

Clinical Policy: Aflibercept (Eylea)

Reference Number: LA.PHAR.184

Effective Date:

Last Review Date: 06.21

Line of Business: Medicaid

Coding Implications

Revision Log

See Important Reminder at the end of this policy for important regulatory and legal information.

Description

Aflibercept (Eylea[®]) is a vascular endothelial growth factor (VEGF) inhibitor.

FDA Approved Indication(s)

Eylea is 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)

Policy/Criteria

Prior authorization is required. 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 Louisiana Healthcare Connections that Eylea 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, b, c, or d):
 - a. Neovascular (wet) AMD;
 - b. Macular edema following RVO;
 - c. DME;
 - d. DR;
2. Prescribed by or in consultation with an ophthalmologist;
3. Age \geq 18 years;
4. For all indications, except for DME in members with baseline visual acuity worse than 20/50: Member must use 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. AMD: 2 mg (1 vial) every 4 weeks for the first 3 months, then every 8 weeks thereafter;
 - b. DME and DR: 2 mg (1 vial) every 4 weeks for the first 5 injections, then every 8 weeks thereafter;

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- c. **RVO: 2 mg (1 vial) every 4 weeks.**

Approval duration: 6 months

B. Other diagnoses/indications

- 1. **Refer to the off-label use policy if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): LA.PMN.53 for Medicaid.**

II. Continued Therapy

A. Ophthalmic Disease (must meet all):

- 1. **Currently receiving medication via Louisiana Healthcare Connections benefit or member has previously met initial approval criteria;**
- 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. **If request is for a dose increase, new dose does not exceed:**
 - a. **DME and DR: 2 mg (1 vial) every 8 weeks;**
 - b. **RVO: 2 mg (1 vial) every 4 weeks;**
 - c. **AMD: One of the following (i or ii):**
 - i. **Dose does not exceed 2 mg (1 vial) every 8 weeks;**
 - ii. **Member meets both of the following (a and b):**
 - a) **Documentation supports evidence of continued disease activity;**
 - b) **New dose does not exceed 2 mg (1 vial) every 4 weeks.**

Approval duration: 6 months

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

- 1. **Currently receiving medication via Louisiana Healthcare Connections benefit and documentation supports positive response to therapy.**
Approval duration: Duration of request or 6 months (whichever is less); or
- 2. **Refer to the off-label use policy if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): LA.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 policy –LA.PMN.53 for Medicaid or or evidence of coverage documents.**

IV. Appendices/General Information

Appendix A: Abbreviation/Acronym Key

AMD: age-related macular degeneration

DME: diabetic macular edema

DR: diabetic retinopathy

FDA: Food and Drug Administration

RVO: retinal vein occlusion

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 require prior authorization.

| <u>Drug Name</u> | <u>Dosing Regimen</u> | <u>Dose Limit/ Maximum Dose</u> |
|-----------------------------------|--|-------------------------------------|
| <u>Bevacizumab (Avastin®)</u> | <u>Neovascular (wet) AMD:</u> <u>1.25 to 2.5 mg administered by intravitreal injection every 4 weeks.</u> | <u>2.5 mg/month</u> |
| | <u>Macular edema secondary to RVO:</u> <u>1 mg to 2.5 mg administered by intravitreal injection every 4 weeks</u> | <u>2.5 mg/month</u> |
| | <u>DR:</u> <u>1.25 mg administered by intravitreal injection every 6 weeks</u> | <u>1.25 mg/6 weeks</u> |
| | <u>DME:</u> <u>1.25 mg administered by intravitreal injection every 6 weeks</u> | <u>1.25 mg/6 weeks</u> |

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 infection
 - Active intraocular inflammation
 - Hypersensitivity
- Boxed warning(s): none reported

