	metabolizer	39 Year	s 🚽 01/29/2020	•	
*Confirmation	*Classification	*Status	Cancel Reason		
Possible	- Medical		•	-	
Axis I diagnosis Axis II diagnosis Axis III diagnosis Axis IV diagnosis Complaint of Confirmed Differential	Resolved At: Age	Resolved: Data			
Probable	wiver Relationships S	econdary Description	Related Problems		
Provisional Rule out	Severity C	Class •	Severity	Course	
Status date:	Day - 01/29/2020	-			

Confirmation Status of "Possible"

*Problem CYP2C19 poor metabolizer		Free Text	Laterality -	Responsible Provider	
Display As CYP2C19 possible poor me	tabolizer	At:Age 39 Year:	Conset: Date 01/29/2020	Comments	
*Confirmation Possible +	*Classification Medical	*Status ← Active	Cancel Reason	-	
Relate Existing Problem Hide Additional Details Status Details Careging	Resolved At: Age	Hesolved: Date	Related Problems		
Qualifier	Severity Clas	:S	Severity	Course	
Status date:	Day • 01/29/2020	▲			

Possible CYP2C19 Poor Metabolizer in Problem List

Problems					
🕂 Add 🛒 Modify 🐃 Convert 🚯 No Chronic Problems	s Displa	ay: Active	▼.		
Annotated Display 🔺	Ranking	Vocabulary	Code	Classification	<u>u</u>
Asymptomatic human immunodeficiency virus infection		SNOMED CT	152322017	Medical	
CYP2C19 possible poor metabolizer		SNOMED CT	3533642015	Medical	
G CYP2C9 poor metabolizer		SNOMED CT	3532694010	Medical	
CYP2D6 ultra-rapid metabolizer		SNOMED CT	3532664016	Medical	
OPYD poor metabolizer		SNOMED CT	3532686015	Medical	
G6PD - Glucose-6-phosphate dehydrogenase deficiency		SNOMED CT	1220687019	Medical	
NUDT15 POOR METABOLIZER		SNODO Codes	NUDT15 POOR	Medical	
Opioid contract exists		IMO	24288042	Medical	
Opioid use agreement exists		IMO	24288036	Medical	
ID 123456 SID 789456				Medical	
UGT1A1 poor metabolizer		SNOMED CT	3532666019	Medical	
()				Medical	
6		Multum Allergy Category			

CDS alert wording for clopidogrel for CYP2C19 Possible PM



WARNING

Based on the genotype result, this patient MAY be a CYP2C19 poor metabolizer. If clopidogrel is prescribed to a CYP2C19 poor metabolizer for some indications, decreased effectiveness is likely. The use of an alternative agent is recommended. Please consult a clinical pharmacist, review the pharmacogenetics tab or click on the link below for more information.

Alert Action

History

Cancel Clopidogrel order

Continue Clopidogrel order

Add'l info

ΟK

CDS alert wording for clopidogrel for CYP2C19 PM

WARNING

Based on the genotype result, this patient is predicted to be a CYP2C19 poor metabolizer. If clopidogrel is prescribed to a CYP2C19 poor metabolizer for some indications, decreased effectiveness is likely. The use of an alternative agent is recommended. Please consult a clinical pharmacist, review the pharmacogenetics tab or click on the link below for more information.

Alert Action

History

Cancel Clopidogrel order

Continue Clopidogrel order

Add'l info

ΟK