

Clinical Policy: Brolucizumab-dbl (Beovu)

Reference Number: CP.PHAR.445

Effective Date: 03.01.20

Last Review Date: 02.22

Line of Business: Commercial, HIM, Medicaid

[Coding Implications](#)

[Revision Log](#)

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

Description

Brolucizumab-dbl (Beovu[®]) is a vascular endothelial growth factor (VEGF) inhibitor.

FDA Approved Indication(s)

Beovu is indicated for the treatment of:

- Neovascular (wet) age-related macular degeneration (AMD)
- Diabetic macular edema (DME)

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[®] that Beovu is **medically necessary** when the following criteria are met:

I. Initial Approval Criteria

A. Ophthalmic Disease (must meet all):

1. Diagnosis of one of the following (a or b):
 - a. Neovascular (wet) AMD;
 - b. DME;
2. Prescribed by or in consultation with an ophthalmologist;
3. Age \geq 18 years;
4. Failure of bevacizumab intravitreal solution, unless contraindicated or clinically significant adverse effects are experienced;
**Prior authorization may be required for bevacizumab intravitreal solution. Requests for IV formulations of Avastin, Mvasi, and Zirabev will not be approved*
5. Dose does not exceed (a or b):
 - a. Neovascular AMD: 6 mg (1 vial) every 4 weeks for the first 3 doses;
 - b. DME: 6 mg (1 vial) every 6 weeks for the first 5 doses.

Approval duration:

Neovascular (wet) AMD – 4 months (3 loading doses only)

DME – 6 months (5 loading doses only)

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. Ophthalmic Disease (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 as evidenced by one of the following (a, b, c, or d):
 - a. Detained neovascularization;
 - b. Improvement/stabilization in visual acuity;
 - c. Maintenance of corrected visual acuity from prior treatment;
 - d. Supportive findings from optical coherence tomography or fluorescein angiography;
3. Member meets one of the following (a or b):
 - a. Dose does not exceed 6 mg (1 vial) every 12 weeks;
 - b. If request is for a dose increase, both of the following (i and ii):
 - i. Documentation supports evidence of continued disease activity;
 - ii. New dose does not exceed 6 mg (1 vial) every 8 weeks.

Approval duration: 6 months

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
- 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:

- 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

AMD: age-related macular degeneration

DME: diabetic macular edema

FDA: Food and Drug Administration

VEGF: vascular endothelial growth factor

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.

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
bevacizumab (Avastin [®])	1.25 to 2.5 mg administered by intravitreal injection every 4 weeks	2.5 mg/month

Therapeutic alternatives are listed as Brand name[®] (generic) when the drug is available by brand name only and generic (Brand name[®]) when the drug is available by both brand and generic.

Appendix C: Contraindications/Boxed Warnings

- Contraindication(s):
 - Ocular or periocular infections
 - Active intraocular inflammation
 - Known hypersensitivity to Beovu or any of the excipients
- Boxed warning(s): none reported

Appendix D: General Information

- In the HAWK study, brolucizumab 3 mg and 6 mg groups demonstrated non-inferiority to aflibercept 2 mg in terms of mean best corrected visual acuity (BCVA) to week 48 with respective least squares mean changes in BCVA from baseline being: 6.1, 6.6, and 6.8. Similar results were seen in HARRIER, in which brolucizumab 6 mg demonstrated non-inferiority to aflibercept 2 mg with a least squares mean change in BCVA from baseline being 6.9 and 7.6 respectively.
- Disease activity was assessed 8 weeks after the loading phase period, revealing a formal demonstration of superiority versus aflibercept in the HAWK study regarding duration of

effect. Additionally, in both studies, this advantage of brolucizumab 6 mg v. aflibercept was reflected in anatomic assessment; similar results were observed at Week 16 and Week 48.

- Based on observations from the HAWK and HARRIER studies, it was estimated that for brolucizumab-treated eyes, the probabilities for exclusively maintaining q12w dosing after loading through Week 48 were 49.4% and 55.6% in HAWK for 3 mg and 6 mg respectively and 51% in HARRIER. Time-to-event analyses revealed that most q8w treatment needed was identified during the first q12w interval (at Weeks 16 and 20).

