Clinical Policy: Tisotumab Vedotin-tftv (Tivdak)

Reference Number: CP.PHAR.561
Effective Date: 12.01.21
Last Review Date: 11.25
Line of Business: Commercial, HIM, Medicaid

Coding Implications
Revision Log

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

Description

Tisotumab vedotin-tftv (Tivdak®) is a tissue factor-directed antibody and microtubule inhibitor conjugate.

FDA Approved Indication(s)

Tivdak is indicated for the treatment of adult patients with recurrent or metastatic cervical cancer with disease progression on or after 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 Centene Corporation® that Tivdak is **medically necessary** when the following criteria are met:

I. Initial Approval Criteria

A. Cervical Cancer, Vaginal Cancer (off-label) (must meet all)**:**

1. Diagnosis of cervical cancer or vaginal cancer;
2. Prescribed by or in consultation with an oncologist;
3. Age ≥ 18 years;
4. Disease is recurrent or metastatic;
5. Disease has progressed on or after prior chemotherapy (*see Appendix B for examples*);
6. Prescribed in one of the following ways (a or b):
   a. As a single agent;
   b. In combination with Keytruda for cervical cancer that is programmed death-ligand 1 (PD-L1) positive (combined positive score [CPS] ≥ 1);
7. Documentation of member's current weight in kg;
8. Request meets one of the following (a or b):*
   a. Dose does not exceed 2 mg/kg (up to a maximum dose of 200 mg for members ≥ 100 kg) every 3 weeks;
   b. Dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (*prescriber must submit supporting evidence*).

**Prescribed regimen must be FDA-approved or recommended by NCCN*

Approval duration:
Medicaid/HIM – 12 months
Commercial – 6 months or to the member's renewal date, whichever is longer

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. All Indications in Section I (must meet all)**:**
1. Currently receiving medication via Centene benefit, or documentation supports that member is currently receiving Tivdak for a covered indication and has received this medication for at least 30 days;
2. Member is responding positively to therapy;
3. Prescribed as a single agent;
4. Documentation of member's current weight in kg;
5. Dose is at least 0.9 mg/kg every 3 weeks;
6. If request is for a dose increase, request meets one of the following (a or b):*
   a. New dose does not exceed 2 mg/kg (up to a maximum dose of 200 mg for patients ≥ 100 kg) every 3 weeks;
   b. New dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (*prescriber must submit supporting evidence*).

   **Prescribed regimen must be FDA-approved or recommended by NCCN*

Approval duration:
Medicaid/HIM – 12 months
Commercial – 6 months or to the member's renewal date, whichever is longer

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 policies – 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/Acronym 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 Name	Dosing Regimen	Dose Limit/ Maximum Dose
paclitaxel/cisplatin ± bevacizumab (Avastin®, Mvasi®, Zirabev™)	• Paclitaxel: 135 mg/m2 or 175 mg/m2 IV on Day 1 • Cisplatin: 50 mg/m^2 IV on Day 1 or 2 • With or without bevacizumab: 15 mg/kg IV on day Repeat every 3 weeks until disease progression or unacceptable toxicity	Varies
paclitaxel/carboplatin ± bevacizumab (Avastin®, Mvasi®, Zirabev™)	• Paclitaxel 135 mg/m^2 IV over 3 hours • Carboplatin target AUC 5 IV • With or without bevacizumab: 15 mg/kg IV on day Repeat every 3 weeks until disease progression or unacceptable toxicity	Varies
topotecan (Hycamtin®) /paclitaxel ± bevacizumab (Avastin®, Mvasi®, Zirabev™)	• Paclitaxel: 175 mg/m^2 on day 1 • Topotecan: 0.75 mg/m^2 on days 1, 2, and 3 • With or without bevacizumab: 15 mg/kg IV on day	Varies

