Vaginal cuff brachytherapy: do we need to treat to more than a two-centimeter active length?

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Abstract

Purpose: American Brachytherapy Society (ABS) guidelines recommend using a 3-5 cm active length (AL) when treating vaginal cuff (VC) in adjuvant setting of endometrial cancer (EC). The purpose of this study was to evaluate local control and toxicity, using an AL of 1 or 2 cm and immobilization with a traditional table-mounted (stand) or patient-mounted (suspenders) device.

Material and methods: Between 2005 and 2019, 247 patients with EC were treated with adjuvant high-dose-rate vaginal cuff (HDR-VC) brachytherapy with or without external beam radiation (EBRT). Treatment was prescribed to a $0.5\,\mathrm{cm}$ depth, with an AL of 1 or 2 cm, using stand or suspenders. VC boost after EBRT was typically administered with 2 fractions of $5.5\,\mathrm{Gy}$, while VC brachytherapy alone was typically applied with 3 fractions of $7\,\mathrm{Gy}$ or $5\,\mathrm{fractions}$ of $5.5\,\mathrm{Gy}$.

Results: The combination of suspender immobilization and an AL of 2 cm (n = 126, 51%) resulted in 5-year local control of 100%. An AL of 2 cm compared to 1 cm correlated with better local control (99.1% vs. 88.5%, p = 0.0479). Regarding immobilization, suspenders correlated with improved local control compared to stand (100% vs. 86.7%, p = 0.0038). Immobilization technique was significantly correlated with AL (p < 0.0001). Only 5 (2.0%) patients experienced grade \geq 3 toxicity, all of whom received EBRT.

Conclusions: In the present series, an AL of 2 cm provided excellent local control, while 1 cm was inadequate. Suspender immobilization was a practical alternative to stand immobilization in HDR brachytherapy of the vaginal cuff.

J Contemp Brachytherapy 2021; 13, 3: 294–301 DOI: https://doi.org/10.5114/jcb.2021.105971

Received: 21.11.2020

Accepted: 09.03.2021

Published: 07.05.2021

Key words: vaginal cuff, brachytherapy, immobilization, active length, endometrial.

Purpose

Endometrial cancer is the most common gynecologic malignancy in the United States, with an estimated 65,620 new cases in 2020 [1]. Many women diagnosed at an early stage have excellent survival with surgery alone. A significant risk of recurrent disease remains for women with stage I disease and high-intermediate risk features, stage II disease [2], or in more advanced stages. Vaginal cuff is the most common site of post-operative failure, and vaginal cuff brachytherapy (VCB) has become an integral part of adjuvant therapy, either with or without external beam radiotherapy (EBRT) to the pelvis, to obviate the risk of vaginal cuff recurrence in appropriately selected patients [3-7].

There is a marked lack of consensus regarding vaginal cuff treatment parameters [7-10]. This has resulted in innumerable acceptable treatment options by the Ameri-

can Brachytherapy Society (ABS) [11, 12]. ABS guidelines recommend treating with an active length (AL) of 3-5 cm VCB, following hysterectomy [11]. Practices vary broadly, but the most common fixed and fractional length prescriptions in endometrial cancer are 4 cm or the proximal half of the vagina, respectively [8]. Wide practice variances are often due to a lack of data. However, length of vaginal irradiation as well as volume have been known to be associated with vaginal shortening and stenosis for decades [13-16]. Less described in the literature is a relationship between the length of vagina irradiation and vaginal recurrence [14, 17]. The common prescription depth of 5 mm, though potentially more toxic than prescribing to the surface, can be justified by location of vaginal lymphatics [8, 18, 19]. Without similar evidence, and in the absence of residual disease, common VBT prescription lengths may be excessive.

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Vaginal cuff movement can be significant from bladder and rectum volume variation [20, 21]. The vaginal cylinder itself can also cause displacement and altered dose distribution depending on how it is positioned [22]. Traditional table-mounted immobilization (i.e., stand) of the vaginal cylinder is susceptible to inter- and intra-fraction changes in patient positioning. Custommounted patient immobilization devices (e.g., suspenders), may better control for these inter- and intra-fraction changes.

