

## **Clinical Policy: Celecoxib (Celebrex, Elyxyb)**

Reference Number: CP.PMN.122

Effective Date: 01.01.07

Last Review Date: 05.22

Line of Business: Commercial, HIM, Medicaid

[Revision Log](#)

See [Important Reminder](#) at the end of this policy for important regulatory and legal information.

### **Description**

Celecoxib (Celebrex<sup>®</sup>, Elyxyb<sup>™</sup>) is a nonsteroidal anti-inflammatory drug (NSAID).

### **FDA Approved Indication(s)**

Celebrex is indicated for the treatment of:

- Osteoarthritis
- Rheumatoid arthritis
- Juvenile rheumatoid arthritis in patients 2 years and older
- Ankylosing spondylitis
- Acute pain
- Primary dysmenorrhea

Elyxyb is indicated for the acute treatment of migraine with or without aura in adults.

### **Policy/Criteria**

*Provider must submit documentation (such as office chart notes, lab results or other clinical information) supporting that member has met all approval criteria.*

It is the policy of health plans affiliated with Centene Corporation<sup>®</sup> that Celebrex and Elyxyb are **medically necessary** when the following criteria are met:

#### **I. Initial Approval Criteria**

##### **A. All Indications (must meet all):**

1. For Celebrex requests, both of the following (a and b):
  - a. Age  $\geq$  2 years;
  - b. For brand Celebrex requests, member must use generic celecoxib, unless contraindicated or clinically significant adverse effects are experienced;
2. For Elyxyb requests, both of the following (a and b):
  - a. Age  $\geq$  18 years;
  - b. Diagnosis of migraine;
3. Member meets one of the following (a, b, c, d, or e):
  - a. Age  $>$  65 years;
  - b. Current use of a corticosteroid;
  - c. Current use of an anticoagulant (e.g., aspirin, warfarin, low molecular weight heparin, direct thrombin inhibitors, factor Xa inhibitors, clopidogrel);
  - d. Prior gastrointestinal bleed or active peptic ulcer disease (not gastroesophageal reflux disease [GERD]);

- e. Both of the following (i and ii):
  - i. Failure of a  $\geq 4$  week trial of meloxicam at up to maximally indicated doses, unless contraindicated or clinically significant adverse effects are experienced;
  - ii. Failure of a  $\geq 4$  week trial of one additional generic NSAID at up to maximally indicated doses, unless clinically significant adverse effects are experienced or all are contraindicated;
4. Dose does not exceed one of the following (a or b):
  - a. Celebrex: 800 mg (2 capsules) per day;
  - b. Elyxyb: 120 mg (4.8 mL) per day.

**Approval duration:**

**Medicaid/HIM** – 12 months

**Commercial** – Length of Benefit

**B. Other diagnoses/indications** (must meet 1 or 2):

1. If this drug has recently (within the last 6 months) undergone a label change (e.g., newly approved indication, age expansion, new dosing regimen) that is not yet reflected in this policy, refer to one of the following policies (a or b):
  - a. For drugs on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the no coverage criteria policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.33 for health insurance marketplace, and CP.PMN.255 for Medicaid; or
  - b. For drugs NOT on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the non-formulary policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.103 for health insurance marketplace, and CP.PMN.16 for Medicaid; or
2. If the requested use (e.g., diagnosis, age, dosing regimen) is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized) AND criterion 1 above does not apply, refer to the off-label use policy for the relevant line of business: CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid.

**II. Continued Therapy**

**A. All Indications** (must meet all):

1. Member meets one of the following (a or b):
  - a. Currently receiving medication via Centene benefit or member has previously met initial approval criteria;
  - b. Member is currently receiving medication and is enrolled in a state and product with continuity of care regulations (*refer to state specific addendums for CC.PHARM.03A and CC.PHARM.03B*);
2. Member is responding positively to therapy;
3. If request is for a dose increase, new dose does not exceed one of the following (a or b):
  - a. Celebrex: 800 mg (2 capsules) per day;
  - b. Elyxyb: 120 mg (4.8 mL) per day.

**Approval duration:**

**Medicaid/HIM** – 12 months

**Commercial – Length of Benefit**

**B. Other diagnoses/indications (must meet 1 or 2):**

1. If this drug has recently (within the last 6 months) undergone a label change (e.g., newly approved indication, age expansion, new dosing regimen) that is not yet reflected in this policy, refer to one of the following policies (a or b):
  - a. For drugs on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the no coverage criteria policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.33 for health insurance marketplace, and CP.PMN.255 for Medicaid; or
  - b. For drugs NOT on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the non-formulary policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.103 for health insurance marketplace, and CP.PMN.16 for Medicaid; or
2. If the requested use (e.g., diagnosis, age, dosing regimen) is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized) AND criterion 1 above does not apply, refer to the off-label use policy for the relevant line of business: CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid.

**III. Diagnoses/Indications for which coverage is NOT authorized:**

- A. Non-FDA approved indications, which are not addressed in this policy, unless there is sufficient documentation of efficacy and safety according to the off label use policies – CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid, or evidence of coverage documents.