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
	Repeat every 3 weeks until disease progression or unacceptable toxicity	
paclitaxel/cisplatin	• Paclitaxel: 135 mg/m^2 over 24 hours • Cisplatin: 50 mg/m^2 on day 1 Repeat every 3 weeks for a maximum of 6 cycles in non-responders or until disease progression or unacceptable toxicity	Varies
paclitaxel/carboplatin	• Paclitaxel 135 mg/m^2 IV over 3 hours on day 1 until disease progression or unacceptable toxicity • Carboplatin: Target AUC 5 IV every 3 weeks for 6 to 9 cycles	Varies
cisplatin/topotecan (Hycamtin®)	• Cisplatin: 50 mg/m^2 IV on day 1 • Topotecan: 0.75 mg/m^2/day IV for days 1, 2, and 3 Repeat every 3 weeks for a maximum of 6 cycles in nonresponders or until disease progression or unacceptable toxicity	Varies
paclitaxel/topotecan (Hycamtin®)	• Paclitaxel: 175 mg/m^2 on day 1 • Topotecan: 0.75 mg/m^2 on days 1, 2, and 3 Repeat every 3 weeks until disease progression or unacceptable toxicity	Varies
Keytruda® (pembrolizumab) + paclitaxel/cisplatin ± bevacizumab (Avastin®, Mvasi®, Zirabev™) for PD-L1-positive tumors	Varies	Varies
cisplatin	40 mg/m^2 over 4 hours to radiation therapy on days 1, 8, 15, 22, 29, and 36	Varies
carboplatin	400 mg/m^2 on day 1 every 28 days	Varies
paclitaxel	Varies	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): none reported
- Boxed warning(s): ocular toxicity

V. Dosage and Administration

Indication	Dosing Regimen	Maximum Dose
Cervical cancer	2 mg/kg IV over 30 minutes every 3 weeks until disease progression or unacceptable toxicity	2 mg/kg, 200 mg for members ≥ 100 kg

VI. Product Availability

Intravenous powder for solution, single-dose vial: 40 mg

VII. References

1. Tivdak Prescribing Information. Bothell, WA: Seagen Inc.; April 2024. Available at: https://www.tivdakhcp.com. Accessed July 14, 2025.
2. National Comprehensive Cancer Network Drugs and Biologics Compendium. Available at: http://www.nccn.org/professionals/drug_compendium. Accessed July 14, 2025.
3. National Comprehensive Cancer Network. Cervical Cancer Version 4.2025. Available at: https://www.nccn.org/professionals/physician_gls/pdf/cervical.pdf. Accessed July 14, 2025.
4. National Comprehensive Cancer Network. Vaginal Cancer Version 5.2025. Available at: https://www.nccn.org/professionals/physician_gls/pdf/vaginal.pdf. Accessed July 14, 2025.

Coding Implications

HCPCS Codes	Description
J9273	Injection, tisotumab vedotin-tftv, 1 mg

Reviews, Revisions, and Approvals	Date	P&T Approval Date
Policy created	10.18.21	11.21
4Q 2022 annual review: added criterion for single-agent therapy per NCCN; clarified previous failure criterion to remove the "with or without bevacizumab" as the example regimens are listed in Appendix B; removed the no more than two prior systemic regimens in the recurrent or metastatic setting criterion as neither NCCN nor the PI restrict previous number of therapies; updated HCPCS code; references reviewed and updated. Template changes applied to other diagnoses/indications.	08.04.22	11.22
4Q 2023 annual review: no significant changes; references reviewed and updated.	08.10.23	11.23
RT4: converted FDA approved indication for cervical cancer from accelerated approval to full approval per PI; added off-label vaginal cancer indication per NCCN; for continued therapy, updated approval duration for Commercial line of business to standard language of "6 months or to the member's renewal date, whichever is longer"; references reviewed and updated.	05.08.24	
4Q 2024 annual review: no significant changes; added Section III, Diagnoses/Indications for which coverage is NOT authorized per current template; references reviewed and updated.	08.08.24	11.24

Reviews, Revisions, and Approvals	Date	P&T Approval Date
4Q 2025 annual review: added allowance for combination use with Keytruda for PD-L1 positive cervical cancer per NCCN; extended initial approval duration for HIM/Medicaid from 6 to 12 months; references reviewed and updated.	07.14.25	11.25

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.