The aim of the present study was to evaluate the efficacy, in terms of local control and toxicity, of using a 1 or 2 cm AL in the treatment of endometrial cancer with VCB. Similarly, we evaluated the impact of immobilization technique (table- vs. patient-mounted) within the same population.

Material and methods

After obtaining institutional review board approval (IRB 017-227), 247 patients were retrospectively identified

in the electronic medical record of two institutions who met all study criteria. Included patients were diagnosed with endometrial cancer and treated with adjuvant highdose-rate (HDR) brachytherapy to the vaginal cuff, following hysterectomy with or without EBRT to the vagina and pelvis between January 2005 and November 2019. Tables 1 and 2 present patients' and treatments' characteristics, respectively. Patients were excluded if they had gross residual disease in the vagina or endometrial sarcoma. Treatment of the study population was delivered via vaginal cylinder prescribed to a 0.5 cm depth, with an AL of 1 cm in 118 (47.7%) patients and 2 cm in 129 (52.2%) patients. Only 7 patients were identified who were treated with an AL of greater than 2 cm, and these patients were excluded from the analysis due to paucity of representation. Immobilization during brachytherapy was with a stand in 109 (44.1%) patients and suspenders in 138 (55.9%) patients. Suspender immobilization uses an in-house adapter attached to the vaginal cylinder central channel, with two strips of twill tape threaded into the adaptor and tied over the shoulders of the patient (Figure 1).

Table 1. Patient characteristics

| Variable | | <i>P</i> -value | | |
|--------------------------------|-------------------|---------------------|-------------------|----------|
| • | All, n (%) | 1 cm, n (%) | 2 cm, n (%) | |
| • | 247 (100) | 118 (47.8) | 129 (52.2) | |
| Immobilization | | | | < 0.0001 |
| Stand | 109 (44.1) | 106 (90.0) | 3 (2.3) | |
| Suspenders | 138 (55.9) | 12 (10.2) | 126 (97.7) | |
| Surgery type | | | | < 0.0001 |
| TAH/BSO | 57 (52.3) | 16 (13.6) | 41 (31.8) | |
| TAH/BSO/PLN | 49 (20.3) | 8 (6.8) | 41 (31.8) | |
| TAH/BSO/PLN/PaLN | 117 (48.4) | 81 (68.6) | 36 (27.9) | |
| Other | 19 (7.9) | 10 (8.5) | 9 (7.0) | |
| Stage | | | | 0.01 |
| IA | 94 (38.2) | 56 (47.5) | 38 (29.5) | |
| IB | 73 (29.7) | 32 (27.1) | 41 (31.8) | |
| II | 17 (6.9) | 3 (2.5) | 14 (10.9) | |
| III | 58 (23.6) | 25 (21.2) | 33 (25.6) | |
| IV | 4 (1.6) | 2 (1.7) | 2 (1.6) | |
| Pathology | | | | 0.0081 |
| Adenocarcinoma | 178 (72.1) | 91 (77.1) | 87 (67.4) | |
| Papillary serous/clear cell | 45 (18.2) | 23 (19.5) | 22 (17.1) | |
| Adenosquamous | 18 (7.3) | 4 (3.4) | 14 (10.9) | |
| Other | 6 (2.4) | 0 (0.0) | 6 (4.7) | |
| Grade | | | | 0.9683 |
| 1 | 33 (14.7) | 12 (10.2) | 21 (16.3) | |
| 2 | 82 (36.6) | 42 (35.6) | 40 (31.0) | |
| 3 | 109 (48.6) | 48 (40.7) | 61 (47.3) | |
| Follow-up time, median (range) | 36 months (3-144) | 46.5 months (3-113) | 28 months (3-144) | |

TAH – total abdominal hysterectomy, BSO – bilateral salpingo-oophorectomy, PLN – pelvic lymph node dissection, Plan – paraaortic lymph node dissection

