

Comparison of Quality of Life and Sexuality between Cervical Cancer Survivors and Healthy Women

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Purpose

The purpose of this study is to compare quality of life (QoL) and sexual functioning between sexually active cervical cancer survivors and healthy women.

Materials and Methods

In this cross-sectional study, propensity-score-matched cervical cancer survivors (n=104) and healthy women (n=104) were compared. All women had engaged in sexual activity within the previous 3 months, and cervical cancer survivors showed no evidence of disease after primary treatment. QoL and sexual functioning were assessed using three questionnaires; the European Organization for Research and Treatment of Cancer Quality of Life Questionnaire Core 30 (EORTC QLQ-C30), Cervical Cancer Module (EORTC QLQ-CX24), and the Female Sexual Function Index (FSFI).

Results

Significantly higher scores for lymphedema were observed in the cervical cancer survivors group compared with the healthy women group (mean, 20.2 vs. 12.2; $p < 0.05$). Sexuality, both in terms of sexual activity, sexual enjoyment, and sexual worry (EORTC QLQ-CX24), and in terms of desire, arousal, lubrication, orgasm, satisfaction, and pain (FSFI) were similar between the groups. When the scale of sexual/vaginal functioning in EORTC QLQ-CX24 was divided into individual questions, cervical cancer survivors reported shorter vaginal length than the control group, but without statistical significance (mean, 80.6 vs. 85.4; $p=0.077$).

Conclusion

Compared with healthy women, sexuality was not impaired in cervical cancer survivors who showed no evidence of disease after primary treatment and engaging in sexual activity. Further prospective cohort studies are warranted to confirm this finding.

Key words

Uterine cervical neoplasms, Quality of life, Sexuality, Survivors

Introduction

In 2015, cervical cancer was predicted to account for 2.2% of all new cancers (n=3,100) and 2.6% of all cancer deaths (n=761) among Korean women [1]. The incidence of cervical cancer is much higher than that of other gynecologic malignancies such as endometrial cancer and ovarian cancer [2]. As the recent survival of cervical cancer patients has improved, quality of life (QoL), including sexuality, has

become an important issue for survivors [3].

Reports on sexuality in Korean women have been limited [4-6]. For women with gynecologic cancer, survivors' QoL and sexual functioning worsened after treatment [3,7]. Treatment for cervical cancer is composed of radiotherapy, chemotherapy, and radical surgery, and various combinations of these modalities are frequently administered. Considering these aspects, some studies have evaluated the extent of deterioration according to the type of treatment [8]. However, small size of the study population, heterogeneity

in patient's characteristics, and use of different questionnaires among the studies hinder inferences from the results. In addition, it is necessary to measure cervical cancer survivors' sexuality using authorized questionnaire sets and to perform a comparison with that of a proper control group.

Sexual-wellbeing is an important part of cancer survivorship. Therefore, precise information on this issue, which might be affected by treatment, should be provided to cancer survivors as well as to their partners. Thus, the aim of this study was to compare QoL and sexual functioning between cervical cancer survivors and healthy women.

Materials and Methods

After obtaining approval from the Institutional Review Board (NCCNCS-13-685), the current study was conducted in the outpatient clinic of the National Cancer Center, Korea during the period between February 2013 and January 2014. To be eligible for inclusion, cervical cancer survivors had to have no evidence of disease, be 18 years of age, be able to understand Korean, and had to have engaged in sexual activity within 3 months of the study. Women who had not been diagnosed with any type of cancer in their medical records and who had engaged in sexual activity within the previous 3 months were assigned to the control group. Those who refused to participate or failed to complete the questionnaires were excluded.

Enrolled women's demographic data, including age, education, family income, marital status, and occupational status, were collected by interview and review of the medical records. Women's QoL was assessed using the European Organization for Research and Treatment of Cancer Quality of Life Questionnaire Core 30 (EORTC QLQ-C30) and Cervical Cancer module (EORTC QLQ-CX24).

The EORTC QLQ-C30 is a 30 item, cancer-specific questionnaire for assessment of five functioning scales (physical, role, emotional, cognitive, and social functions), three symptom scales (fatigue, pain, and nausea and vomiting), a scale for global health status and overall QoL, and six single-items (dyspnea, insomnia, appetite loss, constipation, diarrhea, and financial difficulties) [9]. The scales and single-item measures are scored from 0 to 100. In the functional and global health scales, higher scores indicate better functioning or status, while higher scores on the symptom scales and for the single-items reflect more problematic symptoms.

