

| *National Imaging Associates, Inc. | |
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| Clinical guideline | Original Date: June 2013 |
| CERVICAL CANCER | |
| Radiation Oncology | Last Revised Date: May 2023 |
| Guideline Number: NIA_CG_127 | Implementation Date: January 2024 |

GENERAL INFORMATION

- It is an expectation that all patients receive care/services from a licensed clinician. All appropriate supporting documentation, including recent pertinent office visit notes, laboratory data, and results of any special testing must be provided. If applicable: All prior relevant imaging results and the reason that alternative imaging cannot be performed must be included in the documentation submitted.
- Where a specific clinical indication is not directly addressed in this guideline, medical necessity determination will be made based on widely accepted standard of care criteria. These criteria are supported by evidence-based or peer-reviewed sources such as medical literature, societal guidelines and state/national recommendations.

INDICATIONS FOR RADIATION THERAPY AND TREATMENT OPTIONS

Definitive/Preoperative Radiation Therapy¹

- Stage IA-IA2– Brachytherapy (LDR or HDR) +/- 2D/3D-CRT (40-50 Gy; 28 fx max)
- Stage IB1 Pelvic 2D/3D-CRT (40-50 Gy; 28 fx max) + brachytherapy boost
- Stage IB2-IIIA Pelvic radiation therapy 2D/3D-CRT (40-50 Gy; 28 fx max) + brachytherapy boost) and concomitant chemotherapy +/- adjuvant hysterectomy
- Stage IIB-IVA Pelvic and/or para-aortic 2D/3D-CRT + brachytherapy + concurrent chemotherapy
- Stage IVB 2D/3D-CRT +/- brachytherapy for palliation only (symptom control)

Grossly involved unresected nodes may be evaluated for boosting with an additional 10-15 Gy. IMRT is approvable when the para-aortic nodes are being treated.

Post-operative (Adjuvant) Radiation Therapy¹

- Patients found to have deep cervical stromal invasion, lymphovascular invasion and/or bulky primary tumors
 - Pelvic 2D/3D-CRT/IMRT (45-50 Gy; 28 fx max) +/-concurrent chemotherapy

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- Patients with positive nodes, positive margins and/or parametrial invasion -
 - Pelvic 2D/3D-CRT/IMRT (45-50 Gy; 28 fx max) + concurrent chemotherapy
 - Pelvic 2D/3D-CRT/IMRT (45-50 Gy; 28 fx max) +/- vaginal brachytherapy boost (LDR or HDR) can be considered in women with a positive margin

Grossly involved unresected nodes may be evaluated for boosting with an additional 10-15 Gy. Unless otherwise indicated, standard radiation fractionation consists of 1.8 Gy to 2.0 Gy per day.

Local /Regional Recurrence¹

- No previous RT or outside previous RT fields
 - 2D/3D-CRT + chemotherapy +/- brachytherapy
- Previous RT
 - Intraoperative Radiation Therapy (IORT) for centralized disease
 - Possible Brachytherapy (LDR or HDR) for centralized disease < 2cm Tumor directed 2D/3D-CRT +/- chemotherapy if noncentral disease
 - o External Beam Radiation Therapy

TREATMENT OPTIONS REQUIRING ADDITIONAL CLINICAL REVIEW¹:

Intensity modulated radiation therapy (IMRT)

IMRT is not indicated as a standard treatment option and should not be used routinely for the delivery of radiation therapy for cervical cancer. IMRT is strictly defined by the utilization of inverse planning modulation techniques. IMRT may be appropriate for circumstances in which radiation therapy is indicated and

- Non-IMRT techniques cannot adequately deliver the radiation prescription without exceeding normal tissue radiation tolerance. The non-IMRT delivery is anticipated to contribute to potential late toxicity
- Tumor volume dose heterogeneity from non-IMRT techniques is such that unacceptable hot or cold spots are created

Requests for IMRT treatment delivery to the cervix will be reviewed for medical necessity prior to authorization based on the above criteria. Clinical rationale and documentation for performing IMRT rather than non-IMRT techniques must be provided for review. This includes a statement of medical necessity from the requesting provider and a dosimetric comparison plan addressing the approval criteria above.

The plan will:

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- Demonstrate how non-IMRT treatment planning cannot produce a satisfactory treatment plan (as stated above) via the use of patient-specific dose volume histograms and isodose plans.
- Provide tissue constraints for both the target and affected critical structures.



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IMRT for Post-operative Radiation

IMRT for post-operative radiation therapy is approvable. If there is gross residual disease and the area(s) can be sufficiently utilized, a boost can be added to a total dose of 60-70Gy, respecting normal tissue sensitivity. For gross nodal disease, consider boost to 60-65Gy while respecting normal tissue constraints.¹⁻³

Stereotactic Body Radiation Therapy (SBRT) Stereotactic Body Radiation Therapy is not a standard treatment option for the treatment of cervical cancer.

