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Section:	Prescription Drugs	**Effective Date:**	January 1, 2026
Subsection:	Topical Products	**Original Policy Date:**	February 17, 2017
Subject:	Ophthalmic VEGF Inhibitors	**Page:**	1 of 9

Last Review Date: December 12, 2025

Ophthalmic VEGF Inhibitors

Description

Beovu (brolucizumab-dbll)
Eylea (aflibercept), Ahzantive* (aflibercept-mrbb), Enzeevu* (aflibercept-abzv), Eydenzelt* (aflibercept-boav), Opuviz* (aflibercept-yszy), Pavblu** (aflibercept-ayyh), Yesafili* (aflibercept-jbvf)
Eylea HD (aflibercept)
Vabysmo (faricimab-svoa)

*These medications are included in this policy but are not available on the market as of yet

**Product covered on the medical benefit only

Background
Ophthalmic VEGF inhibitors are vascular endothelial growth factor (VEGF) inhibitors used to treat patients with a variety of ocular conditions. The VEGF inhibitors block the effects of VEGF-A and prevents the interaction of VEGF-A with its receptors (VEGFR1 and VEGFR2) on the surface of endothelial cells, reducing endothelial cell growth, vascular leakage, and new blood vessel formation (1-10).

Regulatory Status
FDA-approved indications: (1-10)

1. **Eylea** and its biosimilars are vascular endothelial growth favor (VEGF) inhibitors indicated for the treatment of patients with:
    - Neovascular (Wet) Age-Related Macular Degeneration (AMD)

- Macular Edema following Retinal Vein Occlusion (RVO)
- Diabetic Macular Edema (DME)
- Diabetic Retinopathy (DR)
- Retinopathy of Prematurity (ROP)

2. **Eylea HD** is a vascular endothelial growth favor (VEGF) inhibitor indicated for the treatment of patients with:
   - Neovascular (Wet) Age-Related Macular Degeneration (AMD)
   - Diabetic Macular Edema (DME)
   - Diabetic Retinopathy (DR)
3. **Beovu** is a vascular endothelial growth factor (VEGF) inhibitor indicated for the treatment of:
   - Neovascular (Wet) Age-Related Macular Degeneration (AMD).
   - Diabetic Macular Edema (DME)
4. **Vabysmo** is a vascular endothelial growth factor receptor (VEGF) and angiopoietin-2 (Ang-2) inhibitor indicated for the treatment of patients with:
   - Neovascular (Wet) Age-Related Macular Degeneration (AMD)
   - Diabetic Macular Edema (DME)
   - Macular Edema following Retinal Vein Occlusion (RVO)

VEGF inhibitors are contraindicated in ocular or periocular infections and in patients with active intraocular inflammation (1-10).

VEGF inhibitors must only be administered by a qualified physician. Adequate anesthesia and a topical broad-spectrum microbicide should be given prior to the injection. Increases in intraocular pressure have been seen within 30-60 minutes of an intravitreal injection (1-10).

The safety and effectiveness of Beovu, Eylea HD, and Vabysmo in pediatric patients have not been established. The safety and effectiveness of Eylea and its biosimilars has been demonstrated in pre-term infants with ROP (1-10).

Related policies
Bevacizumab, Lucentis, Susvimo

Policy

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

Ophthalmic VEGF inhibitors may be considered **medically necessary** if the conditions indicated below are met.

Ophthalmic VEGF inhibitors may be considered **investigational** for all other indications.

Prior-Approval Requirements

Age 18 years of age or older

Diagnoses

Eylea, Ahzantive, Enzeevu, Eydenzelt, Opuviz, Pavblu, and Yesafili only

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

1. Neovascular (wet) age-related macular degeneration (AMD)
2. Macular edema following retinal vein occlusion (RVO)
3. Diabetic macular edema (DME)
4. Diabetic retinopathy (DR)

Eylea HD only

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

1. Neovascular (wet) age-related macular degeneration (AMD)
2. Diabetic macular edema (DME)
3. Diabetic retinopathy (DR)

Beovu only

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

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

Vabysmo only

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

1. Neovascular (wet) age-related macular degeneration (AMD)
2. Diabetic macular edema (DME)
3. Macular edema following retinal vein occlusion (RVO)

AND ALL of the following for **ALL** medications:

a. Documented baseline visual acuity test
b. **NO** active intraocular inflammation

c. **NO** ocular or periocular infection
d. **NOT** used in combination with other vascular endothelial growth factor (VEGF) inhibitors for ocular indications (see Appendix 1)

Age No age requirement

Diagnosis

Eylea, Ahzantive, Enzeevu, Eydenzelt, Opuviz, Pavblu, and Yesafili only

Patient must have the following:

1. Retinopathy of prematurity (ROP)

AND ALL of the following:

a. **NO** active intraocular inflammation
b. **NO** ocular or periocular infection
c. **NOT** used in combination with other vascular endothelial growth factor (VEGF) inhibitors for ocular indications (see Appendix 1)

Prior – Approval *Renewal* Requirements

Age 18 years of age or older

Diagnoses

Eylea, Ahzantive, Enzeevu, Eydenzelt, Opuviz, Pavblu, and Yesafili only

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

1. Neovascular (wet) age-related macular degeneration (AMD)
2. Macular edema following retinal vein occlusion (RVO)
3. Diabetic macular edema (DME)
4. Diabetic retinopathy (DR)

Eylea HD only

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

1. Neovascular (wet) age-related macular degeneration (AMD)
2. Diabetic macular edema (DME)
3. Diabetic retinopathy (DR)

Beovu only

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

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

Vabysmo only

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

1. Neovascular (wet) age-related macular degeneration (AMD)
2. Diabetic macular edema (DME)
3. Macular edema following retinal vein occlusion (RVO)