The EORTC QLQ-CX24 is a 24 item, cervical cancer-specific questionnaire composed of three multi-item scales (symptom experience, body image, and sexual/vaginal functioning) and six single-item scales (lymphedema, peripheral

neuropathy, menopausal symptoms, sexual activity, sexual worry, and sexual enjoyment) [10]. The scales and single-item measures are also scored from 0 to 100. In sexual activity and sexual enjoyment, higher scores indicate better functioning, however, in the other scales and single-items, higher scores reflect more problematic functioning or status.

For assessment of women's sexual functioning, the Korean version of the Female Sexual Function Index (FSFI) was also used [11]. This 19-item questionnaire includes six domains: desire, arousal, lubrication, orgasm, satisfaction, and pain. A higher score in each domain indicates better status.

Statistical analysis was performed using the R statistical software ver. 2.12.1 (R Foundation for Statistical Computing, Vienna, Austria; ISBN 3-900051-07-0; <http://www.Rproject.org>). Because the demographic characteristics of the cervical cancer survivors and control groups were different, the two groups were matched in age, education, family income, marital status, occupational status, and exercise using propensity score matching. Count percentage, and quartile were used for analysis of the demographic and clinical characteristics of the participants. *t* tests were performed for comparison of continuous variables between cervical cancer survivors and control groups.

Results

Among the women who agreed to participate in the study, 135 cervical cancer survivors and 220 women in the control group provided informed consent and completed the questionnaires (Fig. 1). After the propensity score matching, demographic data, including age, education, family income, marital status, occupational status, and regular exercise were not significantly different between the cervical cancer survivors and control group (Table 1).

The clinical characteristics of cervical cancer survivors are shown in Table 2. The majority of cervical cancer survivors (95.2%) were International Federation of Gynecology and Obstetrics stage I and II diseases. Squamous cell carcinoma was the most frequent histologic type (77.9%). The median interval from the diagnosis of cervical cancer to the current study was 45 months (range, 5 to 211 months). The proportions of the survivors according to the intervals between diagnosis to survey were as follows: < 2 years, 29.8%; 2-5 years, 37.5%; and > 5 years, 32.7%.

In terms of the treatment, radical hysterectomy was performed in 63.5% of the cervical cancer patients. Twelve point five percent of the survivors received all three treatment modalities: radical hysterectomy, radiotherapy, and chemotherapy.

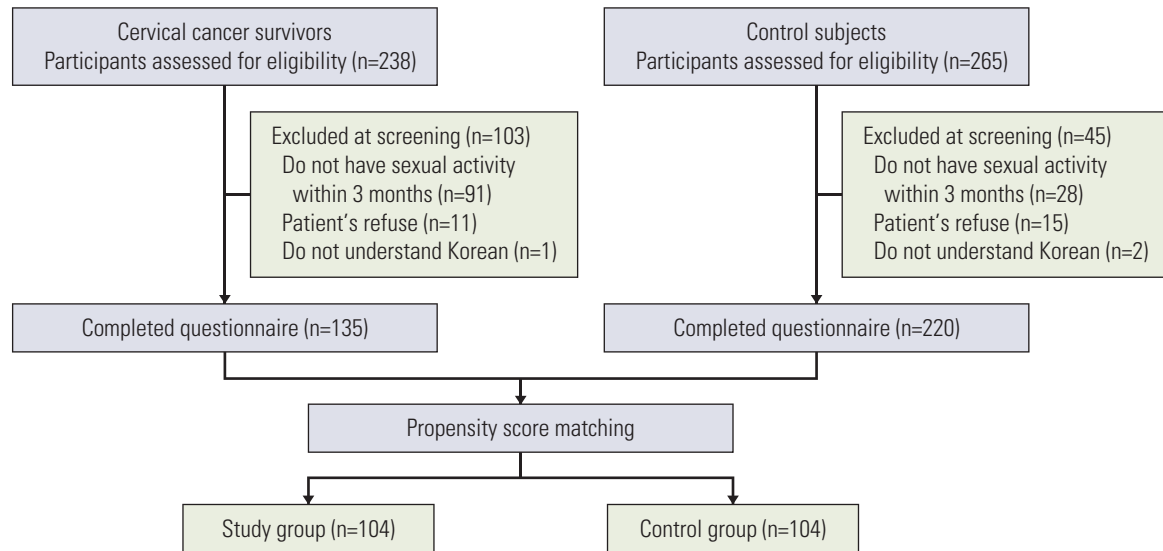


Fig. 1. Flow chart of enrollment.

