

## Medical Policy Manual **Approved Rev: Do Not Implement until 10/31/2026**

**Denosumab Products: Denosumab (Prolia®); Denosumab-nxxp (Bildyos®); Denosumab-mobz (Boncresa); Denosumab-kyqq (Bosaya™); Denosumab-bnht (Conexxence®); Denosumab-qbde (Enoby™); Denosumab-bbdz (Jubbonti®); Denosumab-dssb (Ospomyv™); Denosumab-desu (Osvyrti); **Denosumab-adet (Ponlimsi™); Denosumab-bmwo (Stoboclo®)****

### **IMPORTANT REMINDER**

We develop Medical Policies to provide guidance to Members and Providers. This Medical Policy relates only to the services or supplies described in it. The existence of a Medical Policy is not an authorization, certification, explanation of benefits or a contract for the service (or supply) that is referenced in the Medical Policy. For a determination of the benefits that a Member is entitled to receive under his or her health plan, the Member's health plan must be reviewed. If there is a conflict between the medical policy and a health plan or government program (e.g., TennCare), the express terms of the health plan or government program will govern.

### **POLICY**

#### **INDICATIONS**

The indications below including FDA-approved indications and compendial uses are considered a covered benefit provided that all the approval criteria are met and the member has no exclusions to the prescribed therapy.

#### FDA-Approved Indications

- Treatment of postmenopausal women with osteoporosis at high risk for fracture, defined as a history of osteoporotic fracture, or multiple risk factors for fracture; or patients who have failed or are intolerant to other available osteoporosis therapy.
- Treatment to increase bone mass in men with osteoporosis at high risk for fracture, defined as a history of osteoporotic fracture, or multiple risk factors for fracture; or patients who have failed or are intolerant to other available osteoporosis therapy.
- Treatment of glucocorticoid-induced osteoporosis in men and women at high risk of fracture who are either initiating or continuing systemic glucocorticoids in a daily dosage equivalent to 7.5 mg or greater of prednisone and expected to remain on glucocorticoids for at least 6 months. High risk of fracture is defined as a history of osteoporotic fracture, multiple risk factors for fracture, or patients who have failed or are intolerant to other available osteoporosis therapy.
- Treatment to increase bone mass in men at high risk for fracture receiving androgen deprivation therapy (ADT) for non-metastatic prostate cancer.
- Treatment to increase bone mass in women at high risk for fracture receiving adjuvant aromatase inhibitor therapy for breast cancer.

#### Compendial Uses

- For treatment-related bone loss in patients with prostate cancer receiving androgen deprivation therapy (ADT).
- Treatment in postmenopausal (natural or induced) patients with breast cancer receiving adjuvant aromatase inhibition therapy to maintain or improve bone mineral density and reduce risk of fractures.
- To inhibit progression of bone erosion in patients with rheumatoid arthritis.

All other indications are considered experimental/investigational and not medically necessary.

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### **DOCUMENTATION**

#### **Postmenopausal Osteoporosis, Osteoporosis in Men, Glucocorticoid-Induced Osteoporosis**

- Chart notes or medical record documentation indicating a history of fractures, T-score, and Fracture Risk Assessment Tool (FRAX) probability (if applicable).
- Chart notes, medical record documentation, or claims history supporting previous medications tried (if applicable), including response to therapy.

#### **Prostate Cancer**

Chart notes, medical record documentation, or claims history supporting use of androgen deprivation therapy (ADT).

#### **Breast Cancer**

Chart notes, medical record documentation, or claims history supporting use of aromatase inhibition therapy.

### **COVERAGE CRITERIA**

#### **Postmenopausal Osteoporosis**

Authorization of 12 months may be granted to postmenopausal members with osteoporosis when EITHER of the following criteria is met:

- Member has a history of fragility fractures (e.g., low trauma fracture from force similar to a fall from standing position).
- Member has a pre-treatment T-score  $\leq -2.5$  OR member has osteopenia (i.e., pre-treatment T-score between -1 and -2.5) with a high pre-treatment FRAX probability (see Appendix) and meets ANY of the following criteria:
  - Member has indicators of very high fracture risk (e.g., advanced age, frailty, glucocorticoid use, very low T-scores [ $\leq -3$ ], or increased fall risk)
  - Member has had an inadequate response or intolerance to previous injectable osteoporosis therapy (e.g., zoledronic acid [Reclast], teriparatide [Forteo], abaloparatide [Tymlos])
  - Member has had an inadequate response or intolerance to previous oral bisphosphonate therapy.

