Prior Authorization Review Panel

CHC-MCO Policy Submission

A separate copy of this form must accompany each policy submitted for review.
Policies submitted without this form will not be considered for review.

Plan: PA Health & Wellness	**Submission Date: 05/01/2022**
Policy Number: PA.CP.PHAR.418	**Effective Date: 01/2020** **Revision Date: 04/2022**
Policy Name: Dexrazoxane (Zinecard, Totect)	

Type of Submission – Check all that apply:

- ☐ **New Policy**
- ✓ **Revised Policy***
- ☐ **Annual Review - No Revisions**
- ☐ **Statewide PDL -** *Select this box when submitting policies for Statewide PDL implementation and when submitting policies for drug classes included on the Statewide PDL.*

***All revisions to the policy must be highlighted using track changes throughout the document.**

Please provide any changes or clarifying information for the policy below:

2Q 2022 annual review: per NCCN added off-label supported uses in patients under 18 years of age in Ph-negative ALL, aggressive mature B-cell lymphomas, Hodgkin lymphoma, or Wilms Tumor (nephroblastoma); removed appendix D that provided references to studies with inconclusive doxorubicin thresholds for use in pediatric patients as such use is supported by NCCN; references reviewed and updated.

Name of Authorized Individual (Please type or print): **Venkateswara R. Davuluri, MD**	**Signature of Authorized Individual:**

Clinical Policy: Dexrazoxane (Zinecard, Totect)

Reference Number: PA.CP.PHAR.418
Effective Date: 01/2020
Last Review Date: 04/2022

Coding Implications
Revision Log

Description
Dexrazoxane (Zinecard®, Totect®) is a cytoprotective agent.

FDA Approved Indications
Totect and Zinecard are indicated for reducing the incidence and severity of cardiomyopathy associated with doxorubicin in women with metastatic breast cancer who have received a cumulative doxorubicin dose of 300 mg/m^2 and will continue receiving doxorubicin to maintain tumor control.

Totect is indicated for the treatment of extravasation resulting from intravenous anthracycline chemotherapy.

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 PA Health & Wellness® that Zinecard and Totect are **medically necessary** when the following criteria are met:

I. Initial Approval Criteria

A. Doxorubicin-Induced Cardiomyopathy (must meet all)**:**

1. Prescribed to reduce the incidence or severity of cardiomyopathy associated with doxorubicin;
2. Prescribed by or in consultation with an oncologist or hematologist;
3. One of the following:
    a. Age ≥ 18 years and member has received a cumulative doxorubicin dose of ≥ 300 mg/m^2;
    b. Request is for Ph-negative ALL as part of the DFCI ALL Protocol 11-001 or 16-001 in members with an anticipated cumulative anthracycline dose ≥ 250 mg/m^2 of doxorubicin equivalent or radiation with potential impact to the heart (e.g., radiation to chest, abdomen, spine, or total body irradiation) (off-label);
    c. Request is for pediatric aggressive mature B-cell lymphomas or Hodgkin lymphoma (off-label);
    d. Request is for Wilms Tumor (nephroblastoma) and member has a planned cumulative dose of doxorubicin ≥ 150 mg/m^2 (off-label);
4. Will be used concurrently with doxorubicin;
5. Request meets one of the following (a or b):
    a. Dose does not exceed 10 times the dose of doxorubicin (e.g., dexrazoxane 500 mg/m^2 for member receiving doxorubicin 50 mg/m^2) given with each doxorubicin dose;

b. Dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (*prescriber must submit supporting evidence*).

Approval duration: 12 months or duration of doxorubicin therapy, whichever is less

B. Anthracycline-Induced Extravasation (must meet all)**:**

1. Diagnosis of anthracycline-induced extravasation;
2. Prescribed by or in consultation with an oncologist or hematologist;
3. Age ≥ 18 years;
4. Request meets one of the following (a or b):
   a. Dose does not exceed 2,000 mg per day on days 1 and 2, and 1,000 mg on day 3.
   b. Dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (*prescriber must submit supporting evidence*).

Approval duration: 3 days

C. 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): PA.CP.PMN.53

II. Continued Therapy

A. Doxorubicin-Induced Cardiomyopathy (must meet all)**:**

1. Currently receiving medication via PA Health & Wellness benefit and documentation supports positive response to therapy or the Continuity of Care policy (PA.LTSS.PHAR.01) applies;
2. Member continues to receive doxorubicin;
3. Member is responding positively to therapy;
4. Request meets one of the following (a or b):
   a. Dose does not exceed 10 times the dose of doxorubicin (e.g., dexrazoxane 500 mg/m^2 for member receiving doxorubicin 50 mg/m^2) given with each doxorubicin dose;
   b. Dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (*prescriber must submit supporting evidence*).

Approval duration: 12 months or duration of doxorubicin therapy, whichever is less

B. Anthracycline-Induced Extravasation

1. Re-authorization is not permitted. Member must meet the initial approval criteria.

Approval duration: Not applicable

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

1. Currently receiving medication via PA Health & Wellness benefit and documentation supports positive response to therapy or the Continuity of Care policy (PA.LTSS.PHAR.01) applies.
   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): PA.CP.PMN.53

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 policies – PA.CP.PMN.53.