Table 1. Demographic characteristics of cervical cancer survivors and control group

Characteristic	Cervical cancer survivor (n=104)	Control group (n=104)	p-value
Age (yr)	46.95±7.98	48.86±8.53	0.098
Marital status			
Single/Separated/Widowed	14 (13.5)	11 (10.6)	0.670
Married	90 (86.5)	93 (89.4)	
Education (yr)			
≤ 9	12 (11.5)	14 (13.5)	0.834
> 9	92 (88.5)	90 (86.5)	
Family income (\$/mo)			
≤ 2,000	13 (12.5)	16 (15.4)	0.833
> 2,000 to ≤ 4,000	45 (43.3)	44 (42.3)	
> 4,000	46 (44.2)	44 (42.3)	
Occupational status			
Yes	45 (43.3)	49 (47.1)	0.676
No	59 (56.7)	55 (52.9)	
Menopause status			
Yes	76 (73.1)	67 (64.4)	0.231
No	28 (26.9)	37 (35.6)	
Regular exercise			
Yes	50 (48.1)	45 (43.3)	0.578
No	54 (51.9)	59 (56.7)	

Values are presented as mean±standard deviation or number (%).

The comparison of the EORTC QLQ-C30 and CX24 scores between cervical cancer survivors and control group is shown in Table 3. Among the scales, only scores for lymphedema (mean, 20.2 vs. 12.2; $p=0.022$) and body image

(mean, 73.1 vs. 79.6; $p=0.036$) were significantly different between the two groups. When the scale of sexual/vaginal functioning in EORTC QLQ-CX24 was divided into individual questions, cervical cancer survivors reported shorter

Table 2. Clinical characteristics of cervical cancer survivors

Characteristic	Cervical cancer survivor (n=104)
FIGO stage	
I	81 (77.9)
II	18 (17.3)
III	4 (3.8)
IV	1 (1.0)
Histology	
Squamous cell carcinoma	81 (77.9)
Adenocarcinoma	22 (21.2)
Small cell carcinoma	1 (1.0)
Survival period (yr)	
≤ 2	31 (29.8)
> 2 to ≤ 5	39 (37.5)
> 5	34 (32.7)
Type of treatment	
Conization only	14 (13.5)
Simple hysterectomy only	8 (7.7)
Radical hysterectomy only	43 (41.3)
Conization+R+C	2 (1.9)
Simple hysterectomy+C	1 (1.0)
Simple hysterectomy+R+C	2 (1.9)
Radical hysterectomy+C	4 (3.8)
Radical hysterectomy+R	6 (5.8)
Radical hysterectomy+R+C	13 (12.5)
Staging surgery only+R	3 (2.9)
R only	1 (1.0)
R+C	7 (6.7)
Type of surgery	
Conization	16 (15.4)
Simple hysterectomy	11 (10.6)
Radical hysterectomy	66 (63.5)
Staging surgery	3 (2.9)
No surgery	8 (7.7)
Status of ovaries	
Ovaries <i>in situ</i>	79 (76.0)
Ovarian transposition	1 (1.0)
Bilateral oophorectomy	18 (17.3)
Unilateral oophorectomy	6 (5.8)

Values are presented as number (%). Staging surgery was defined as performance of pelvic lymphadenectomy and para-aortic lymphadenectomy without hysterectomy. FIGO, International Federation of Gynecology and Obstetrics; R, radiotherapy; C, chemotherapy.

vaginal length than the control group, but without statistical significance (mean, 80.6 vs. 85.4; $p=0.077$). Other scale scores of the EORTC QLQ-CX24 including sexual activity (32.1 vs. 30.1, $p=0.583$), sexual enjoyment (40.1 vs. 37.8, $p=0.537$), and sexual worry (26.6 vs. 22.0, $p=0.214$) were similar between cervical cancer survivors and control group.