Table 2. Radiation treatment characteristics

| Variable | 1 cm active length | 2 cm active length | <i>P</i> -value |
|---|---|---|-----------------|
| Most common treatment regimens to VC only (BED ₂) | 5.5 Gy × 5 ($n = 37, 35.5 \text{ Gy}_2$), 5.5 Gy × 4 ($n = 16, 28.4 \text{ Gy}_2$) | 7 Gy × 3 ($n = 39, 29.8 \text{ Gy}_2$), 5.5 Gy × 4 ($n = 7, 28.4 \text{ Gy}_2$) | N.A. |
| Most common boost treatment regimen with $EBRT$ (BED_2) | 5.5 Gy × 2 following 50.4 (n = 27) or 45.0 Gy (n = 22) of EBRT (58.5-63.8 Gy ₂) | 5.5 Gy × 2 following 50.4 (n = 31) or 45.0 Gy (n = 12) of EBRT (58.5-63.8 Gy ₂) | N.A. |
| VC dose per fraction | 550 cGy (400-708.3) | 550 cGy (400-700) | < 0.0001 |
| VC total dose | 2050 cGy (400-4250) | 1100 cGy (550-3500) | < 0.0001 |
| VC number of fractions | 3 (1-6) | 2 (1-5) | < 0.0001 |
| EBRT | | | |
| Yes | 57 (23.1) | 50 (20.2) | 0.0412 |
| No | 61 (24.7) | 79 (32.0) | |
| EBRT dose, median (range) | 5040 cGy (4500-5040) | 5040 cGy (4500-5625) | 0.0159 |
| Total VC BED ₂ , median (range) | 4899 cGy (933-7756) | 5846 cGy (992-8344) | 0.3254 |

VC – vaginal cuff, EBRT – external beam radiotherapy, BED_2 – biologic equivalent dose at 2 Gy, N.A. – not applied



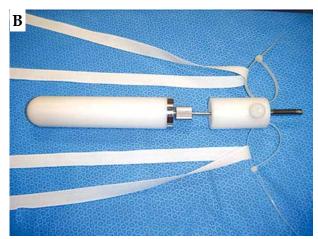




Fig. 1. Immobilization by **A**) suspenders with **B**) custom in-house device, or with **C**) stand

Prior to treatment planning, fiducial markers were placed at the vaginal cuff, usually at 3 and 9 o'clock. Treatment planning was either 2D- or 3D-based, without clinical target volume (CTV) delineation. For 179 pa-

tients, 2D atlas-based library plans were used with fluoroscopy. For 68 patients, 3D planning was used with a computed tomography (CT) scan performed prior to each fraction to ensure correct placement and diameter of the cylinder. A remote afterloader tested for channel obstruction and dwell times were independently checked. Daily QA included verification of the source position and interlocks. At the time of source exchange, after calibration, verification of the source positioning was performed. All vaginal cylinders were examined annually for physical integrity, with scans used to check for center channel damage and to verify offsets used in the treatment planning system. Cylinder diameter was chosen based on patient anatomy. Treatment plans were optimized using treatment planning software to ensure 100% isodose line coverage at 5 mm depth. On extremely rare occasions, isodose lines were manually modified to avoid the bladder or rectum.

After completion of radiation, the patients were followed every 3 months during the first 2 years and then, every 6 months up to 5 years or more. Median follow-up time was 36 months (range, 3-144 months). They were evaluated for recurrence and side effects by clinical and gynecological examinations as well as imaging studies. The patients were not censored for vaginal recurrence at the time of distant or nodal recurrence and continued to receive monitoring.

Toxicity was assessed using common terminology criteria for adverse events v. 5.0 guidelines. The highest grade of vaginal toxicity during treatment or on follow-up with a radiation oncologist or gynecologic oncologist was recorded. For all women, vaginal dilators were provided but compliance was infrequently reported and thus precluded from toxicity analysis.

Chi-square, Fisher's exact, or Wilcoxon rank-sum tests were used to compare characteristics of the 1 and 2 cm AL populations. Bivariate analysis was used to assess overall survival by AL and vaginal recurrence as well as vaginal recurrence by AL and immobilization method. Chi-square or Fisher's exact tests were used to test for an association between AL and toxicity. Low number of events precluded multivariate analysis. SAS version 9.4 (SAS Institute Inc., Cary, NC, USA) was used for statistical analysis.

Results

The patients' and treatments' characteristics are presented in Tables 1 and 2. The patients received different fractionation schedules in terms of total dose and number of fractions, all of which were converted to a biologically effective dose (BED)₂ at 0.5 cm depth using the L/Q model (Table 2). Those who received EBRT, typically obtained 4500 or 5040 cGy with subsequent brachytherapy to two fractions of 550 cGy (58.5-63.8 Gy₂ total), while patients who did not, typically received 3 fractions of 700 cGy or 5 fractions of 550 cGy (35.5 or 29.8 Gy₂, respectively). Patients treated with an AL of 2 cm were significantly more likely to be immobilized with suspenders than a stand (97.7% vs. 2.3%, p < 0.0001).

