

## Clinical Policy: Mitomycin ~~for Pyelocalyceal~~ Instillation Solution (Jelmyto, Zusduri)

Reference Number: LA.PHAR.495

Effective Date: 06.20.24

Last Review Date: ~~08.20.24~~ 11.20.24

Line of Business: Medicaid

[Coding Implications](#)

[Revision Log](#)

See [Important Reminder](#) at the end of this policy for important regulatory and legal information.

**\*\*Please note: This policy is for medical benefit\*\***

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### Description

Mitomycin (for pyelocalyceal solution ~~(Jelmyto<sup>®</sup>)~~ and for intravesical solution (Zusduri<sup>™</sup>)) is an alkylating drug.

### FDA Approved Indication(s)

Jelmyto is indicated for the treatment of adult patients with low-grade upper tract urothelial cancer (LG-UTUC).

Zusduri is indicated for the treatment of adult patients with recurrent low-grade intermediate-risk non-muscle invasive bladder cancer (LG-IR-NMIBC).

### 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 Louisiana Healthcare Connections<sup>®</sup> that Jelmyto ~~is~~ and Zusduri are medically necessary when the following criteria are met:

#### I. Initial Approval Criteria

##### A. Low-Grade Upper Tract Urothelial Cancer (must meet all):

1. Request is for Jelmyto;
- ~~1. Newly diagnosed or recurrent LG-UTUC that is both (a and b):~~
  - ~~a. Above the ureteropelvic junction;~~
- ~~b. 2. Nonnon~~-metastatic;
- ~~2. 3.~~ Prescribed by or in consultation with an oncologist or urologist;
- ~~3. 4.~~ Age ≥ 18 years;
- ~~4. 5.~~ Lesion(s) measure ≤ 15 mm;
- ~~5. For the affected kidney(s), member does not have a recent history (within the last year) of carcinoma in situ in the urinary tract, invasive urothelial carcinoma, or high-grade papillary urothelial carcinoma;~~
6. Member is not a candidate for or is not seeking nephroureterectomy as definitive treatment;
7. One of the following (a or b):
  - a. Member has had complete or near complete endoscopic resection or ablation;

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- b. Member is not a candidate for endoscopic/surgical intervention;

#### 8. Prescribed as monotherapy:

#### 8-9. Request meets one of the following (a or b):\*

- a. Dose does not exceed 60 mg once weekly for 6 instillations per kidney;
- 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: 3 months (6 instillations per kidney)**

#### **B. Non-Muscle Invasive Bladder Cancer (must meet all):**

1. Request is for Zuesduri;
2. Diagnosis of NMIBC characterized as both of the following (a and b):
  - a. Ta low-grade;
  - b. Intermediate-risk (see Appendix D);
3. Prescribed by or in consultation with an oncologist or urologist;
4. Age > 18 years;
5. Member has previously undergone transurethral resection of bladder tumor (TURBT);
6. Request meets one of the following (a or b):\*
  - a. Dose does not exceed 75 mg once weekly for 6 instillations into the bladder;
  - 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: 3 months (up to 6 total intravesical instillations)**

#### **B-C. 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 LA.PMN.255
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: LA.PMN.53 for Medicaid.~~

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## II. Continued Therapy

#### **A. Low-Grade Upper Tract Urothelial Cancer All Indications in Section I** (must meet all):

1. Currently receiving medication via Louisiana Healthcare Connections benefit, or documentation supports that member is currently receiving ~~Jelmyto~~ mitomycin instillation solution for a covered indication and has received this medication for at least 30 days;

#### 2. For Jelmyto, both of the following (a and b):

- 2-a. If member has received 6 instillations, complete response (CR) has been achieved at 3 months after initiation of therapy as evidenced by complete absence of tumor lesions on urine cytology and ureteroscopy;

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- ~~3.b.~~ Member has not received more than 17 instillations;
3. For Zuspuri, member has not received  $\geq 6$  instillations;
4. If request is for a dose increase, request meets one of the following (a, b, or c):\*
  - a. For Jelmyto requests, one of the following (i or ii):
    - ~~a.i.~~ If member has completed  $< 6$  weekly instillations: New dose does not exceed 60 mg once weekly for up to 6 instillations per kidney;
    - ~~b.ii.~~ If member has completed  $\geq 6$  weekly instillations: New dose does not exceed 60 mg once monthly for up to 11 instillations per kidney;
  - b. For Zuspuri requests: New dose does not exceed 75 mg once weekly for 6 instillations into the bladder;
  - c. 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~~

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

#### Approval duration:

Jelmyto – 12 months (up to 17 total instillations per kidney)

Zuspuri – 3 months (up to 6 total intravesical instillations per lifetime)

#### 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 LA.PMN.255
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: 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 ~~ies – LA.PMN.53 for Medicaid, or evidence of coverage documents.~~

### IV. Appendices/General Information

#### Appendix A: Abbreviation/Acronym Key

FDA: Food and Drug Administration

LG-UTUC: low-grade upper tract

urothelial cancer

NCCN: National Comprehensive Cancer Network

NMIBC: non-muscle invasive bladder cancer

TURBT: transurethral resection of bladder tumor

#### Appendix B: Therapeutic Alternatives

Not applicable

#### Appendix C: Contraindications/Boxed Warnings

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- Contraindication(s): perforation of the bladder or upper urinary tract
  - Jelmyto: perforation of the bladder or upper urinary tract
  - Zusduri: perforation of the bladder, prior hypersensitivity reaction to mitomycin or any component of the product
- Boxed warning(s): none reported

#### Appendix D: General Information

- NCCN Compendium currently recommend Jelmyto with a Category 2A recommendation for primary treatment for a non-metastatic, residual, low-grade, low volume (5-15 mm), solitary tumor in the upper urinary tract for a patient who is not a candidate for or not seeking nephroureterectomy as a definitive treatment. Complete or near complete endoscopic resection or ablation is recommended prior to mitomycin ureteral gel application.
- Intermediate-risk NMIBC is defined as having one or two of the following: the presence of multiple tumors, a solitary tumor > 3 cm, and/or early or frequent recurrence (≥ 1 occurrence of low-grade NMIBC within 1 year of current diagnosis).