The comparison of the FSFI scores between two groups is also shown in Table 3. Scale scores for desire (3.0 vs. 2.9, $p=0.617$), arousal (3.5 vs. 3.2, $p=0.232$), lubrication (4.1 vs. 4.0, $p=0.696$), orgasm (3.9 vs. 3.7, $p=0.356$), satisfaction (4.0 vs. 3.9, $p=0.363$), and pain (4.5 vs. 4.1, $p=0.214$) were not significantly different between cervical cancer survivors and control group.

Discussion

In the current study, lymphedema was more problematic in cervical cancer survivors compared to the control group. Sexuality, both in terms of sexual activity, sexual enjoyment, and sexual worry (EORTC QLQ-CX24), and in terms of desire, arousal, lubrication, orgasm, satisfaction, and pain (FSFI) were similar between the two groups.

Data from previously published studies on cervical cancer survivors' sexuality which used the FSFI are summarized in Table 4 [12-17]. Sexual functioning deteriorated after radiotherapy, radical hysterectomy, and radical trachelectomy [14-16]. Older age and lower level of education were associated with more deteriorated sexual functioning of the survivors [17]. Importantly, impact of the type of treatment on sexuality was inconsistent among the previous studies [12, 14]. In the current study, with 45 months of median interval from the diagnosis, cervical cancer survivors showed similar total FSFI scores compared to the control group (23.0 vs. 21.9, $p=0.300$).

It has been reported that survivors of gynecologic malignancies express a reluctance to engage in sexual activity due to the fear of recurrence [18]. In the current study, the proportion of women excluded because of absence of sexual activity within 3 months was 38.2% (91/238) in cervical cancer survivors, whereas it was only 10.6% (28/265) in the control group. In addition, as Asians are known to be more sexually conservative, Korean women are considered to have relatively lower levels of sexual interest and activity [19]. These features might result in similar sexual functioning between cervical cancer survivors and control group.

In the current study, the target participants were limited to women who had engaged in sexual activity within 3 months. Although the diagnosis of sexual dysfunction is usually evaluated sexual problems within the last 6 months at

Table 3. Comparison of quality of life and sexual function between cervical cancer survivors and control group

Domain	Cervical cancer survivor (n=104)	Control group (n=104)	p-value
EORTC QLQ-C30			
Physical functioning	79.3±15.5	80.6±14.5	0.539
Role functioning	85.7±19.8	87.3±17.1	0.533
Emotional functioning	77.9±20.9	76.8±21.9	0.706
Cognitive functioning	77.9±17.6	78.4±16.9	0.841
Social functioning	83.2±24.3	88.1±21.1	0.117
Fatigue	35.1±22.7	39.9±23.9	0.147
Nausea and vomiting	5.8±13.3	8.7±16.9	0.173
Pain	15.2±19.6	19.4±21.7	0.147
Dyspnea	12.2±20.8	12.8±19.9	0.820
Insomnia	23.4±27.8	25.3±26.9	0.613
Appetite loss	11.2±19.5	10.3±18.0	0.712
Constipation	25.0±28.2	21.8±28.5	0.416
Diarrhea	14.7±21.2	11.9±22.2	0.339
Financial difficulties	11.2±20.6	10.3±21.3	0.741
Global health status	64.5±20.2	59.9±20.5	0.108
EORTC QLQ-CX24			
Body image	73.1±25.2	79.6±18.8	0.036
Symptom experience	12.1±10.1	10.5±8.9	0.244
Lymphedema	20.2±28.0	12.2±21.8	0.022
Peripheral neuropathy	22.4±28.4	17.0±23.2	0.132
Menopausal symptoms	20.5±27.2	17.3±23.2	0.362
Sexual activity	32.1±24.5	30.1±26.6	0.583
Sexual enjoyment	40.1±24.3	37.8±25.9	0.537
Sexual/Vaginal functioning	80.6±20.6	85.4±16.8	0.077
Sexual worry	26.6±28.0	22.0±24.9	0.214
FSFI			
Desire	3.0±1.0	2.9±1.1	0.617
Arousal	3.5±1.3	3.2±1.6	0.232
Lubrication	4.1±1.7	4.0±1.9	0.696
Orgasm	3.9±1.5	3.7±1.8	0.356
Satisfaction	4.0±0.7	3.9±1.2	0.363
Pain	4.5±1.7	4.1±2.0	0.214
Total	23.0±6.6	21.9±8.2	0.300

