PDL DRUG REVIEW

Proprietary Name: Nurtec® ODT
Common Name: rimegepant tablet, orally disintegrating
PDL Category: CGRP Inhibitors

Comparable Products	Preferred Drug List Status
Reyvow	Non-Preferred with Conditions
Sumatriptan	Preferred
Ubrelvy	Non-Preferred with Conditions

Summary

Pharmacology/Usage: Rimegepant, the active ingredient of Nurtec® ODT, is a calcitonin gene-related peptide (CGRP) receptor antagonist.

Indication: For the acute treatment of migraine with or without aura in adults. Nurtec® ODT is not indicated for the preventive treatment of migraine.

There is no pregnancy category for this medication; however, the risk summary indicates that there are no adequate data on the developmental risk associated with use in pregnant women. The safety and efficacy of use in the pediatric population have not been established.

Dosage Form: Orally Disintegrating Tablets: 75mg

Recommended Dosage: Remove the ODT from the blister and place on the tongue. Alternatively, the ODT may be placed under the tongue. The ODT will disintegrate in saliva so that it can be swallowed without additional liquid. The recommended dose is 75mg PO, with the maximum dose in a 24-hour period being 75mg. In addition, the safety of treating more than 15 migraines in a 30-day period has not been established.

Dose adjustments are not required with mild or moderate hepatic impairment; however, avoid use in patients with severe hepatic impairment. Dose adjustments are not required with mild, moderate, or severe renal impairment; however, avoid use in patients with end-stage renal disease.

Drug Interactions: Avoid concomitant use of Nurtec® ODT with strong inhibitors of CYP3A4. Avoid another dose of Nurtec® ODT within 48 hours when it is concomitantly administered with moderate inhibitors of CYP3A4.

Avoid concomitant use of Nurtec® ODT with strong or moderate inducers of CYP3A. In addition, avoid concomitant use of Nurtec® ODT with inhibitors of P-gp or BCRP.

Box Warning: There is no box warning listed with this product.

Common Adverse Drug Reactions: *Listed % incidence for adverse drug reactions= reported % incidence for drug (Nurtec®) minus reported % incidence for placebo. Please note that an incidence of 0% means the incidence was the same as or less than placebo.* The most frequently reported adverse events included nausea (1.6%).

Hypersensitivity reactions, including dyspnea and rash, have occurred with Nurtec® ODT in clinical studies. Hypersensitivity reactions can occur days after use and delayed serious hypersensitivity has occurred. If a hypersensitivity reaction occurs, discontinue Nurtec® ODT and start appropriate therapy.

Contraindications: In patients with a history of hypersensitivity reactions to rimegepant or any of its components

Manufacturer: Biohaven Pharmaceuticals, Inc

Analysis: The efficacy of Nurtec® ODT for the acute treatment of migraine with or without aura in adults was assessed in a randomized, double-blind, placebo-controlled study.

In study 1, patients were instructed to treat a migraine of moderate to severe headache pain intensity, being randomized to either Nurtec® ODT (N=732) or placebo (N=734). Rescue medications (such as APAP, NSAIDs, and/or an antiemetic) were allowed 2 hours after the initial treatment, but other forms of rescue medication such as triptans were not allowed within 48 hours of initial treatment. About 14% were taking preventive medications for migraine at baseline; however, none in study 1 were on concomitant preventive medication that act on the CGRP pathway.

The primary endpoints were pain freedom and most bothersome symptoms (MBS) freedom at 2 hours after dosing. Pain freedom was defined as a reduction of moderate or severe headache pain to no headache pain, while MBS freedom was defined as the absence of the self-identified MBS (i.e. photophobia, phonophobia, nausea). Among patients who selected an MBS, the most commonly selected symptom was photophobia (54%), followed by nausea (28%) and phonophobia (15%). Results of this study suggested that the percentage of patients achieving headache pain freedom and MBS freedom 2 hours after a single dose was statistically significantly greater in patients who received Nurtec® ODT as compared with placebo.

Results are summarized in the table below, which was adapted from the prescribing information. The table also includes some secondary endpoints assessed.

	Nurtec® ODT	Placebo
Pain Free at 2 hours		
number of responders/Number in group	142/669	74/682
% Responders	21.2%	10.9%
Difference from placebo; p-value	10.3%; p<0.001	
Calculated NNT per CHC	10	
MBS Free at 2 hours		
number of responders/Number in group	235/669	183/682
% Responders	35.1%	26.8%
Difference from placebo; p-value	8.3%; p=0.001	
Calculated NNT per CHC	13	
Pain Relief at 2 hours		
number of responders/Number in group	397/669	295/682
% Responders	59.3%	43.3%
Difference from placebo; p-value	16.1%; p<0.001	
Calculated NNT per CHC	7	
Sustained Pain Freedom 2-48 hours		
number of responders/Number in group	90/669	37/682
% Responders	13.5%	5.4%

	Nurtec® ODT	Placebo
Difference from placebo; p-value	8%; p<0.001	
Calculated NNT per CHC	13	
Use of Rescue Medication within 24 hours		
number of responders/Number in group	95/669	199/682
% Responders	14.2%	29.2%
Difference from placebo; p-value	-15%; p<0.001	
Calculated NNT per CHC	7	
% of Patients reporting normal function at 2 hours		
number of responders/Number in group	255/669	176/682
% Responders	38.1%	25.8%
Difference from placebo; p-value	12.3%; p<0.001	
Calculated NNT per CHC	9	

Place in Therapy: Nurtec® ODT is a CGRP receptor antagonist indicated for the acute treatment of migraine with or without aura in adults. It is not indicated for the preventive treatment of migraine. In a clinical trial, Nurtec® ODT was found to be significantly more effective than placebo for pain freedom at 2 hours and most bothersome symptom freedom at 2 hours. A 2019 network meta-analysis by Xu et al[2] included 10 trials to assess the safety and efficacy of CGRP receptor antagonists. While olcegepant (OR 4.09; not available in the US) and ubrogepant (OR 2.11) were more effective than placebo and olcegepant treatment was superior to the other five treatments, significant differences were not seen between telcagepant (not available in the US), olcegepant, and rimegepant for adverse events.

There is no evidence at this time to support that Nurtec® ODT is safer or more effective than the currently preferred medications. It is therefore recommended that Nurtec® ODT remain non-preferred and require prior authorization and be available to those who are unable to tolerate or who have failed on preferred medications.

PDL Placement:
- ☐ **Preferred**
- ☒ **Non-Preferred with Conditions**
- ☒ **Refer to DUR for PA Criteria**

References

[1] Nurtec ODT [package insert]. New Haven, CT: Biohaven Pharmaceuticals, Inc; 2020.
[2] Xu F, Sun W. Network meta-analysis of calcitonin gene-related peptide receptor antagonists for the acute treatment of migraine. *Front Pharmacol*. 2019; 10:795.