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5.70.003

Section: Prescription Drugs Effective Date: April 1, 2024

Subsection: Analgesics and Anesthetics Original Policy Date: March 5, 2005

Subject: 5-HT1 Agonists (Triptans) Page: 1 of 8

Last Review Date: March 8, 2024

5-HT1 Agonists (Triptans)

Description

Almotriptan
Amerge (naratriptan)
Frova (frovatriptan)
Imitrex, Onzetra Xsail, Tosymra NS, Treximet (sumatriptan)
Imitrex, Zembrace (sumatriptan injection)
Relpax (eletriptan)
Zomig, Zomig-ZMT (zolmitriptan)

Background

The selective serotonin receptor agonists, or "triptans", are a class of medications that have the ability to stop a migraine. Triptans work by binding to serotonin receptors in the brain. Specifically, per Drug Facts and Comparisons pharmacology of the Serotonin 5-HT₁ Receptor Agonists (Triptans): The vascular 5-HT₁ receptor subtype is present on the human basilar artery and in the vasculature of isolated human dura mater. Current theories on the etiology of migraine headaches suggest that symptoms are caused by local cranial vasodilation or the release of vasoactive and proinflammatory peptides from sensory nerve endings in an activated trigeminal system. The therapeutic activity of the serotonin 5-HT₁ receptor agonists in migraine most likely can be attributed to agonist effects at 5-HT_{1B/1D} receptors on the extracerebral, intracranial blood vessels that become dilated during a migraine attack and on nerve terminals in the trigeminal system. Activation of these receptors results in cranial vessel constriction, inhibition of neuropeptide release, and reduced transmission in trigeminal pain pathways (1).

Regulatory Status

^{*}Maxalt is found in a separate policy.

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FDA-approved indication: Triptan medications are indicated for the acute treatment of migraine attacks with or without aura in adults (2-14).

<u>Limitations of Use</u>: (2-14)

- Use only after a clear diagnosis of migraine has been established
- Not intended for the prophylactic therapy of migraine
- Not indicated for the treatment of cluster migraine

Triptan medications are not intended for use in the management of hemiplegic or basilar migraine (2-14).

Off-Label Use: (15)

Triptans have been found to be safe and effective in the pediatric and adolescent population.

Related policies

Butalbital analgesics, Dihydroergotamine Nasal Sprays, Elyxyb, Maxalt, Migraine CGRP Antagonists IV, Migraine CGRP Antagonists Nasal, Migraine CGRP Antagonists Oral, Migraine CGRP Antagonists SC, Migraine Powders

Policy

This policy statement applies to clinical review performed for pre-service (Prior Approval, Precertification, Advanced Benefit Determination, etc.) and/or post-service claims.

Triptan medications may be considered **medically necessary** if the conditions indicated below are met.

Triptan medications may be considered **investigational** for all other indications.

Prior-Approval Requirements

Age 6 years of age or older

Ages 6-11 must be prescribed by a neurologist

Diagnoses

Patient must have **ONE** of the following:

- 1. Migraine, with aura (classic)
- 2. Migraine, without aura (common)
- 3. Sumatriptan injection **only**: Cluster headache acute treatment

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AND ALL of the following:

- a. Patient is currently using migraine prophylactic therapy **OR** the patient has had an inadequate treatment response, intolerance, or contraindication to migraine prophylactic therapy (e.g., divalproex sodium, topiramate, valproate sodium, metoprolol, propranolol, etc.)
- b. **NO** hemiplegic migraine
- c. NO basilar migraine
- d. **NO** dual therapy with a calcitonin gene related peptide (CGRP) antagonist for acute migraine treatment (e.g., Nurtec ODT, Ubrelvy)
- e. NO dual therapy with Reyvow (lasmiditan) or Elyxyb (celecoxib)
- f. NO other PA on file for any triptan agent

Prior-Approval Renewal Requirements

Age 6 years of age or older

Ages 6-11 must be prescribed by a neurologist

Diagnoses

Patient must have **ONE** of the following:

- 1. Migraine, with aura (classic)
- 2. Migraine, without aura (common)
- 3. Sumatriptan injection only: Cluster headache acute treatment

AND ALL of the following:

- a. NO hemiplegic migraine
- b. NO basilar migraine
- NO dual therapy with a calcitonin gene related peptide (CGRP) antagonist for acute migraine treatment (e.g., Nurtec ODT, Ubrelvy)
- d. **NO** dual therapy with Reyvow (lasmiditan) or Elyxyb (celecoxib)
- e. NO other PA on file for any triptan agent

Policy Guidelines

Pre - PA Allowance

Age 12 years of age or older

No Pre-PA Allowance for under 12 years of age

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Quantity

Patients are allowed Pre-PA quantities of <u>up to TWO</u> triptan medications only.