Values are presented as mean±standard deviation. EORTC QLQ-C30, The European Organization for Research and Treatment of Cancer Quality of Life Questionnaire-Core 30; EORTC QLQ-CX24, The European Organization for Research and Treatment of Cancer Quality of Life Questionnaire-Cervical Cancer Module; FSFI, Female Sexual Function Index.

DSM-5, the Japanese researchers who developed the Japanese version of the FSFI proposed sexual activity within 3 months [20]. Accordingly, women who had sexual activity within 3 months, rather than 6 months, were enrolled in this study.

The QoL of cervical cancer survivors has improved over time [21]. A similar level of QoL between cervical cancer survivors and control groups may be explained by the relatively long interval from the diagnosis to the survey (median, 45 months), as well as by the survivors' disease-free state.

Body image is a part of the QoL. The majority of rectal, gastric, and head and neck cancer patients have shown a deteriorated body image [22,23]. The aggravated body image during cancer treatment may recover gradually after treatment. However, body image is often not fully recovered, even 5 years after treatment [23]. Body image is also known to be affected by socioeconomic status.

Interestingly, the cervical cancer survivors in the current study showed better body image than women in the control group (73.1 vs. 79.6, $p=0.036$). Possible reasons for this find-

Table 4. Summary of studies using the Female Sexual Function Index (FSFI) to assess sexuality in cervical cancer

Study (year)	Study period	Inclusion criteria	Exclusion criteria	No. of participants	Age at study	FSFI total score	Years since diagnosis	Measurement	Result
Frumovitz et al. (2005) [14], USA	NA	Stage I	Combination of surgical and radiation therapies	RH (n=37)	43.6	25.1	7.5	FSFI, SF-12, BSI-18, menopause scale, A-DAS, CARES	Radiotherapy deteriorated sexual functioning
		Age < 55 yr at the treatment	CCRT	Radiotherapy (n=37)	46.9	17.1	7.0	-	63% of those sexually active before having cancer remained sexually active after treatment
		Treatment at least 5 yr previously	Disease recurrence	Control (n=40)	42.8	26.4	-	-	Sexual functioning and QoL did not differ significantly between surgery and control
		Disease-free state	Treated for other malignancies	-	-	-	-	-	-
		Surgery or radiotherapy	-	-	-	-	-	-	-
Carter et al. (2008) [13], USA	Feb 2004- Jan 2008	Stage I	NA	RT (n=30)	18-49	NA	NA	FSFI, FACT-CX, CES-D	At the 6-mo follow-up assessment, 70% of the women were sexually active
		Age 18-49 yr	-	-	-	-	-	-	Postoperative dyspareunia decreased over time
Serati et al. (2009) [15], Italy	Jan 2003- Dec 2007	Stage IB or IIA	Non-sexually active women	RH (n=38)	47 (24-71)	17.2	NA	FSFI	Radical hysterectomy deteriorated sexual functioning
		Received RH	Severe mental illness	Laparoscopy (n=20)	45.0 (24-69)	10.8	-	-	Laparoscopic surgery did not show any benefit on sexuality over the abdominal surgery
		-	Severe physical handicap	Laparotomy (n=18)	44.5 (25-68)	21.9	-	-	-
		-	Pregnant	Control (n=35)	45 (24-69)	30.7	-	-	-
Carter et al. (2010) [12], USA	Feb 2004- Jan 2008	Stage IA1-IB2	NA	RT (n=33)	32.6	Baseline vs. after 2 yr 17.4 vs. 21.9	NA	FSFI, FACT-CX, CES-D, IES	Sexual functioning and QoL did not differ significantly by surgical type
		Age 18-45 yr	-	RH (n=19)	37.6	Baseline vs. after 2 yr 15.7 vs. 22.6	-	-	-
Tsai et al. (2011) [17], Taiwan	NA	CIS or above age ≥ 20 yr	NA	CC (n=105)	54.3 (28-78)	NA	NA	FSFI	Older age (CI, 1.07-1.25)
		Sexual activity 3 yr before the diagnosis	-	-	-	-	-	-	Level of education (≤ 9th grade) (CI, 1.51-10.37)

