Aspects of Therapy for Cervical Cancer in Germany 2012 – Results from a Survey of German Gynaecological Hospitals

Aspekte der Therapie des Zervixkarzinoms in Deutschland 2012 – Ergebnisse einer Umfrage unter den deutschen Kliniken für Gynäkologie

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Key words
- staging of cervical cancer
- radiochemotherapy for cervical cancer
- radical hysterectomy
- questionnaire on treatment situation
- therapeutic concepts
- stage-dependent therapy

Schlüsselwörter
- Staging bei Zervixkarzinom
- Radiochemotherapie bei Zervixkarzinom
- radikale Hysterektomie
- Umfrage zur Behandlungssituation
- Therapiekonzepte
- stadienabhängige Therapie

Abstract

Introduction: In spite of the existence of guidelines and international recommendations, many aspects in the diagnosis, therapy and follow-up of patients with cervical cancer are not based on validated data. A broad spectrum of different opinions and procedures concerning the therapy for patients with cervical cancer is under controversial discussion by the responsible gynaecologists in German hospitals.

Methods: The present study is intended to picture the current treatment situation for cervical cancer in Germany. For this purpose a specially developed questionnaire with questions divided into 19 subsections was sent to all 688 gynaecological hospitals in Germany.

Results: The response rate to the questionnaire was 34.91% of the hospitals treated between 0 and 25 patients with cervical cancer per year. 7.5% treated between 26 and 50 and 1.4% of the hospitals more than 50 patients per year. The bimanual examination was the most frequently used staging method (98%); PET-CT was the least used method (2.3%). Interestingly 48% of the hospitals used surgical staging. The great majority of the hospitals (71%) used abdominal radical hysterectomy (Wertheim-Meigs operation) to treat their patients. TMMR via laparotomy was used by 13%. 16% of the hospitals performed laparoscopic or robot-assisted radical hysterectomies. The sentinel concept was hardly used even in the early stages. It must be emphasised that in 74% of the hospitals radical hysterectomies were performed even in cases with positive pelvic lymph nodes and in 43% also in cases with positive paraaortic lymph nodes. The therapy of choice for FIGO IIB cancers is primary radiochemotherapy (RCTX) in 21% of the hospitals; operative staging followed by radiochemotherapy in 24% and treatment by radical hysterectomy followed by adjuvant RCTX was employed in this sit-

Zusammenfassung


Methoden: Die vorliegende Studie sollte die derzeitige Behandlungssituation des Zervixkarzinoms in Deutschland abbilden. Dazu wurde ein speziell dafür entwickelter Fragebogen, der Fragen zu 19 Teilbereichen abdeckte, an alle 688 deutschen Kliniken für Gynäkologie versandt.

Resultate: Die Rücklaufquote der Fragebögen war 34.91% der Kliniken behandelten zwischen 0 und 25 Patientinnen/Jahr. 7,5% zwischen 26 und 50 und 1,4% der Kliniken mehr als 50 Patientinnen/Jahr. Die bimanuelle Untersuchung war die am häufigsten eingesetzte Stagingmethode (98%), das PET-CT wurde zum Staging am seltensten beansprucht (2,3%). Interessanterweise setzten 48% der Kliniken ein operatives Staging um. Die große Mehrheit der Kliniken (71%) erhielten die Patientinnen per abdominaler radikaler Hysterektomie (Wertheim-Meigs-Operation). Die TMMR per Laparotomie wurde von 13% eingesetzt, 16% der Kliniken operierten laparoskopisch oder roboterassistiert per radikaler Hysterektomie. Das Sentinel-Konzept wurde auch bei Frühstadien kaum verwandt. Hervorzuheben ist, dass in 74% der Kliniken radikale Hysterektomien auch bei positiven pelvinen Lymphknoten und in 43% auch bei positiven paraaortalen Lymphknoten durchgeführt wurden. Die Therapie der Wahl bei FIGO-IIB-Karzinomen war die primäre Radiochemotherapy (RCTX) in 21% der Kliniken, ein operatives Staging gefolgt von einer Radiochemotherapy in 24% und
A standardisation of therapy is needed. An update of national guidelines could help to achieve more transparency and a standardisation of treatment for patients with cervical cancer in Germany. A broad spectrum of differing staging and therapy concepts is in use for patients with cervical cancer.