For all the patients, the overall survival (OS) rates at 2 and 5 years was 95.5% and 77.0%, respectively. An OS difference was not detected by AL (p = 0.8050). Patients with a vaginal recurrence had a 5-year OS of 61.7% vs. 77.7% for those without (p = 0.2654).

For all the patients, the vaginal recurrence rates at 2 and 5 years were 1.9% and 8.0%, respectively. Overall recurrence rates at 2 and 5 years were 11.9% and 20.3%, respectively. Patients treated with an AL of 1 cm had a significantly worse 5-year vaginal recurrence free survival (VRFS) than those with an AL of 2 cm (88.5% vs. 99.1%, p=0.0479). Patients treated with suspender immobilization had a higher 5-year VRFS (100.0% vs. 86.7%, p=0.0038). The most common combination of 2 cm AL with suspenders (n=126, 51%) had a 100% local control at 5 years (Figure 2). Immobilization technique was significantly correlated with AL (p < 0.0001). Patients treated with or without EBRT did not have a significantly different VRFS (94.3% vs. 91.5%, p=0.4354). Low number of events prevented log-rank analysis by EBRT and AL (Figure 3).

Table 3 shows patients' toxicities. Grades 1-2 vaginal toxicity were experienced in 21.7% and 14.6% of patients with and without EBRT, respectively. No grade 3 or 4 vaginal toxicities were documented. Grades 1-3 rectal toxicity were experienced in 63.4% and 8.2% of patients with and without EBRT, respectively. Grades 1-3 bladder

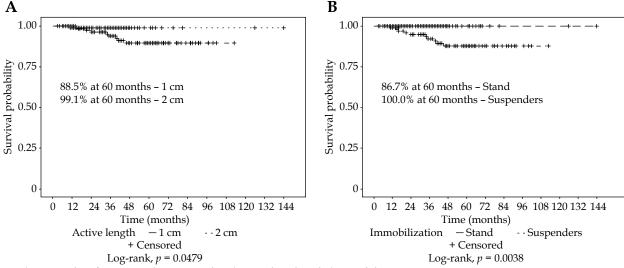


Fig. 2. Freedom from vaginal recurrence by A) active length and B) immobilization

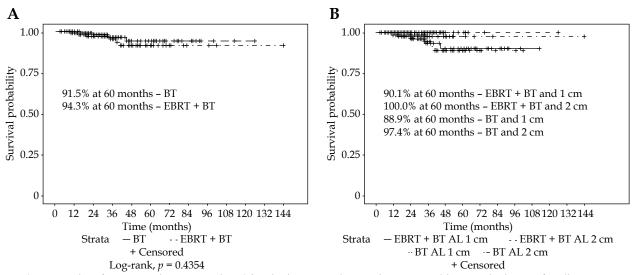


Fig. 3. Freedom from vaginal recurrence by A) brachytherapy with or without external beam radiotherapy for all patients, or by B) active length