AND ALL of the following for **ALL** medications:

a. Patient has demonstrated a positive clinical response to therapy (e.g., improvement or maintenance in best corrected visual acuity [BCVA] or visual field, or a reduction in the rate of vision decline or the risk of more severe vision loss)
b. **NO** active intraocular inflammation
c. **NO** ocular or periocular infection
d. **NOT** used in combination with other vascular endothelial growth factor (VEGF) inhibitors for ocular indications (see Appendix 1)

Age No age requirement

Diagnosis

Eylea, Ahzantive, Enzeevu, Eydenzelt, Opuviz, Pavblu, and Yesafili only

Patient must have the following:

1. Retinopathy of prematurity (ROP)

AND ALL of the following:

a. Patient has demonstrated a positive clinical response to therapy (e.g., no clinically significant reactivations of ROP)
b. **NO** active intraocular inflammation
c. **NO** ocular or periocular infection
d. **NOT** used in combination with other vascular endothelial growth factor (VEGF) inhibitors for ocular indications (see Appendix 1)

Policy Guidelines

Pre - PA Allowance

None

Prior - Approval Limits

Duration 12 months

Prior – Approval *Renewal* Limits

Same as above

Rationale

Summary

VEGF inhibitors prevent the binding and activation of VEGF receptors leading to a decrease in the neovascularization and vascular permeability associated with neovascular AMD and macular edema following RVO, DR, and DME. Patients taking VEGF inhibitors must be monitored and managed for intravitreal injection procedure associated effects, elevated intraocular pressure, and appropriate perfusion of the optic nerve head. VEGF inhibitors must only be administered by a retina trained ophthalmologist (1-10).

Prior approval is required to ensure the safe, clinically appropriate, and cost-effective use of VEGF Inhibitors while maintaining optimal therapeutic outcomes.

References

1. Eylea [package insert]. Tarrytown, NY: Regeneron Pharmaceuticals, Inc.; October 2024.
2. Eylea HD [package insert]. Tarrytown, NY: Regeneron Pharmaceuticals, Inc.; October 2024.
3. Beovu [package insert]. East Hanover, NJ: Novartis Pharmaceuticals Corporation; July 2024.
4. Vabysmo [package insert]. South San Francisco, CA: Genentech, Inc.; July 2024.
5. Opuviz [package insert]. Cambridge, MA: Biogen MA Inc.; May 2024.
6. Yesafili [package insert]. Cambridge, MA: Biocon Biologics Inc.; May 2024.
7. Ahzantive [package insert]. Martinsried, Germany: Formycon AG; June 2024.
8. Pavblu [package insert]. Thousand Oaks, CA: Amgen, Inc.; August 2024.
9. Enzeevu [package insert]. Princeton, NJ: Sandoz Inc.; August 2024.
10. Eydenzelt [package insert]. Jersey City, NJ: Celltrion USA, Inc.; October 2025.

Policy History

Date	Action
February 2017	Addition to PA
June 2017	Removal of Lucentis and the addition of Macugen Addition of the requirement: not be used in combination therapy with other vascular endothelial growth factor (VEGF) inhibitors
September 2017	Annual review
September 2018	Annual review and reference update
May 2019	Change to Eylea indication: patients with diabetic retinopathy (DR) no longer required to have concurrent DME
June 2019	Annual review
October 2019	Addition of Beovu
December 2019	Annual review
March 2020	Annual review and reference update
March 2021	Revised renewal requirement from “no loss of greater than 15 letters in visual acuity” to “patient has demonstrated a positive clinical response to therapy (e.g., improvement or maintenance in best corrected visual acuity [BCVA] or visual field, or a reduction in the rate of vision decline or the risk of more severe vision loss)”. Also removed the letter count requirement for initiation per FEP
June 2021	Annual review and reference update. Removed Macugen from policy and Appendix 1 due to being discontinued
March 2022	Annual review and reference update. Addition of Vabysmo to policy and added Susvimo and Vabysmo to Appendix 1
June 2022	Annual review and reference update. Per PI update, addition of indication for Beovu of diabetic macular edema
September 2022	Annual review
December 2022	Annual review
March 2023	Per PI update, added indication for Eylea of retinopathy of prematurity (ROP)
June 2023	Annual review
September 2023	Addition of Eylea HD to policy
December 2023	Annual review. Per PI update, added indication of macular edema following RVO to Vabysmo
March 2024	Annual review and reference update
May 2024	Added requirement not to use in combination with Syfovre
August 2024	Addition of Eylea biosimilars Ahzantive, Opuviz, and Yesafili
September 2024	Annual review and reference update
November 2024	Addition of Eylea biosimilar Pavblu
February 2025	Per FEP, removed requirement not to use in combination with Syfovre for ocular indications

March 2025	Annual review and reference update
April 2025	Addition of Eylea biosimilar Enzeevu
June 2025	Annual review
October 2025	Addition of Eylea biosimilar Eydenzelt
December 2025	Annual review

Keywords

This policy was approved by the FEP® Pharmacy and Medical Policy Committee on December 12, 2025 and is effective on January 1, 2026.

Appendix 1 - List of VEGF Inhibitors for Ocular Indications

Generic Name	Brand Name
aflibercept	Eylea/Eylea HD
bevacizumab	Avastin
brolucizumab-dbll	Beovu
faricimab-svoa	Vabysmo
ranibizumab	Lucentis
ranibizumab	Susvimo