## Abbreviations

- **FIGO**: Fédération Internationale de Gynécologie et d’Obstétrique
- **GOG**: Gynecologic Oncology Group
- **NCCN**: National Comprehensive Cancer Network
- **OS**: Overall survival
- **PFS**: Progression-free survival
- **RCTX**: Radiochemotherapy
- **RH**: Radical hysterectomy
- **TMMR**: Total mesometrial resection
- **VALRH**: Vaginal-assisted laparoscopic radical hysterectomy
- **LARVH**: Laparoscopic assisted radical vaginal hysterectomy

## Introduction

The treatment of patients with invasive cervical cancer in Germany should be oriented on the currently valid AGO guidelines [1]. These guidelines, as well as international recommendations (e.g., NCCN) are based on prospective and retrospective monocentric data, since only few prospective randomised studies are available [2]. Even for the radical hysterectomy (RH) which has been practiced for more than one hundred years there are today still no randomised studies comparing the various surgical techniques [99]. Also the classification of radicality by various groups has not led to a standardisation of the surgical methods [3,4]. In addition, because of the declining incidence of invasive cervical cancer – with 4880 newly diagnosed cases and 1600 deaths in Germany, fewer and fewer patients per hospital are being treated which will lead to appreciable problems with regard to experience and training [5].

On the other hand novel diagnostic [100], surgical and radiooncological procedures are being introduced and need to be evaluated [101]. The staging system for cervical cancer according to FIGO is today based solely on gynaecological examinations and cystoscopy/rectoscopy. Although MRI and CT have clear limitations for the staging of women with cervical cancer they are being employed more and more [6]. Whether or not PET-CT can provide improved data for staging is currently not clear [7–10]. Also the oncological relevance of surgical staging is still being discussed controversially and is being checked in the Uterus-11 study of AGO [11–13]. The sentinel concept appears to be applicable with highly sensitive and detection rates for tumours < 2 cm; even so there are reservations due to the low prevalence of positive lymph nodes in cases of tumours in stages IA1–IB1 [14,15].

More recently the laparoscopic assisted RH [26–28] and TMMR [29] have been developed and provided highly promising oncological results. Nerve-sparing operations also help to markedly reduce the postoperative morbidity. Also from the radiotherapy side, considerable improvements in the treatment of patients with cervical cancer have been achieved. Beside the usual combination of chemotherapy and radiation (radiochemotherapy = RCTX), which achieves a significant improvement in survival for the patient [30–34], it is above all the use of innovative techniques of radiation that clearly reduces the rate of side effects caused by the therapy [35,36]. The sole randomised study to compare radical hysterectomy with radiotherapy alone by Landoni in 1997 must now be considered as outdated with regard to the radiotherapy-associated side effects [37].

In the most recent publications on RH more than 50% of the patients receive a postoperative adjuvant radiochemotherapy, which is often associated with an increased morbidity [38,39]. The criteria for an adjuvant therapy are applied in widely different ways in spite of the GOG-109 data [40–43]. Furthermore, different algorithms are used in the follow-up [44–46]. The large number of open and controversial topics prompted our group to develop a questionnaire to assess the current status in the treatment of women with cervical cancer, even though we do not make any claims to completeness for this compilation.

## Material and Methods

Between February 2012 and June 2012 a questionnaire comprising 19 topic complexes was sent to the heads of all gynaecological hospitals nationwide in Germany (Questionnaire – Fig. 1). The data were evaluated using SPSS. Percentage values each refer to the number of usable replies to the particular topic.

## Results

### General

From 688 hospitals to which the questionnaire was sent 234 (34%) answered, of these 26 were university hospitals (11%), 113 were medical centres (49%) and 93 were regional and general hospitals (40%). 28% (n = 63) of these hospitals are certified as centres for gynaecological oncology. All hospitals were subdivided into 3 groups according to the number of patients with cervical cancer treated per year: 1: 0–25 patients, 2: 26–50, 3: more than 50. The university hospitals could be assigned as fol-
Questionnaire on therapy for cervical cancer 2012
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(It should not take more than 15 minutes to answer all questions; solely question 13 is somewhat more extensive but concerns an extremely important and controversially discussed topic.)

### General

1. What type of hospital do you work in?
   - University hospital
   - Medical centre/specialist hospital
   - General regional hospital

2. Is your department certified as a centre for gynaecological oncology?
   - Yes
   - No

3. How many patients with invasive cervical cancer of the following FIGO stages did you treat last year (2011)?
   - IA1
   - IA2
   - IB1
   - IB2
   - II A
   - II B
   - III A/III B
   - IV A/IV B

Have the answers to this question been taken from a register or are they estimated?

### Staging

4. Which imaging/staging examinations do you usually perform from stage IB1 (please check – multiple answers are possible)?
   - Bimanual exam
   - Cystoscopy
   - Rectoscopy
   - CT
   - MRI
   - PET-CT
   - Surgical staging

5. If you regularly perform surgical staging which operative approach do you use in the majority of cases (please name only one procedure)?
   - Open transperitoneal
   - Open retroperitoneal
   - Laparoscopic transperitoneal
   - Laparoscopic extraperitoneal
   - Robot-assisted transperitoneal
   - Robot-assisted retroperitoneal

6. Do you perform sentinel lymph node procedures in women with cervical cancer (please give only one answer)?
   - In all stages routinely
   - Only for tumours < 2 cm, but then for all patients
   - Only within studies
   - Only if requested by the patient