Appendix D: General Information

- In the VEGF Trap-Eye: Investigation of Efficacy and Safety in Wet Age-Related Macular Degeneration (VIEW)-1 trial, the difference in the number of patients who lost fewer than 15 letters at 52 weeks between Eylea every 8 weeks compared to Lucentis was 0.6% (95.1% CI -0.32, 4.4). In terms of the number of patients who gained at least 15 letters, the mean difference between Eylea every 8 weeks was 6.6% (95.1% CI -1.0, 14.1). There were no adverse events that were found to be significant from the Lucentis arm.
- In a trial comparing Eylea, Avastin and Lucentis, the Diabetic Retinopathy Clinical Research Network found in patients with diabetic macular edema that when the initial visual-acuity letter score was 78 to 69 (equivalent to approximately 20/32 to 20/40) (51% of participants), the mean improvement was 8.0 with Eylea, 7.5 with Avastin, and 8.3 with Lucentis (p > 0.50 for each pair wise comparison). When the initial letter score was less than 69 (approximately 20/50 or worse), the mean

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improvement was 18.9 with Eylea, 11.8 with Avastin, and 14.2 with Lucentis (p < 0.001 for Eylea vs. Avastin, p = 0.003 for Eylea vs. Lucentis, and p = 0.21 for Lucentis vs. Avastin).

- In clinical trials for the treatment of AMD, DME, and DR, additional efficacy was not demonstrated in most patients when Eylea was dosed every 4 weeks as a maintenance dose, compared to every 8 weeks. Maintenance dosing at every 8 weeks should be attempted before increasing the intravitreal injection frequency to every 4 weeks.

V. Dosage and Administration

| Indication | Dosing Regimen | Maximum Dose |
|------------------------------------|---|-------------------|
| <u>AMD</u> | <u>2 mg (1 vial) administered by intravitreal injection once a month for 3 months then 2 mg every 2 months</u> <u>Although Eylea may be dosed as frequently as 2 mg every 4 weeks (monthly), additional efficacy was not demonstrated in most patients when Eylea was dosed every 4 weeks compared to every 8 weeks. Some patients may need every 4 week (monthly) dosing after the first 12 weeks (3 months).</u> | <u>2 mg/month</u> |
| <u>Macular edema following RVO</u> | <u>2 mg (1 vial) administered by intravitreal injection once every 4 weeks (monthly)</u> | <u>2 mg/month</u> |
| <u>DME, DR</u> | <u>2 mg (1 vial) administered by intravitreal injection once a month for the first 5 injections, followed by 2 mg via intravitreal injection once every 2 months</u> <u>Although Eylea may be dosed as frequently as 2 mg every 4 weeks (monthly), additional efficacy was not demonstrated in most patients when Eylea was dosed every 4 weeks compared to every 8 weeks. Some patients may need every 4 week (monthly) dosing after the first 20 weeks (5 months).</u> | <u>2 mg/month</u> |

VI. Product Availability

Single-dose vial and pre-filled syringe: 2 mg/0.05 mL solution

VII. References

1. Eylea Prescribing Information. Tarrytown, NY: Regeneron Pharmaceuticals, Inc.; August 2019. Available at: www.eylea.com. Accessed September 17, 2020.
2. American Academy of Ophthalmology Retina/Vitreous Panel. Preferred Practice Pattern® Guidelines. Age-Related Macular Degeneration. San Francisco, CA: American

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Academy of Ophthalmology; October 2019. Available at: www.aao.org/ppp. Accessed September 17, 2020.

3. **American Academy of Ophthalmology Retina/Vitreous Panel. Preferred Practice Pattern® Guidelines. Retinal Vein Occlusions. San Francisco, CA: American Academy of Ophthalmology; October 2019. Available at: www.aao.org/ppp. Accessed September 17, 2020.**
4. **American Academy of Ophthalmology Retina/Vitreous Panel. Preferred Practice Pattern® Guidelines. Diabetic Retinopathy. San Francisco, CA: American Academy of Ophthalmology; October 2019. Available at: www.aao.org/ppp. Accessed September 17, 2020.**
5. **Wells JA, Glassman AR, Ayala AR, et al. Aflibercept, bevacizumab, or ranibizumab for diabetic macular edema. N Engl J Med. 2015 Mar 26;372(13):1193-203. Doi: 10.1056/NEJMoa1414264.**

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.

| <u>HCPCS Codes</u> | <u>Description</u> |
|--------------------|-------------------------------------|
| J0178 | Injection, aflibercept, 1 mg |

| <u>Reviews, Revisions, and Approvals</u> | <u>Date</u> |
|--|----------------|
| Converted corporate to local policy | 06.2021 |
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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. LHCC 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

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policy is consistent with standards of medical practice current at the time that this clinical policy was approved.

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