# Clinical Policy: Estradiol Vaginal Ring (Femring)

Reference Number: CP.PMN.263
Effective Date: 01.01.22
Last Review Date: 08.24
Line of Business: Commercial, HIM, Medicaid

Revision Log

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

## Description

Estradiol vaginal ring (Femring®) is an estrogen steroid.

## FDA Approved Indication(s)

Femring is indicated for the treatment of moderate to severe vasomotor symptoms due to menopause and moderate to severe vulvar and vaginal atrophy, due to menopause.

## 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 Femring is **medically necessary** when the following criteria are met:

**I. Initial Approval Criteria**

**A. Dyspareunia** (must meet all)**:**

1. Diagnosis of dyspareunia (vulvar and vaginal atrophy) or vasomotor symptoms due to menopause;
2. Age ≥ 18 years;
3. Failure of two vaginal lubricants or vaginal moisturizers, unless contraindicated or clinically significant adverse effects are experienced (*see Appendix B*);
4. Failure of ≥ 4 week trial of one vaginal estrogen (e.g., estradiol vaginal cream (Estrace®), estradiol vaginal insert (Vagifem®), Premarin® vaginal cream), unless contraindicated or clinically significant adverse effects are experienced (*see Appendix B*);
5. Dose does not exceed one vaginal ring every 3 months.

**Approval duration:**
**Medicaid/HIM** – 12 months
**Commercial -** 12 months or duration of request, whichever is less

**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. Dyspareunia** (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 (e.g., dyspareunia symptom reduction);
3. If request is for a dose increase, new dose does not exceed one vaginal ring every 3 months.

**Approval duration:**
**Medicaid/HIM** – 12 months
**Commercial -** 12 months or duration of request, whichever is less

**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 policy – 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 Key*

FDA: Food and Drug Administration

*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 | Dosing Regimen | Dose Limit/ Maximum Dose |
|---|---|---|
| estradiol vaginal cream (Estrace®) | Initial: 2 to 4 gm vaginally QD for 1 to 2 weeks, gradually reduce to 50% of initial dose for 1 to 2 weeks<br>Maintenance: 1 gm 1 to 3 times a week | Varies |
| Premarin® (conjugated estrogens) vaginal cream | 0.5 gm intravaginally twice per week continuously | Varies |
| estradiol vaginal insert (Vagifem®) | 1 insert intravaginally daily for 2 weeks, followed by 1 insert twice weekly | 1 insert/day |
| Vaginal Lubricants:<br>*Water-based* Astroglide, FemGlide, Just Like Me, K-Y Jelly, Pre-Seed, Slippery Stuff, Summer's Eve<br>*Silicone-based* ID Millennium, Pink, Pjur, Pure Pleasure | Apply intravaginally before sex | Varies |
| Vaginal moisturizers:<br>Fresh Start, K-Y Silk-E, Moist Again, Replens, K-Y Liquibeads | Apply intravaginally before sex | Varies |

*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): undiagnosed abnormal genital bleeding; known, suspected, or history of breast cancer; known or suspected estrogen-dependent neoplasia; active DVT, PE, or history of these conditions; active arterial thromboembolic disease (for example, stroke and MI), or a history of these conditions; known anaphylactic reaction or

angioedema to Femring; known liver impairment or disease; known protein C, protein S, or antithrombin deficiency, or other known thrombophilic disorders; known or suspected pregnancy.
- Boxed warning(s): endometrial cancer, cardiovascular disorders, breast cancer, and probable dementia

## V. Dosage and Administration

| Drug Name | Dosing Regimen | Maximum Dose |
|---|---|---|
| Dyspareunia/ vasomotor symptoms due to menopause | Administer one vaginal ring every 3 months | 1 vaginal ring/3 months |

## VI. Product Availability

Vaginal ring: 12.4 mg, 24.8 mg

## VII. References

1. Femring Prescribing Information. Irvine, CA: Allergan USA, Inc., August 2018. Available at: www.femring.com. Accessed May 13, 2024.
2. American College of Obstetricians and Gynecologists Committee on Practice Bulletins-Gynecology. ACOG Practice Bulletin No. 213: Female sexual dysfunction. Obstet Gynecol. 2019;134(1):e1-e18.
3. Management of symptomatic vulvovaginal atrophy: 2013 position statement of The North American Menopause Society. Menopause. 2013;20:888-902.
4. The North American Menopause Society Recommendations for Clinical Care of Midlife Women, Chapter 3: Clinical Issues. Menopause: The Journal of The North American Menopause Society. 2014;21(10). Available at: https://www.menopause.org/docs/default-source/2014/nams-recomm-for-clinical-care. Accessed May 22, 2024.
5. Micromedex® Healthcare Series [Internet database]. Greenwood Village, Colo: Thomson Healthcare. Updated periodically. Accessed May 22, 2024.
6. The 2020 genitourinary syndrome of menopause position statement of The North American Menopause Society. Menopause: The Journal of The North American Menopause Society. 27 (9): 976-992.

| Reviews, Revisions, and Approvals | Date | P&T Approval Date |
|---|---|---|
| Policy created per March SDC and prior clinical guidance. | 03.26.21 | 11.21 |
| 3Q 2022 annual review: no significant changes; references reviewed and updated. | 05.04.22 | 08.22 |
| Template changes applied to other diagnoses/indications and continued therapy section. | 10.07.22 | |
| 3Q 2023 annual review: no significant changes; for commercial line of business, approval duration changed from “length of benefit” to | 04.11.23 | 08.23 |

| Reviews, Revisions, and Approvals | Date | P&T Approval Date |
|---|---|---|
| "12 months or duration of request, whichever is less"; references reviewed and updated. | | |
| 3Q 2024 annual review: no significant changes; references reviewed and updated. | 05.13.24 | 08.24 |

**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.