Table 4. Continued

Study (year)	Study period	Inclusion criteria	Exclusion criteria	No. of participants	Age at study	FSFI total score	Years since diagnosis	Measurement	Result
Song et al. (2012) [16], Korea	Aug 2010- Nov 2010	Stage IA-IB1	Refuse to participate in the survey	Conization (n=39)	34.7	30.7	2.5	FSFI	Radical trachelectomy and hysterectomy deteriorated sexual function than conization
		Age < 40 yr	Laparotomic approach	RT (n=18)	34.7	21.7	2.5	-	-
		Sexual activity preoperative	Oophorectomy with radical surgery	RH (n=24)	36.8	22.4	2.3	-	-
		-	Adjuvant radiation and / or chemotherapy	-	-	-	-	-	-
		-	Disease recurrence	-	-	-	-	-	-
Lee et al., Korea	Feb 2013- Jan 2014	Stage I-IV	Refuse to participate in the survey	CC (n=104)	47.0 (24-70)	23.0	3.7	FSFI, EORTC QLQ-C30, EORTC QLQ-CX24	Body image and lymphedema were more problematic in cervical cancer survivors
		Age ≥ 18 yr	-	Control (n=104)	48.9 (24-66)	21.9	-	-	Sexual functioning was not impaired
		Sexual activity within 3 mo	-	-	-	-	-	-	-
		Disease-free state	-	-	-	-	-	-	-

NA, not available; RH, radical hysterectomy; SF-12, Short Form 12; BSI, Brief Symptom Index-18; A-DAS, Abbreviated Dyadic Adjustment Scale; CARES, Cancer Rehabilitation Evaluation System; CCRT, concurrent chemoradiotherapy; QoL, quality of life; RT, radical trachelectomy; FACT, Functional Assessment of Cancer Therapy; CES-D, Center for Epidemiology Studies Depression; IES, Impact of Event Scale; CIS, carcinoma *in situ*; CC, cervical cancer; CI, confidence interval; EORTC QLQ-C30, The European Organization for Research and Treatment of Cancer Quality of Life Questionnaire-Core 30; EORTC QLQ-CX24, EORTC QLQ-Cervical Cancer Module.

ing may be as follows. (1) Through the three concise questions for measurement of body image, quantitative or detailed measurement of body image was limited. (2) Korean women tend to be sensitive regarding their appearance and to have more body dissatisfaction. According to the previous study comparing body satisfaction across groups of college women from Korea, China, and the United States, the highest dissatisfaction was observed in Korean women [24]. In the Korea National Health and Nutrition Examination Survey, 2007-2010, a quarter of underweight adult Korean women (25.4%) have tried to control weight, and this attempt was influenced by distorted body image [25]. Therefore, even Korean women who are healthy and without disease may have poor body image. (3) Last, because of propensity score matching, socioeconomic status, known to be associated with altered body image, was not different between the two groups.

The symptoms of lymphedema were aggravated in cervical cancer survivors compared to the control group, as treatment for cervical cancer includes pelvic lymph node dissection as part of surgical treatment, chemotherapy, or radiotherapy in the pelvic area. After surgery and radiotherapy, a damaged lymphatic system is not fully recovered. The prolonged or persistent lymphedema-related symptoms have been also known to cause deterioration of the patients' QoL [21].

The current study had several limitations. Only women who had engaged in sexual activity within 3 months were included. Propensity score matching was used to make an adjustment for demographic characteristics of the cervical cancer survivors and control group. Inevitable issues, such as selection bias, might exist in this case-control study.

Conclusion

Compared with healthy women, sexual functioning was not impaired in cervical cancer survivors who showed no evidence of disease after primary treatment and engaging in sexual activity. However, in terms of QoL, lymphedema was more problematic in cervical cancer survivors than in the control group. Prospective cohort studies are warranted to confirm these findings.

Conflicts of Interest

Conflict of interest relevant to this article was not reported.

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