Table 3. Toxicity

| Туре | Grade | Patients, n (%) | | P-value ¹ With EBRT, n (| | RT, n (%) | <i>P</i> -value ² | Without EBRT, n (%) | | <i>P</i> -value ¹ | |
|---------|-------|-----------------|------------|-------------------------------------|----------|-----------|------------------------------|---------------------|-----------|------------------------------|--------|
| | | Total | With | Without | | 1 cm AL | 2 cm AL | | 1 cm AL | 2 cm AL | |
| | | | EBRT | EBRT | | | | | | | |
| Vaginal | None | 198 (81.5) | 105 (43.2) | 93 (38.3) | 0.1646 | 48 (35.8) | 57 (42.5) | 0.0798 | 55 (50.5) | 38 (34.9) | 0.0383 |
| | 1 | 35 (14.4) | 29 (11.9) | 16 (6.6) | | 8 (6.0) | 21 (15.7) | | 5 (4.6) | 11 (10.1) | |
| | 2 | 10 (4.1) | | | | | | | | | |
| Rectal | None | 149 (61.3) | 49 (20.2) | 100 (41.2) | < 0.0001 | 22 (16.4) | 27 (20.2) | 0.5798 | 55 (50.5) | 45 (41.3) | 1.0000 |
| | 1 | 43 (17.7) | 85 (35.0) | 9 (3.7) | | 34 (25.4) | 51 (38.1) | | 5 (4.6) | 4 (3.7) | |
| | 2 | 47 (19.3) | | | | | | | | | |
| | 3 | 4 (1.7) | | | | | | | | | |
| Bladder | None | 183 (75.3) | 86 (35.4) | 97 (39.9) | < 0.0001 | 38 (28.4) | 48 (35.8) | 0.4518 | 50 (45.9) | 47 (43.1) | 0.0368 |
| | 1 | 43 (17.7) | 48 (19.8) | 12 (4.9) | | 18 (13.4) | 30 (22.4) | | 10 (9.2) | 2 (1.8) | |
| | 2 | 16 (6.6) | | | | | | | | | |
| | 3 | 1 (0.4) | | | | | | | | | |

¹ With EBRT vs. without EBRT, any grade toxicity, 21 cm vs. 2 cm, any grade toxicity, EBRT – external beam radiotherapy, AL – active length prescription

toxicity were experienced in 35.8% and 11.0% of patients with and without EBRT, respectively.

Discussion

Post-operative irradiation decreases locoregional relapses in high/intermediate-risk endometrial cancer, and vaginal cuff brachytherapy achieves vaginal control similar to EBRT with lower toxicity [3, 4]. Even though not demonstrated in randomized trials, the National Cancer Database and the Surveillance, Epidemiology, and End Results Database have shown evidence for improved survival in intermediate- and high-risk groups [23-25]. Following the PORTEC-2 demonstrating equivalent local control of vaginal cylinder to EBRT in early-stage patients with decreased toxicity [3], it makes sense to find the least toxic way to administer vaginal radiation without sacrificing local control. This is especially true without a well-established survival benefit.

In patients treated with an AL of 2 cm with suspenders, no patient developed a recurrence of the vaginal cuff. Seldom described is the effect of vaginal irradiation length on vaginal recurrence and survival. Kloetzer et al. observed no difference in survival or vaginal recurrence treating only proximal half of the vagina [17]. Surprisingly, one analysis found that overall recurrence was correlated with increased prescription length, though this was not discussed in the paper, and it was most likely due to differences in disease [14]. We found a significant difference in vaginal cuff recurrence between patients treated with an AL of 1 and 2 cm (5-year recurrence rates of 11.5% and 0.9%, respectively). There were many significant differences between the 1 and 2 cm populations, but most of those differences strengthen this finding. Patients treated with an AL of 2 cm had significantly higher stages, less extensive nodal dissections, less EBRT at lower doses, and lower total brachytherapy dose. A difference that weakens our finding was the decreased amount of