#### **Osteoporosis in Men**

Authorization of 12 months may be granted to male members with osteoporosis when any of the following criteria is met:

- Member has a history of an osteoporotic vertebral or hip fracture
- Member has a pre-treatment T-score  $\leq -2.5$  OR member has osteopenia (i.e., pre-treatment T-score between -1 and -2.5) with a high pre-treatment FRAX probability (see Appendix).
- Member has had an inadequate response or intolerance to previous bisphosphonate therapy.

#### **Glucocorticoid-Induced Osteoporosis**

Authorization of 12 months may be granted for members with glucocorticoid-induced osteoporosis when BOTH of the following criteria are met:

- Member is currently receiving or will be initiating glucocorticoid therapy at an equivalent prednisone dose of  $\geq 2.5$  mg/day for  $\geq 3$  months.

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- Member meets EITHER of the following criteria:
  - Member has had an inadequate response or intolerance to previous bisphosphonate therapy
  - Member meets ANY of the following criteria:
    - Member has a history of a fragility fracture (e.g., low trauma fracture from force similar to a fall from standing position).
    - Member has a pre-treatment T-score  $\leq -2.5$ .
    - Member has osteopenia (i.e., pre-treatment T-score between -1 and -2.5) with a high pre-treatment FRAX probability (see Appendix).

### **Prostate Cancer**

Authorization of 12 months may be granted to members who are receiving androgen deprivation therapy (ADT) for prostate cancer.

### **Breast Cancer**

Authorization of 12 months may be granted to members who are receiving adjuvant aromatase inhibition therapy for breast cancer.

### **Rheumatoid Arthritis**

Authorization of 12 months may be granted to inhibit progression of bone erosion in members with rheumatoid arthritis.

### **CONTINUATION OF THERAPY**

Authorization of 12 months may be granted for all members (including new members) who are currently receiving the requested medication through a previously authorized pharmacy or medical benefit, who meet either of the following:

- Member has received less than 24 months of therapy and has not experienced clinically significant adverse events during therapy
- Member has received 24 months of therapy or more and meets both of the following:
  - Member has experienced clinical benefit (i.e., improvement or stabilization in T-score since the previous bone mass measurement)
  - Member has not experienced any adverse effects

### **MEDICATION QUANTITY LIMITS**

| Drug Name                   | Diagnosis/Diagnoses                                               | Maximum Dosing Regimen                                       |
|-----------------------------|-------------------------------------------------------------------|--------------------------------------------------------------|
| Bildyos<br>(Denosumab-nxxp) | Breast Cancer                                                     | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Bildyos<br>(Denosumab-nxxp) | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced) | Route of Administration: Subcutaneous<br>60mg every 6 months |



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| Bildyos<br>(Denosumab-nxxp)   | Prostate Cancer                                                   | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Bildyos<br>(Denosumab-nxxp)   | Rheumatoid Arthritis                                              | Route of Administration: Subcutaneous<br>60mg every 3 months |
| Boncrea<br>(Denosumab-mobz)   | Breast Cancer                                                     | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Boncrea<br>(Denosumab-mobz)   | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced) | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Boncrea<br>(Denosumab-mobz)   | Prostate Cancer                                                   | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Boncrea<br>(Denosumab-mobz)   | Rheumatoid Arthritis                                              | Route of Administration: Subcutaneous<br>60mg every 3 months |
| Bosaya<br>(Denosumab-kyqq)    | Breast Cancer                                                     | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Bosaya<br>(Denosumab-kyqq)    | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced) | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Bosaya<br>(Denosumab-kyqq)    | Prostate Cancer                                                   | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Bosaya<br>(Denosumab-kyqq)    | Rheumatoid Arthritis                                              | Route of Administration: Subcutaneous<br>60mg every 3 months |
| Conexence<br>(Denosumab-bnht) | Breast Cancer                                                     | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Conexence<br>(Denosumab-bnht) | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced) | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Conexence<br>(Denosumab-bnht) | Prostate Cancer                                                   | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Conexence<br>(Denosumab-bnht) | Rheumatoid Arthritis                                              | Route of Administration: Subcutaneous<br>60mg every 3 months |