V. Dosage and Administration

Indication	Dosing Regimen	Maximum Dose
Neovascular (wet) AMD	6 mg (1 vial) via intravitreal injection every 4 weeks for the first 3 months, then every 8 or 12 weeks thereafter	6mg (1 vial) every 2 months after loading period

VI. Product Availability

- Single-dose vial: 6 mg/0.05 mL (120 mg/mL solution in 0.05 mL total volume)
- Single-dose, preservative free, pre-filled syringe: 6 mg/0.05 mL (120 mg/mL solution in 0.05 mL total volume)

VII. References

1. Beovu Prescribing Information. East Hanover, NJ: Novartis Pharmaceuticals, Inc.; May 2022. Available at: https://www.accessdata.fda.gov/drugsatfda_docs/label/2020/761125s004lbl.pdf. Accessed June 14, 2022.
2. American Academy of Ophthalmology Retina/Vitreous Panel. Preferred Practice Pattern[®] Guidelines. Age-related macular degeneration. San Francisco, CA: American Academy of Ophthalmology; October 2019. Available at: www.aao.org/ppp. Accessed November 9, 2021.
3. Dugel PA, Koh A, Ogura Y, et al. HAWK and HARRIER: Phase 3, multicenter, randomized, double-masked trials of brolucizumab for neovascular age-related macular degeneration. Presented at: the American Academy of Ophthalmology Annual Meeting, Nov 11-14, 2017, New Orleans, LA. Accessed: 23 Oct. 2019. [https://www.aaojournal.org/article/S0161-6420\(18\)33018-5/](https://www.aaojournal.org/article/S0161-6420(18)33018-5/)
4. Brown DM, Emanuelli A, Bandello F, et al. KESTREL and KITE: 52-week results from two phase III pivotal trials of brolucizumab for diabetic macular edema. American Journal of Ophthalmology; June 2022; 238:157-175. <https://doi.org/10.1016/j.ajo.2022.01.004>.
5. American Academy of Ophthalmology Retina/Vitreous Panel. Preferred Practice Pattern[®] Guidelines. Diabetic Retinopathy. San Francisco, CA: American Academy of Ophthalmology; September 2019. Available at: www.aao.org/ppp. Accessed June 14, 2022.

Coding Implications

Codes referenced in this clinical policy are for informational purposes only. Inclusion or exclusion of any codes does not guarantee coverage. Providers should reference the most up-to-date sources of professional coding guidance prior to the submission of claims for reimbursement of covered services.

HCPCS Codes	Description
J0179	Injection, brolucizumab-dbl, 1mg

Reviews, Revisions, and Approvals	Date	P&T Approval Date
Policy created	11.26.19	02.20
Finalize HIM line of business per March SDC and prior clinical guidance.	03.03.20	
Ad Hoc update: clarified redirection from bevacizumab to Avastin as compounding pharmacies often break standard Avastin vials into smaller dosages specifically for ophthalmic use and there is a temporary CPT code not currently available to biosimilars.	10.01.20	
1Q 2021 annual review: no significant changes; references to HIM.PHAR.21 revised to HIM.PA.154; references reviewed and updated.	12.01.20	02.21
Ad Hoc update: updated redirection to “bevacizumab intravitreal solution” given availability of generic bevacizumab intravitreal solution and considering goal was to minimize use of IV bevacizumab products, most notably biosimilars; converted redirection language to “must use”	03.04.21	
Ad Hoc update: converted redirection language from “must use” to “Failure of” bevacizumab intravitreal solution.	08.03.21	
1Q 2022 annual review: no significant changes; references reviewed and updated.	11.09.21	02.22
RT4: criteria added for new FDA approved indication of DME; references reviewed and updated.	06.15.22	
Template changes applied to other diagnoses/indications and continued therapy section.	09.27.22	

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.

This clinical policy is the property of the Health Plan. Unauthorized copying, use, and distribution of this clinical policy or any information contained herein are strictly prohibited. Providers, members and their representatives are bound to the terms and conditions expressed herein through the terms of their contracts. Where no such contract exists, providers, members 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.

©2020 Centene Corporation. All rights reserved. All materials are exclusively owned by Centene Corporation and are protected by United States copyright law and international copyright law. No part of this publication may be reproduced, copied, modified, distributed, displayed, stored in a retrieval system, transmitted in any form or by any means, or otherwise published without the prior written permission of Centene Corporation. You may not alter or remove any trademark, copyright or other notice contained herein. Centene[®] and Centene Corporation[®] are registered trademarks exclusively owned by Centene Corporation.