Table 4. Vaginal recurrence in the literature

| Vaginal recurrence following adjuvant vaginal cuff brachytherapy | | | | | | | | | | |
|--|-------------------------------------|------|--------------|--------------------|-------------|--|-------------------------------------|-----------------------|--------------------------------|--|
| Authors | AL (cm or vaginal fraction) | Year | Patients (n) | FIGO stage | EBRT (%) | Brachytherapy treat- ment (dose × fractions) | Depth (cm) | Median FU (months) | Vaginal recur- rence (%) | |
| Current study | 21 | 2020 | 126 | IA-IV | 61.2 | Median 7 Gy × 3 or 5.5 Gy × 2 (w/EBRT) ³ | 0.5 | 28.0 | 0.0 | |
| | 1 | 2020 | 118 | IA-IV | 48.3 | Median 5.5 Gy × 5 or × 2 (w/EBRT) ⁴ | 0.5 | 47.0 | 7.6 | |
| Hou <i>et al</i> . [26] ² | Proximal 1/2 | 2019 | 390 | stage I (IR-HR) | 37.2 | 5 Gy × 5-6 or 4-6 Gy × 2-3 (w/EBRT) | 0.5 | 48.0 | 1.0 | |
| Zhang <i>et al</i> . [27] | 2.5 | 2019 | 325 | IA-IVB | 100.0 | 4-6 Gy × 3, 5-6 Gy × 2, or 7 Gy × 1 | 0.5 | 51.0-95.0 | 1.8 | |
| Rovirosa <i>et al</i> . [28] | 2.5 | 2017 | 146 | IA-II (IR) | 0.0 | 4-6 Gy × 6, 4-5 Gy × 4, or 6 Gy × 3 | 0.5 | 41.0-88.0 | 0.0 | |
| Sorbe <i>et al</i> . [29] | Proximal 2/3 | 2012 | 527 | IA-IC (IR) | 50.0 | 3 Gy × 6, 5.9 Gy × 3, or 20 Gy (LDR) | 0.5 | 62.0 | 3.3 | |
| Diavolitsis et al. [30] | 3-5 | 2012 | 169 | IA | 0.0 | 7 Gy × 3, 5.5 Gy × 4, or 70 Gy (LDR) | 0.5 or ovoid surface (LDR) | 103.0 | 0.6 | |
| Nout <i>et al</i> . (PORTEC-2) [3] | 4.7 (mean) | 2010 | 213 | IB-2A (HIR) | 0.0 | 7 Gy × 3 or 30 Gy (LDR) | 0.5 | 45.0 | 1.8 | |
| Alektiar <i>et al</i> . [31] | Proximal 1/2 to 2/3 (> if G3) | 2005 | 382 | IB-IIB (IR) | 0.0 | 6-7 Gy × 3 | 0.5 | 48.0 | 1.8 | |
| Horowitz <i>et al</i> . [32] | 5 | 2002 | 164 | IB-IIB (IR) | 0.0 | 7 Gy × 3 | 0.5 | 65.0 | 1.2 | |
| Petereit <i>et al</i> . [33] | Not reported | 1999 | 191 | IA-C (LR-IR) | 0.0 | 16.2 Gy × 2 | Ovoid surface | 38.0 | 0.0 | |
| Eltabbakh et al. [34] | Not reported | 1997 | 332 | I (LR) | 0.0 | 30 Gy (LDR) | 0.5 | 97.0 | 0.0 | |
| Sorbe <i>et al</i> . [13] | Proximal 2/3 | 1990 | 404 | IA-IB | 0.0 | 4.5 Gy × 6 to 9 Gy × 4 | 1.0 | Not report- ed | 0.7 | |

AL- active length of prescription, EBRT - external beam radiotherapy, FU- follow-up, IR- intermediate-risk, HR- high-risk, LDR- low-dose-rate, G- grade, 1 with suspenders, 2 with multi-channel applicators, 3 range, 4-7 Gy \times 1-6

follow-up time in the patients treated with an AL of 2 cm compared to 1 cm, which may have resulted in later recurrences not being captured. We were also unable to perform a multivariate analysis due to the low number of events. Table 4 demonstrates our recurrence rates in context of the literature [3, 13, 26-34]. Not all of these papers distinguished between a vaginal cuff specific recurrence and vaginal recurrence overall. In our patients, all recurrences had an epicenter within 2 cm of the vaginal apex, such that there was no difference. These papers also had varying proportions of patients who received EBRT, but receiving EBRT did not significantly impacted VRFS in our study.

The length of vagina irradiation is known to be associated with vaginal shortening and stenosis [13, 35]. A recent analysis showed stenosis rates of 5% and 31%, with prescription lengths of 2.5 cm and 5.0 cm, respectively [14]. This same analysis found that stenosis rates correlated significantly with prescription depth (4% vs. 21% at

the surface and 5 mm, respectively), but not with cylinder diameter, which, unlike prescription point and length, physicians tend to alter based on individual anatomy. Agnes *et al.* retrospectively analyzed 100 patients with endometrial cancer who received exclusive VBT post-operatively, with a median follow-up of 24 months, and found a correlation between an AL \geq 5 cm and increased vaginal toxicity (p = 0.002) in patients who had received VBT without EBRT. In the shortest AL series published, Zhang *et al.* observed that vaginal shortening was \geq 1/3 in 9.7% of patients treated with an AL of 2.5 cm. Late grade 2 vaginal toxicity in post-operative endometrial carcinoma was associated with a 68 Gy dose equivalent to 2 Gy per fraction ($\alpha/\beta = 3$ Gy) at 2 cm³ of the vagina in patients who received EBRT and intracavitary radiation [16, 27].