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| Enoby<br>(Denosumab-qbde)            | Breast Cancer                                                            | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| Enoby<br>(Denosumab-qbde)            | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced)        | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| Enoby<br>(Denosumab-qbde)            | Prostate Cancer                                                          | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| <b>Enoby<br/>(Denosumab-qbde)</b>    | <b>Rheumatoid Arthritis</b>                                              | <b>Route of Administration: Subcutaneous<br/>60mg every 3 months</b> |
| Jubbonti<br>(Denosumab-bbdz)         | Breast Cancer                                                            | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| Jubbonti<br>(Denosumab-bbdz)         | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced)        | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| Jubbonti<br>(Denosumab-bbdz)         | Prostate Cancer                                                          | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| <b>Jubbonti<br/>(Denosumab-bbdz)</b> | <b>Rheumatoid Arthritis</b>                                              | <b>Route of Administration: Subcutaneous<br/>60mg every 3 months</b> |
| Ospomyv<br>(Denosumab-dssb)          | Breast Cancer                                                            | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| Ospomyv<br>(Denosumab-dssb)          | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced)        | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| Ospomyv<br>(Denosumab-dssb)          | Prostate Cancer                                                          | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| <b>Ospomyv<br/>(Denosumab-dssb)</b>  | <b>Rheumatoid Arthritis</b>                                              | <b>Route of Administration: Subcutaneous<br/>60mg every 3 months</b> |
| Osvyrti<br>(Denosumab-desu)          | Breast Cancer                                                            | Route of Administration: Subcutaneous<br>60mg every 6 months         |
| <b>Osvyrti<br/>(Denosumab-desu)</b>  | <b>Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced)</b> | <b>Route of Administration: Subcutaneous<br/>60mg every 6 months</b> |



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| Osvyrti<br>(Denosumab-desu)  | Prostate Cancer                                                   | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Osvyrti<br>(Denosumab-desu)  | Rheumatoid Arthritis                                              | Route of Administration: Subcutaneous<br>60mg every 3 months |
| Ponlinsi<br>(Denosumab-adet) | Breast Cancer                                                     | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Ponlinsi<br>(Denosumab-adet) | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced) | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Ponlinsi<br>(Denosumab-adet) | Prostate Cancer                                                   | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Ponlinsi<br>(Denosumab-adet) | Rheumatoid Arthritis                                              | Route of Administration: Subcutaneous<br>60mg every 3 months |
| Prolia<br>(Denosumab)        | Breast Cancer                                                     | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Prolia<br>(Denosumab)        | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced) | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Prolia<br>(Denosumab)        | Prostate Cancer                                                   | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Prolia<br>(Denosumab)        | Rheumatoid Arthritis                                              | Route of Administration: Subcutaneous<br>60mg every 3 months |
| Stoboclo<br>(Denosumab-bmwo) | Breast Cancer                                                     | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Stoboclo<br>(Denosumab-bmwo) | Osteoporosis (Post-menopausal, in men, or glucocorticoid-induced) | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Stoboclo<br>(Denosumab-bmwo) | Prostate Cancer                                                   | Route of Administration: Subcutaneous<br>60mg every 6 months |
| Stoboclo<br>(Denosumab-bmwo) | Rheumatoid Arthritis                                              | Route of Administration: Subcutaneous<br>60mg every 3 months |

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### APPENDIX

#### FRAX® (Fracture Risk Assessment Tool)

- FRAX® (fracture risk assessment tool) available at: <https://fraxplus.org>
- High FRAX probability: 10-year major osteoporotic fracture probability  $\geq 20\%$  or hip fracture probability  $\geq 3\%$
- FRAX Glucocorticoid correction: If glucocorticoid dose is greater than 7.5 mg (prednisone equivalent) per day, the estimated risk score generated with FRAX should be multiplied by 1.15 for major osteoporotic fracture (including fractures of the spine [clinical], hip, wrist, or humerus) and 1.2 for hip fracture

#### APPLICABLE TENNESSEE STATE MANDATE REQUIREMENTS

BlueCross BlueShield of Tennessee's Medical Policy complies with Tennessee Code Annotated Section 56-7-2352 regarding coverage of off-label indications of Food and Drug Administration (FDA) approved drugs when the off-label use is recognized in one of the statutorily recognized standard reference compendia or in the published peer-reviewed medical literature.

#### ADDITIONAL INFORMATION

For appropriate chemotherapy regimens, dosage information, contraindications, precautions, warnings, and monitoring information, please refer to one of the standard reference compendia (e.g., the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines®) published by the National Comprehensive Cancer Network®, Drugdex Evaluations of Micromedex Solutions at Truven Health, or The American Hospital Formulary Service Drug Information).

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**EFFECTIVE DATE** 10/31/2026

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