IV. Appendices/General Information

Appendix A: Abbreviation/Acronym Key
ALL: acute lymphoblastic leukemia
FDA: Food and Drug Administration

Appendix B: Therapeutic Alternatives
Not applicable

Appendix C: Contraindications/Boxed Warnings
- Contraindication(s):
  - Zinecard: should not be used with non-anthracycline chemotherapy regimens
  - Totect: none reported
- Boxed warning(s): none reported

V. Dosage and Administration

Drug Name	Indication	Dosing Regimen	Maximum Dose
dexrazoxane (Totect, Zinecard)	Doxorubicin-induced cardiomyopathy	Give dexrazoxane at a ratio of 10:1 with the doxorubicin dose as an IV infusion over 15 minutes and within 30 minutes before doxorubicin is given.	Not applicable
dexrazoxane (Totect)	Anthracycline-induced extravasation	Day 1: 1,000 mg/m^2 Day 2: 1,000 mg/m^2 Day 3: 500 mg/m^2 Give Totect as an IV infusion over 1-2 hours and within 6 hours of extravasation. Treatment on days 2 and 3 should start at the same hour (+/- 3 hours) as day 1.	Day 1: 2,000 mg Day 2: 2,000 mg Day 3: 1,000 mg

VI. Product Availability

Drug Name	Availability
dexrazoxane (Zinecard)	Single-dose vial, IV powder for solution: 250 mg, 500 mg
dexrazoxane (Totect)	Single-dose vial, IV powder for solution: 500 mg

VII. References

1. Zinecard Prescribing Information. New York, NY: Pharmacia & Upjohn Co; October 2016. Available at: http://labeling.pfizer.com/ShowLabeling.aspx?format=PDF&id=514 . Accessed February 14,2022.

2. Totect Prescribing Information. Yardville, PA: Clinigen, Inc; November 2020. Available at: https://dailymed.nlm.nih.gov/dailymed/drugInfo.cfm?setid=b098c7f5-07a9-49a3-8e8a-f8adda671eca&audience=consumer. Accessed February 14,2022.
3. American Society of Clinical Oncology 2008 Clinical Practice Guideline Update: Use of Chemotherapy and Radiation Therapy Protectants. Available at: http://ascopubs.org/doi/pdf/10.1200/JCO.2008.17.2627. J Clin Oncol; 27:127-145.
4. Choi HS, Park ES, Kang HJ, et al. Dexrazoxane for preventing anthracycline cardiotoxicity in children with solid tumors. J Korean Med Sci. 2010;25(9):1336-42.
5. Asselin BL, Devidas M, Chen L, et al. Cardioprotection and Safety of Dexrazoxane in Patients Treated for Newly Diagnosed T-Cell Acute Lymphoblastic Leukemia or Advanced-Stage Lymphoblastic Non-Hodgkin Lymphoma: A Report of the Children's Oncology Group Randomized Trial Pediatric Oncology Group 9404. J Clin Oncol. 2016;34(8):854-62.
6. Dexrazoxane. In: National Comprehensive Cancer Network Drugs and Biologics Compendium. Available at: https://nccn.org/. Accessed February 14, 2022.
7. National Comprehensive Cancer Network. Pediatric Aggressive Mature B-Cell Lymphomas Version 2.2021. Available at: https://www.nccn.org/professionals/physician_gls/pdf/ped_b-cell.pdf. Accessed February 14, 2022.
8. National Comprehensive Cancer Network. Pediatric Acute Lymphoblastic Leukemia Version 1.2022. Available at: https://www.nccn.org/professionals/physician_gls/pdf/ped_all.pdf. Accessed February 14, 2022.
9. National Comprehensive Cancer Network. Pediatric Hodgkin Lymphoma Version 3.2021. Available at: https://www.nccn.org/professionals/physician_gls/pdf/ped_hodgkin.pdf. Accessed February 14, 2022.
10. National Comprehensive Cancer Network. Wilms Tumor (Nephroblastoma) Version 2.2021. Available at: https://www.nccn.org/professionals/physician_gls/pdf/wilms_tumor.pdf. Accessed February 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
J1190	Injection, dexrazoxane, 250 mg

Reviews, Revisions, and Approvals	Date	P&T Approval Date
Policy created	01/2020	

Reviews, Revisions, and Approvals	Date	P&T Approval Date
1Q 2021 annual review: updated policy in response to expansion of Totect FDA indication for doxorubicin toxicity, which overlaps with Zinecard; references reviewed and updated	01/2021	
2Q 2021 annual review: updated section V dosing to include Totect for the indication of doxorubicin-induced cardiomyopathy; references reviewed and updated.	04/2021	
2Q 2022 annual review: per NCCN added off-label supported uses in patients under 18 years of age in Ph-negative ALL, aggressive mature B-cell lymphomas, Hodgkin lymphoma, or Wilms Tumor (nephroblastoma); removed appendix D that provided references to studies with inconclusive doxorubicin thresholds for use in pediatric patients as such use is supported by NCCN; references reviewed and updated.	04/2022	