Medication	Strength	Quantity
almotriptan	6.25 mg	48 tablets per 90 days AND/OR
almotriptan	12.5 mg	24 tablets per 90 days AND/OR
Amerge	1 mg	63 tablets per 90 days AND/OR
Amerge	2.5 mg	27 tablets per 90 days AND/OR
Frova	2.5 mg	36 tablets per 90 days AND/OR
Relpax	20 mg	36 tablets per 90 days AND/OR
Relpax	40 mg	18 tablets per 90 days AND/OR
Sumatriptan	5 mg nasal spray	96 units per 90 days AND/OR
Sumatriptan	10 mg nasal spray (Tosymra)	48 units per 90 days AND/OR
Sumatriptan	20 mg nasal spray	24 units per 90 days AND/OR
Sumatriptan	25 mg tablets	99 tablets per 90 days AND/OR
Sumatriptan	50 mg tablets	45 tablets per 90 days AND/OR
Sumatriptan	100 mg tablets	27 tablets per 90 days AND/OR
Sumatriptan	85 mg/ 500 mg (Treximet)	32 tablets per 90 days AND/OR
Sumatriptan (1 kit = 8 doses or 8 pouches containing 2 nosepieces/units per pouch = 16 units per kit)	11mg nasal powder (Onzetra Xsail)	6 x 8-dose kits (96 units) per 90 days AND/OR
Sumatriptan injection (1 kit = 2 injections)	4 mg/0.5ml injection kits	18 kits per 90 days AND/OR
Sumatriptan injection (1 kit = 2 injections)	6 mg/0.5 ml injection kits	12 kits per 90 days AND/OR
Sumatriptan injection	6mg/0.5ml injection vials	25 vials per 90 days AND/OR
Sumatriptan injection	3mg injection (Zembrace)	36 syringes per 90 days AND/OR
Zomig	2.5 mg tablets	36 tablets per 90 days AND/OR
Zomig	2.5 mg nasal spray	36 units per 90 days AND/OR
Zomig	5 mg tablets	18 tablets per 90 days AND/OR
Zomig	5 mg nasal spray	18 units per 90 days

Prior - Approval Limits

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Quantity

Medication	Strength	Quantity
almotriptan	6.25 mg	72 tablets per 90 days OR
almotriptan	12.5 mg	36 tablets per 90 days OR
Amerge	1 mg	90 tablets per 90 days OR
Amerge	2.5 mg	36 tablets per 90 days OR
Frova	2.5 mg	54 tablets per 90 days OR
Relpax	20 mg	54 tablets per 90 days OR
Relpax	40 mg	24 tablets per 90 days OR
Sumatriptan	5 mg nasal spray	144 units per 90 days OR
Sumatriptan	10 mg nasal spray (Tosymra)	72 units per 90 days OR
Sumatriptan	20 mg nasal spray	36 units per 90 days OR
Sumatriptan	25 mg tablets	144 tablets per 90 days OR
Sumatriptan	50 mg tablets	63 tablets per 90 days OR
Sumatriptan	100 mg tablets	36 tablets per 90 days OR
Sumatriptan	85 mg/ 500 mg (Treximet)	45 tablets per 90 days OR
Sumatriptan (1 kit = 8 doses or 8 pouches containing 2 nosepieces/units per pouch = 16 units per kit)	11mg nasal powder (Onzetra Xsail)	9 x 8-dose kits (144 units) per 90 days OR
Sumatriptan injection (1 kit = 2 injections)	4 mg/0.5ml injection kits	27 kits per 90 days OR
Sumatriptan injection (1 kit = 2 injections)	6 mg/0.5 ml injection kits	18 kits per 90 days OR
Sumatriptan injection	6mg/0.5ml injection vials	35 vials per 90 days OR
Sumatriptan injection	3mg injection (Zembrace)	54 syringes per 90 days OR
Zomig	2.5 mg tablets	54 tablets per 90 days OR
Zomig	2.5 mg nasal spray	54 units per 90 days OR
Zomig	5 mg tablets	27 tablets per 90 days OR
Zomig	5 mg nasal spray	24 units per 90 days

Duration 6 months

Prior – Approval Renewal Limits

Same as above

Rationale

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Summary

Triptan medications are indicated for the acute treatment of migraine attacks with or without aura in adults. Triptans are not intended for the prophylactic therapy of migraine or for use in the management of hemiplegic or basilar migraine. Triptans have been found to be safe and effective in the pediatric and adolescent population (1-15).