SBRT is an approach that allows for delivery of very high doses of focused EBRT in 1-5 fractions and may be applied to isolated metastatic sites, considering can be given for limited disease in the reirradiation setting.^{4,5}

Proton Beam Radiation Therapy

Proton beam is not an approved treatment option for cervical cancer. Proton beam has not been proven superior treatment to conventional radiation therapy.

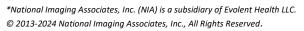
THE FOLLOWING APPLIES TO CMS (MEDICARE) MEMBERS ONLY:

For Proton Beam and Stereotactic Radiotherapy, refer to Local Coverage Determination (LCD), if applicable.

BACKGROUND

The role of radiation therapy in the treatment of cervical cancer has been long established through clinical trial, providing strong evidence of support as an effective cervical cancer treatment. The traditional approach utilizes external beam irradiation therapy to the pelvis ± periaortic lymph nodes, as well as some form of brachytherapy boost, based on clinical and pathologic factors. There have been improvements in radiation therapy technology, reducing dose to normal surrounding tissue (bladder, rectum, and small bowel), but the majority of the experience to date is based on a point A dosing system.

This guideline outlines several methods suitable for the employment of radiation therapy in conjunction with cervical cancer treatment. These include the use of three-dimensional conformal radiation therapy (3D-CRT), intensity-modulated radiation therapy (IMRT), image-guided radiation therapy (IGRT), and internal radiation (brachytherapy). Although intensity-modulated radiation therapy (IMRT) is becoming more widely available, the routine use in treating cervical cancer remains to be validated. IMRT may be useful when high doses are required to treat gross disease in regional lymph nodes. However, IMRT should not be used as routine alternatives to brachytherapy for treatment of central disease in patients with an intact cervix. Although there have been significant advances in imaging, planning, and treatment delivery, this must be tailored to a thorough understanding to the stage of disease, pathways for





dissemination and recurrence risk. Most external beam treatments are delivered using a high-energy linear accelerator. Brachytherapy is generally delivered as either low dose permanent implant or high dose rate implant. Principles of radiation therapy for these guidelines closely follow what is recommended both by the American Brachytherapy Society (Cervical Cancer Brachytherapy Task Group), as well as in National Comprehensive Cancer Network Practice Guidelines for Cervical Cancer.



REFERENCES

1. NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines): Cervical Cancer Version 1.2023. National Comprehensive Cancer Network (NCCN). Updated December 23, 2022. Accessed December 27, 2022. https://www.nccn.org/professionals/physician_gls/pdf/cervical.pdf

2. Klopp A, Yeung A, Deshmukh S, et al. A phase III randomized trial comparing patient-reported toxicity and quality of life (QOL) during pelvic intensity modulated radiation therapy as compared to conventional radiation therapy. *Int J Radiat Oncol Biol Phys.* 2016;96(2):S3.

doi:https://doi.org/10.1016/j.ijrobp.2016.06.024

3. Chino J, Annunziata CM, Beriwal S, et al. Radiation Therapy for Cervical Cancer: Executive Summary of an ASTRO Clinical Practice Guideline. *Pract Radiat Oncol*. Jul-Aug 2020;10(4):220-234. doi:10.1016/j.prro.2020.04.002

4. Choi CW, Cho CK, Yoo SY, et al. Image-guided stereotactic body radiation therapy in patients with isolated para-aortic lymph node metastases from uterine cervical and corpus cancer. *Int J Radiat Oncol Biol Phys.* May 1 2009;74(1):147-53. doi:10.1016/j.ijrobp.2008.07.020

5. Higginson DS, Morris DE, Jones EL, Clarke-Pearson D, Varia MA. Stereotactic body radiotherapy (SBRT): Technological innovation and application in gynecologic oncology. *Gynecol Oncol*. Mar 2011;120(3):404-12. doi:10.1016/j.ygyno.2010.11.042



POLICY HISTORY:

| Date | Summary |
|--------------|--|
| May 2023 | Under Definitive/Preoperative Radiation Therapy |
| | Added: IMRT is approvable when the para-aortic nodes are being treated |
| | • Revised: Pelvic radiation therapy from Stage IB2-IIA to Stage Stage IB2-IIIA |
| | Deleted Additional Resources |
| January 2022 | Added IMRT to Postoperative (Adjuvant) Radiation Therapy |
| | • Moved the following from Local/Regional Recurrence section to |
| | Postoperative (Adjuvant) Radiation Therapy section |
| | \circ Grossly involved unresected nodes may be evaluated for boosting |
| | with an additional 10-15Gy |
| | \circ Unless otherwise indicated, standard radiation fractionation consists |
| | of 1.8 Gy to 2.0 Gy per day |
| | Under Treatment Options Requiring Additional Clinical Review: |
| | Added IMRT for Post-operation Radiation |
| | \circ Clarified that SBRT can be given for limited disease in the re- |
| | irradiation setting |



Reviewed / Approved by NIA Clinical Guideline Committee:

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