## V. Dosage and Administration

Drug Name	Indication	Dosing Regimen	Maximum Dose
<u>Mitomycin for pyelocalyceal solution (Jelmyto)</u>	LG-UTUC	<p>Jelmyto is for pyelocalyceal use only and not for intravenous use, topical use, or oral administration.</p> <p>The dose of Jelmyto to be instilled is 4 mg/mL via ureteral catheter or nephrostomy tube, with total instillation volume based on volumetric measurements using pyelography, not to exceed 15 mL (60 mg of mitomycin).</p> <p>Instill Jelmyto once weekly for six weeks. For patients with a complete response 3 months after Jelmyto initiation, Jelmyto instillations may be administered once a month for a maximum of 11 additional instillations.</p>	60 mg/ <u>instillation</u> ; 17 instillations
<u>Mitomycin for intravesical solution (Zusduri)</u>	NMIBC	<u>75 mg (56 mL) instilled once weekly for six weeks into the bladder via a urinary catheter</u>	<u>75 mg/instillation; 6 instillations</u>

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#### VI. Product Availability

~~For pyelocalyceal solution — carton containing the following:~~

- ~~• Two 40 mg (each) single-dose vials of mitomycin for pyelocalyceal solution~~
- ~~• One vial of 20 mL sterile hydrogel for reconstitution~~

Drug Name	Availability
Mitomycin for pyelocalyceal solution (Jelmyto)	Carton containing the following: <ul style="list-style-type: none"> <li>• Two 40 mg (each) single-dose vials of mitomycin for pyelocalyceal solution</li> <li>• One vial of 20 mL sterile hydrogel for reconstitution</li> </ul>
Mitomycin for intravesical solution (Zusduri)	Kit containing the following: <ul style="list-style-type: none"> <li>• Two 40 mg (each) single-dose vials of mitomycin for intravesical solution</li> <li>• One vial of 60 mL sterile hydrogel for reconstitution</li> </ul>

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#### VII. References

1. Jelmyto Prescribing Information. Princeton, NJ: UroGen Pharma, Inc.; ~~September 2022~~October 2024. Available at <https://www.jelmyto.com/>. Accessed ~~May 6, 2024~~April 9, 2025.
- ~~1-2~~ 2. Zusduri Prescribing Information. Princeton, NJ: UroGen Pharma; ~~June 2025~~. Available at: [https://www.urogen.com/download/pdf/zusduri\\_prescribing.pdf](https://www.urogen.com/download/pdf/zusduri_prescribing.pdf). Accessed ~~June 25, 2025~~.
- ~~2-3~~ 3. Kleinmann N, Matin S, Pierorazio P, et al. Primary chemoablation of low-grade upper tract urothelial carcinoma using UGN-101, a mitomycin-containing reverse thermal gel (OLYMPUS): an open-label, single-arm, phase 3 trial. Lancet Oncol 2020. Published online April 29, 2020. Available at [https://doi.org/10.1016/S1470-2045\(20\)30147-9](https://doi.org/10.1016/S1470-2045(20)30147-9).
4. Prasad SM, Shishkov D, Mihaylov NV, et al. Primary chemoablation of recurrent low-grade intermediate-risk nonmuscle-invasive bladder cancer with UGN-102: A single-arm, open-label, phase 3 trial (ENVISION). J Urol. 2025;213(2):205-216.
- ~~3-5~~ 5. National Comprehensive Cancer Network Drugs and Biologics Compendium. Available at: [http://www.nccn.org/professionals/drug\\_compendium](http://www.nccn.org/professionals/drug_compendium). Accessed May ~~22, 2024~~16, 2025.
- ~~4-6~~ 6. National Comprehensive Cancer Network. Bladder Cancer Version ~~4.2024~~1, 2025. Available at [https://www.nccn.org/professionals/physician\\_gls/pdf/bladder.pdf](https://www.nccn.org/professionals/physician_gls/pdf/bladder.pdf). Accessed ~~May 22, 2024~~June 25, 2025.

#### 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.

HCPSC Codes	Description
J9281	Mitomycin pyelocalyceal instillation, 1 mg
J9999	<u>Not otherwise classified, antineoplastic drugs (Zusduri)</u>
C9399	<u>Unclassified drugs or biologicals (Zusduri)</u>

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Reviews, Revisions, and Approvals	Date	LDH Approval Date
Policy created	05.01.23	08.28.23
Annual review: added criteria that LG-UTUC be non-metastatic; added requirement for endoscopic resection or ablation if member is a candidate per NCCN; references reviewed and updated.	03.15.24	06.20.24
No significant changes; references reviewed and updated.	11.20.24	<u>01.27.25</u>
<u>Annual review: removed requirement for cancer location above the ureteropelvic junction per NCCN; removed exclusion for “recent history of carcinoma in situ in the urinary tract, invasive urothelial carcinoma, or high-grade papillary urothelial carcinoma” as this is not excluded per NCCN or the FDA indication; added requirement for use as monotherapy per NCCN; added criteria for newly approved Zsduri; policy renamed to “Mitomycin Instillation Solution.”; references reviewed and updated</u>	<u>08.20.25</u>	

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

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 LHCC administrative policies and procedures.

This clinical policy is effective as of the date determined by LHCC. 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. LHCC 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.

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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.

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