Prior authorization is required to ensure the safe, clinically appropriate, and cost-effective use of Triptan medications while maintaining optimal therapeutic outcomes.

References

- Serotonin 5-HT1 Receptor Agonists (Triptans). Drug Facts and Comparisons. Facts & Comparisons [database online]. St. Louis, MO: Wolters Kluwer Health, Inc; April 2019.
- 2. Amerge [package insert]. Research Triangle Park, NC: GlaxoSmithKline; October 2020.
- 3. Almotriptan [package insert]. Morgantown, WV: Mylan Pharmaceuticals, Inc.; May 2017.
- 4. Frova [package insert]. Malvern, PA: Endo Pharmaceuticals Inc.; August 2018.
- 5. Relpax [Package Insert]. New York, NY: Pfizer Inc.; March 2020.
- 6. Imitrex nasal spray [package insert]. Research Triangle Park, NC: GlaxoSmithKline; December 2017.
- 7. Imitrex tablets [package insert]. Research Triangle Park, NC: GlaxoSmithKline; December 2020.
- 8. Tosymra nasal spray [package insert]. Maple Grove, MN: Upsher-Smith Laboratories, LLC; February 2021.
- 9. Onzetra Xsail [package insert]. Morristown, NJ: Currax Pharmaceuticals LLC; December 2019.
- 10. Treximet [package insert]. Morristown, NJ: Currax Pharmaceuticals LLC; January 2024.
- 11. Imitrex Injection [package insert]. Research Triangle Park, NC: GlaxoSmithKline; February 2023.
- 12. Zembrace SymTouch [package insert]. Princeton, NJ: Promius Pharma, LLC; February 2021.
- 13. Zomig and Zomig-ZMT [package insert]. Bridgewater, NJ: Amneal Pharmaceuticals LLC; May 2019.
- 14. Zomig Nasal Spray [package insert]. Bridgewater, NJ: Amneal Pharmaceuticals LLC; May 2019.
- 15. Evers S. The Efficacy of Triptans in Childhood and Adolescence Migraine. Curr Pain Headache Rep. 2013 July;17(7)342.

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Policy History	
Date	Action
March 2005	Change in Pre-PA Allowance and Prior – Approval Limits due to manufacturer's packaging. The drug is pre-packaged in boxes of 9 tablets, and some pharmacies will not split packaging – under the old quantity limits, this resulted in member's receiving less medication than they were approved for. This adjustment in PA limits sets the quantity limit at two times the pre-PA quantity limits, which is the same ratio as other drugs in this class.
April 2011 December 2012	Annual editorial review and update Changed quantity limit to 1.5 x FDA-approved dosage Annual editorial review
April 2013	Revised quantity limits to allow mail order to fill correctly
September 2014	Revision of age to allow pediatric and adolescent use Annual editorial review and reference update
June 2015	Annual editorial review and reference update
March 2016	Annual editorial review and reference update Policy number changed from 5.02.03 to 5.70.03
March 2017	Annual review and reference update
March 2018 November 2018	Annual editorial review and reference update Annual editorial review and reference update. Addition of no dual therapy with CGRP antagonist requirement and no dual therapy with another PA for any triptan agent
March 2019	Annual review
September 2019	Revised quantity limits to quantity per 90 days
November 2019 December 2019	Addition of no dual therapy with Reyvow Annual review
March 2020	Annual review and reference update
June 2020	Annual review
March 2021	Annual review and reference update
April 2021	Added no dual therapy with a CGRP antagonist for acute migraine treatment. Revised no dual therapy requirement after 6 months of a prophylactic CGRP antagonist. Added initiation requirement to be on a migraine prophylactic therapy or have an inadequate treatment response, intolerance, or contraindication to migraine prophylactic therapy
June 2021	Annual review
September 2021 March 2022	Annual review and reference update Annual review. Per SME, removed requirement of "no dual therapy after 6 months with a prophylactic CGRP antagonist"

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April 2022 Renamed policy 5-HT1 Agonists (Triptans). Listed Axert as generic

(almotriptan) due to brand being discontinued. Combined policies Amerge, Axert, Frova, Relpax, Sumatriptan, Sumatriptan Injection, and Zomig. Added no dual therapy with Elyxyb. Added Pre-PA quantity statement that

patients are allowed Pre-PA for two triptan medications only

June 2022 Annual review

March 2023 Annual review and reference update. Changed policy number to 5.70.003

September 2023 Annual review

March 2023 Annual review and reference update

Keywords

This policy was approved by the FEP® Pharmacy and Medical Policy Committee on March 8, 2024 and is effective on April 1, 2024.