**IV. Appendices/General Information**

*Appendix A: Abbreviation/Acronym Key*

CABG: coronary artery bypass graft

FDA: Food and Drug Administration

GERD: gastroesophageal reflux disease

NSAID: nonsteroidal anti-inflammatory drug

*Appendix B: Therapeutic Alternatives*

*This table provides a listing of preferred alternative therapy recommended in the approval criteria. The drugs listed here may not be a formulary agent for all relevant lines of business and may require prior authorization.*

<b>Drug Name</b>	<b>Dosing Regimen</b>	<b>Dose Limit/ Maximum Dose</b>
naproxen sodium (Anaprox <sup>®</sup> , Anaprox DS <sup>®</sup> )	275 – 550 mg PO BID	1,650 mg/day
sulindac (Clinoril <sup>®</sup> )	150 – 200 mg PO BID	400 mg/day
salsalate (Disalcid <sup>®</sup> )	500 – 750 mg PO TID, titrated up to 3,000 mg/day	3,000 mg/day
piroxicam (Feldene <sup>®</sup> )	10 – 20 mg PO QD	20 mg/day
indomethacin (Indocin <sup>®</sup> )	25 – 50 mg PO BID -TID	200 mg/day
indomethacin SR (Indocin <sup>®</sup> SR)	75 mg PO QD - BID	150 mg/day

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
meclofenamate (Meclomen <sup>®</sup> )	50 – 100 mg PO Q4-6hr	400 mg/day
meloxicam (Mobic <sup>®</sup> )	7.5 – 15 mg PO QD	15 mg/day
ibuprofen (Motrin <sup>®</sup> )	400 – 800 mg PO Q6-8hr	3,200 mg/day
fenoprofen (Nalfon <sup>®</sup> )	200 mg PO Q4-6hr	3,200 mg/day
naproxen (Naprosyn <sup>®</sup> )	250 – 500 mg PO BID	1,500 mg/day
ketoprofen (Orudis <sup>®</sup> )	25 – 75 mg PO Q6-8hr	300 mg/day
nabumetone (Relafen <sup>®</sup> )	1000 mg PO QD or 500 mg PO BID	2,000 mg/day
tolmetin (Tolmetin <sup>®</sup> DS)	400 mg PO TID, titrated up to 1800 mg/day	1,800 mg/day
diclofenac sodium (Voltaren <sup>®</sup> )	50 mg PO Q6-8hr	200 mg/day
oxaprozin (Daypro <sup>®</sup> )	600 – 1,200 mg PO QD	1,800 mg/day
etodolac (Lodine <sup>®</sup> )	400 – 500 mg PO BID	1,200 mg/day

*Therapeutic alternatives are listed as Brand name<sup>®</sup> (generic) when the drug is available by brand name only and generic (Brand name<sup>®</sup>) when the drug is available by both brand and generic.*

*Appendix C: Contraindications/Boxed Warnings*

- Contraindication(s): hypersensitivity to celecoxib or any components of the drug product; history of asthma, urticaria, or other allergic-type reactions to aspirin or other NSAIDs; in the setting of coronary artery bypass graft (CABG) surgery; allergic-type reactions to sulfonamides
- Boxed warning(s): increased risk of serious cardiovascular thrombotic events, including myocardial infarction, and stroke; increased risk of serious gastrointestinal (GI) adverse events including bleeding, ulceration, and perforation of the stomach or intestines; in the setting of CABG surgery

*Appendix D: General Information*

- The risk vs. benefit of COX-II therapy should be individualized based on patient's previous GI history, other co-morbid conditions (e.g., angina, ischemic heart disease, myocardial infarction (MI), coronary artery disease, stroke), age, concurrent medications (e.g., warfarin, oral corticosteroids), duration and dose.
- Celebrex has been associated with an increased risk of serious adverse cardiovascular (CV) events in a long-term placebo-controlled trial. Based on the currently available data, FDA has concluded that an increased risk of serious adverse CV events appears to be a class effect of NSAIDs. FDA has requested that the package insert for all NSAIDs, including Celebrex, be revised to include a boxed warning to highlight the potential increased risk of CV events and the well described risk of serious, and potentially life-threatening, gastrointestinal bleeding.

