

Dosing INGREZZA® (valbenazine) Capsules and INGREZZA® SPRINKLE (valbenazine) Capsules in Patients Taking Cytochrome P450 2D6 (CYP2D6) Inhibitors

Thank you for contacting Neurocrine Biosciences with your unsolicited Medical Information request regarding the dose of INGREZZA and INGREZZA SPRINKLE in patients taking medications considered strong cytochrome P450 2D6 (CYP2D6) inhibitors.

INGREZZA and INGREZZA SPRINKLE are indicated in adults for the treatment of tardive dyskinesia and for the treatment of chorea associated with Huntington's disease.¹

The full Prescribing Information states the following¹:

DOSAGE AND ADMINISTRATION

2.5 Dosage Recommendations for Concomitant Use with Strong CYP2D6 Inhibitors

Coadministration with Strong CYP2D6 Inhibitors

The recommended dosage for patients receiving strong CYP2D6 inhibitors is INGREZZA or INGREZZA SPRINKLE 40 mg once daily.

WARNINGS AND PRECAUTIONS

5.4 QT Prolongation

INGREZZA and INGREZZA SPRINKLE may prolong the QT interval, although the degree of QT prolongation is not clinically significant at concentrations expected with recommended dosing. In patients taking a strong CYP2D6 or CYP3A4 inhibitor, or who are CYP2D6 poor metabolizers, INGREZZA and INGREZZA SPRINKLE concentrations may be higher and QT prolongation clinically significant. For patients who are CYP2D6 poor metabolizers or are taking a strong CYP2D6 inhibitor, dose reduction may be necessary. For patients taking a strong CYP3A4 inhibitor, reduce the dose of INGREZZA or INGREZZA SPRINKLE to 40 mg once daily. INGREZZA and INGREZZA SPRINKLE should be avoided in patients with congenital long QT syndrome or with arrhythmias associated with a prolonged QT interval. For patients at increased risk of a prolonged QT interval, assess the QT interval before increasing the dosage.

DRUG INTERACTIONS

7.1 Drugs Having Clinically Important Interactions with INGREZZA and INGREZZA SPRINKLE

Strong CYP2D6 Inhibitors	
<i>Clinical Implication:</i>	Concomitant use of INGREZZA or INGREZZA SPRINKLE with strong CYP2D6 inhibitors increased the exposure (C_{max} and AUC) to valbenazine's active metabolite compared with the use of INGREZZA or INGREZZA SPRINKLE alone. Increased exposure of active metabolite may increase the risk of exposure-related adverse reactions.
<i>Prevention or Management:</i>	Reduce INGREZZA or INGREZZA SPRINKLE dose when INGREZZA or INGREZZA SPRINKLE is coadministered with a strong CYP2D6 inhibitor.
<i>Examples^{2,3}:</i>	Paroxetine, fluoxetine, quinidine

This letter and the enclosed material are provided in response to your unsolicited medical information inquiry. Please feel free to contact Neurocrine Medical Information at (877) 641-3461 or medinfo@neurocrine.com if you would like to request additional information.

Reference:

1. INGREZZA [package insert]. San Diego, CA: Neurocrine Biosciences, Inc.
2. Hoefl D. An overview of clinically significant drug interactions between medications used to treat psychiatric and medical conditions. *Mental Health Clinician*. 2014;4(3):118–130.
3. United States Food and Drug Administration. Drug Development and Drug Interactions | Table of Substrates, Inhibitors and Inducers. <https://www.fda.gov/drugs/drug-interactions-labeling/drug-development-and-drug-interactions-table-substrates-inhibitors-and-inducers#table3-2>. Accessed November 6, 2023.

Enclosure:

- A. INGREZZA [package insert]. San Diego, CA: Neurocrine Biosciences, Inc.
- B. INGREZZA [Important Safety Information]. San Diego, CA: Neurocrine Biosciences, Inc.