Vaginal toxicity itself, particularly with regard to sexual function, is not well reported or standardized. Compliance with vaginal dilators, which in some studies has been shown to decrease vaginal atrophy [15, 36, 37], is

often not recorded, and this remained true in our retrospective analysis. A Cochrane review found insufficient reliable evidence for routine vaginal dilator use, but still recommended it in patients not sexually active [38]. Vaginal length changes can be reliably measured but alterations in sexual function are more difficult to quantify [39, 40]. Zhang et al. assessed late toxicity of the rectum and bladder using Radiation Therapy Oncology Group (RTOG) scores, while late vaginal toxicity was evaluated with the objective criteria of LENT-SOMA [41, 42]. Here, we assessed toxicity using CTCAE 5.0 guidelines. Since no clear correlation of time from treatment to development of vaginal stenosis has been found in the literature [15, 35], we recorded the highest grade of vaginal toxicity during the treatment or at any available follow-up with a radiation or gynecologic oncologist. We found a higher rate of both bladder and vaginal toxicity (any grade) in patients treated with an AL of 1 cm. This finding was unexpected and is most likely a result of provider and/or institutional bias, as all 1 cm patients were treated by several radiation oncology physicians, while a single physician treated a majority of 2 cm patients at multiple institutions. Additionally, the significantly shorter follow-up time of 2 cm patients (28 months) may have contributed to less late toxicity events being captured.

Unfortunately, similar institutional bias diminished the efficacy of analysis of toxicity or recurrence by immobilization type. We found a significant difference in vaginal cuff recurrence by stand vs. suspender immobilization technique (5-year recurrence rates of 13.3% and 0.0%, respectively). However, there was a significant correlation between suspender immobilization and a 2 cm AL. Only 3 patients (1.2%) were treated with a 2 cm AL using a stand, with 1 of them having a vaginal recurrence. Interestingly, this was the only vaginal recurrence in the patients treated with a 2 cm AL. Due to mild toxicity in patients who did not receive EBRT, any potential difference in side effects using suspenders would be difficult to detect.

Suspender immobilization was clinically beneficial in several instances. A number of patients moved during a treatment, usually because of pain or coughing, and thus had to be restimulated and planned. This was not the case with suspender immobilization, as the vaginal cylinder "moves" with the patient. At one institution, patients were typically treated in an HDR suite, following CT simulation in the same room. When the CT scanner was down for a repair, patients were able to receive their simulation in a separate room and then be transported for the treatment with the vaginal cylinder in place. Vaginal cylinders and stands can be marked in various ways, but external markings cannot always accurately replicate internal anatomical changes. The VC can move up to 34.5 mm during EBRT [20]. These are largely due to variations in the bladder and rectum volumes between fractions, but a cylinder can also cause different displacement of adjacent organs based on patient's and stand's positioning [21, 22]. With suspender immobilization, the vaginal cylinder neutrally follows and stretches patient's vaginal canal, which minimizes air pockets and improves coverage at 0.5 cm depth, additionally allowing for adjustments that conform to motion of the vaginal canal.

Limitations of this study include its retrospective nature and a heterogeneous cohort, with regard to stage, dose, follow-up time, and surgery. Most significantly, patients treated with an AL of 2 cm had a significantly shorter follow-up time. There was a limited number of patients treated with an AL of 2 cm with a stand immobilization or an AL of 1 cm with suspender immobilization.

Conclusions

Excellent control of the vaginal cuff was obtained using patient-mounted suspender immobilization while treating to an AL of 2 cm. Treating to an AL of 1 cm correlated with a significantly increased frequency of the vaginal cuff recurrence. The suspender immobilization offers several practical advantages and is a viable alternative to the stand immobilization in HDR brachytherapy of the vaginal cuff. Optimal brachytherapy with regards to dose, fractionation, and other treatment parameters remains ill-defined. More research, preferably with randomized phase III trials incorporating more objective and detailed monitoring of vaginal toxicity are needed.

Disclosure

Dr. Barry received an ACRO grant for a survey evaluating promotion of women in radiation oncology. Otherwise, none to disclose.

The authors report no conflict of interest.

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