**V. Dosage and Administration**

<b>Drug Name</b>	<b>Indication</b>	<b>Dosing Regimen</b>	<b>Maximum Dose</b>
Celecoxib (Celebrex)	Osteoarthritis	200 mg PO QD or 100 mg PO BID	800 mg/day
Celecoxib (Celebrex)	Rheumatoid arthritis	100 to 200 mg PO BID	800 mg/day
Celecoxib (Celebrex)	Juvenile rheumatoid arthritis	10-25 kg: 50 mg PO BID > 25 kg: 100 mg PO BID	200 mg/day
Celecoxib (Celebrex)	Ankylosing spondylitis	200 mg PO QD or 100 mg PO BID. If no effect is observed after 6 weeks, a trial of 400 mg (single or divided doses) may be of benefit.	800 mg/day
Celecoxib (Celebrex)	Acute pain or Primary dysmenorrhea	400 mg PO initially, followed by a 200 mg dose if needed on the first day. On subsequent days, 200 mg PO BID as needed	800 mg/day
Celecoxib (Elyxyb)	Migraine	120 mg PO PRN. Use the fewest number of days per month, as needed.	120 mg/day

**VI. Product Availability**

<b>Drug Name</b>	<b>Availability</b>
Celecoxib (Celebrex)	Capsules: 50 mg, 100 mg, 200 mg, and 400 mg
Celecoxib (Elyxyb)	Oral solution: 120 mg/4.8 mL (25 mg/mL)

**VII. References**

1. Celebrex Prescribing Information. New York, NY: G.D. Searle, LLC; May 2019. Available at: <http://www.celebrex.com/>. Accessed January 14, 2022.
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11. Juni, et al. Are selective COX 2 inhibitors superior to traditional non-steroidal anti-inflammatory drugs. *BMJ* 2002;324:1287-1288.
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Reviews, Revisions, and Approvals	Date	P&T Approval Date
2Q 2018 annual review: polices combined for Medicaid, HIM, and commercial lines of business; reference number changed from PPA to PMN; Medicaid: Added age and max dose; increased approval duration from 3/12 to 12/12; HIM: removed specific diagnoses; added age; decreased trials from 3 (meloxicam & 2 NSAIDs) to 2 (meloxicam & 1 NSAID); added a path to approval for those with high risk for gastroduodenal damage (>65 years, current steroid or anticoagulant use, or prior bleed); Commercial: added age; changed trial of 2 NSAIDs to meloxicam and 1 NSAID; references reviewed and updated.	02.20.18	05.18
2Q 2019 annual review: no significant changes. References reviewed and updated.	02.23.19	05.19
2Q 2020 annual review: no significant changes; references reviewed and updated.	02.07.20	05.20
RT4: added Elyxyb to policy.	05.14.20	
2Q 2021 annual review: no significant changes; revised HIM.PHAR.21 to HIM.PA.154; references reviewed and updated.	03.01.21	05.21
2Q 2022 annual review: no significant changes; added redirection to generic celecoxib for brand Celebrex requests per formulary status; limited use of Elyxyb to its FDA labeled indication; references reviewed and updated.	01.14.22	05.22
Template changes applied to other diagnoses/indications and continued therapy section.	10.03.22	

**Important Reminder**

This clinical policy has been developed by appropriately experienced and licensed health care professionals based on a review and consideration of currently available generally accepted standards of medical practice; peer-reviewed medical literature; government agency/program approval status; evidence-based guidelines and positions of leading national health professional organizations; views of physicians practicing in relevant clinical areas affected by this clinical policy; and other available clinical information. The Health Plan makes no representations and accepts no liability with respect to the content of any external information used or relied upon in developing this clinical policy. This clinical policy is consistent with standards of medical practice current at the time that this clinical policy was approved. “Health Plan” means a health plan that has adopted this clinical policy and that is operated or administered, in whole or in part, by Centene Management Company, LLC, or any of such health plan’s affiliates, as applicable.

The purpose of this clinical policy is to provide a guide to medical necessity, which is a component of the guidelines used to assist in making coverage decisions and administering benefits. It does not constitute a contract or guarantee regarding payment or results. Coverage decisions and the administration of benefits are subject to all terms, conditions, exclusions and limitations of the coverage documents (e.g., evidence of coverage, certificate of coverage, policy, contract of insurance, etc.), as well as to state and federal requirements and applicable Health Plan-level administrative policies and procedures.

This clinical policy is effective as of the date determined by the Health Plan. The date of posting may not be the effective date of this clinical policy. This clinical policy may be subject to applicable legal and regulatory requirements relating to provider notification. If there is a discrepancy between the effective date of this clinical policy and any applicable legal or regulatory requirement, the requirements of law and regulation shall govern. The Health Plan retains the right to change, amend or withdraw this clinical policy, and additional clinical policies may be developed and adopted as needed, at any time.

This clinical policy does not constitute medical advice, medical treatment or medical care. It is not intended to dictate to providers how to practice medicine. Providers are expected to exercise professional medical judgment in providing the most appropriate care, and are solely responsible for the medical advice and treatment of members. This clinical policy is not intended to recommend treatment for members. Members should consult with their treating physician in connection with diagnosis and treatment decisions.

Providers referred to in this clinical policy are independent contractors who exercise independent judgment and over whom the Health Plan has no control or right of control. Providers are not agents or employees of the Health Plan.

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and their representatives agree to be bound by such terms and conditions by providing services to members and/or submitting claims for payment for such services.

**Note:**

**For Medicaid members**, when state Medicaid coverage provisions conflict with the coverage provisions in this clinical policy, state Medicaid coverage provisions take precedence. Please refer to the state Medicaid manual for any coverage provisions pertaining to this clinical policy.

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