CYP2D6 Pharmacogenetics Consultation UF Health Personalized Medicine Program

HPI:

Patient full name (MRN 00000), is currently enrolled in the research study, Implementing Genomics in Practice (IGNITE): CYP2D6 Genotype-Guided Pain Management in Patients Undergoing Arthroplasty Surgery (IRB 201800445). This patient provided written consent on --/--/---- and was randomized to the genotype-guided group.

Drug allergies:

Add

CYP2D6 interacting drugs (as of --/--):

Add

Pharmacogenetic test results (--/--):

CYP2D6 Genotype: */*

CYP2D6 Phenotype: Normal Metabolizer (NM); fully functional CYP2D6 enzyme activity

NOTE: Because of a CYP2D6 drug interaction, this patient may resemble an Intermediate Metabolizer

(IM) and have decreased CYP2D6 activity.

Interpretation:

Tramadol, codeine, and to a lesser extent hydrocodone and oxycodone are converted to more potent metabolites by the CYP2D6 enzyme. Based on the genotype result, this patient is predicted to be a normal metabolizer of CYP2D6 substrates. However, drug interactions can alter this patient's metabolizer status. *** moderately inhibits the activity of CYP2D6 and can cause individuals with a normal CYP2D6 genotype to resemble intermediate metabolizers. Intermediate metabolizer status is associated with decreased production of more potent forms of CYP2D6-dependent opioids. Therefore, this patient may get little to no pain relief from tramadol, codeine, hydrocodone, and oxycodone.

Recommendations:

Tramadol, codeine, hydrocodone, and oxycodone are **NOT RECOMMENDED** because this patient is predicted to be an **INTERMEDIATE METABOLIZER** of these drugs based on use of ***. **Consider an alternative opioid** such as morphine or hydromorphone that is not affected by CYP2D6 metabolism status, **a non-opioid** analgesic, **or discontinue use of** *** and start alternative therapy with a non-interacting drug.

For questions regarding this recommendation, please contact the UF Health Personalized Medicine Program:

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References:

- 1. Crews, K.R., et al. CPIC guidelines for *CYP2D6* genotype and codeine therapy: 2014 update. Clin Pharmacol Ther. 2014;95:376-82.
- 2. Stamer, U.M., et al. Impact of CYP2D6 genotype on postoperative tramadol analgesia. Pain 2003;105:231-
- 3. FDA: Drug Development and Drug Interactions: Table of Substrates, Inhibitors and Inducers. https://www.fda.gov/Drugs/DevelopmentApprovalProcess/DevelopmentResources/DrugInteractionsLabeling/ucm093664.htm