### Surgical therapy

7. Which type of radical hysterectomy do you perform mainly?
   - Please check only one or two boxes, if two please give percentages for each of the two methods.
   - Classic Wertheim operation (open surgery)
   - TMMR (open surgery)
   - Laparoscopic-assisted radical vaginal (LARVH) or vaginal-assisted laparoscopic radical hysterectomy (VALRH)
   - Total laparoscopic radical hysterectomy (TLRH)
   - Robot-assisted radical hysterectomy (RRH)
   - Robot-assisted TMMR
   - Others (please name them)

8. Do you as a general rule perform an intraoperative immediate section for microscopy of the resected lymph nodes (please check only one box)?
   - Yes
   - No
   - Only for suspicious enlargement of individual lymph nodes
   - As a rule two-stage procedure (1st OP lymphonodectomy, radical hysterectomy as 2nd OP only when free of tumour)

9. What is your usual procedure for positive pelvic lymph nodes in cases of operable local tumours (please check only one box)?
   - Discontinue the radical HE, paraaortal lymphonodectomy and primary radiochemotherapy (RCTX)
   - Continue the radical HE, paraaortal lymphonodectomy and adjuvant RCTX
   - Continue the radical HE, paraaortal lymphonodectomy and adjuvant chemotherapy
   - Discontinue the radical HE and primary radiochemotherapy, including the paraaortic region (extended field)

10. What is your usual procedure for positive paraaortic lymph nodes in cases of operable local tumours (please check only one box)?
    - Discontinue the radical HE and primary radiochemotherapy, including the paraaortic region (extended field)
    - Discontinue the radical HE and palliative chemotherapy
    - Continue the radical HE and adjuvant radiochemotherapy
    - Continue the radical HE and adjuvant chemotherapy

### Stage-specific questions

11. In stage IA1 L0 do you perform a sentinel lymphonodectomy in addition to re-conisation/simple hysterectomy?
    - Yes
    - No

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Fig. 1 Questionnaire. (continued next page)
12. For a patient with cervical cancer in stage IB1 < 2 cm who still wants to have a child, do you (please check only one box)

- always perform a radical HE because you do not approve of trachelectomy.
- refer the patient to a centre experienced in trachelectomy.
- always perform a radical trachelectomy.

If yes:

- For positive nodes; R1/R2 resection and/or tumour invasion of the parametrium
- Only grade 3 (G3)
- Only tumour size > 4 cm
- Only age < 40 years
- Only by invasion of the lymphovascular space (L1)
- Only adenocarcinoma as histological type

For 2 risk factors:

- Combination G3 + age < 40
- Combination G3 + tumour > 4 cm
- Combination G3 + L1
- Combination G3 and adenocarcinoma
- Combination tumour > 4 cm and age < 40
- Combination tumour > 4 cm and L1
- Combination tumour > 4 cm and adenocarcinoma
- Combination age < 40 and L1
- Combination age < 40 and adenocarcinoma
- Combination L1 and adenocarcinoma

For 3 risk factors:

- Combination G3 + tumour > 4 cm + age < 40
- Combination G3 + tumour > 4 cm + L1
- Combination G3 + tumour > 4 cm + adenocarcinoma
- Combination G3 + age < 40 + L1
- Combination G3 + age < 40 + adenocarcinoma
- Combination G3 + L1 + adenocarcinoma
- Combination tumour > 4 cm + age < 40 + L1
- Combination tumour > 4 cm + age < 40 + adenocarcinoma
- Combination tumour > 4 cm + L1 + adenocarcinoma
- Combination age < 40 + L1 + adenocarcinoma

For 4 risk factors:

- Combination G3 + tumour > 4 cm + age < 40 + L1
- Combination G3 + tumour > 4 cm + age < 40 + adenocarcinoma
- Combination G3 + tumour > 4 cm + L1 + adenocarcinoma
- Combination tumour > 4 cm + age < 40 + L1 + adenocarcinoma
- Combination G3 + age < 40 + adenocarcinoma + L1

13. When do you, as surgeon, recommend adjuvant radiochemotherapy after radical hysterectomy in stages IB1/IB2 (multiple answers possible):

- For positive nodes; R1/R2 resection and/or tumour invasion of the parametrium
- If N0, R0 and stage pT1b1 or pT1B2, do you recommend adjuvant RCT in the presence of the following risk factors (please check as appropriate):

  - Only grade 3 (G3)
  - Only tumour size > 4 cm
  - Only age < 40 years
  - Only by invasion of the lymphovascular space (L1)
  - Only adenocarcinoma as histological type

  For 2 risk factors:

  - Combination G3 + age < 40
  - Combination G3 + tumour > 4 cm
  - Combination G3 + L1
  - Combination G3 and adenocarcinoma
  - Combination tumour > 4 cm and age < 40
  - Combination tumour > 4 cm and L1
  - Combination tumour > 4 cm and adenocarcinoma
  - Combination age < 40 and L1
  - Combination age < 40 and adenocarcinoma
  - Combination L1 and adenocarcinoma

  For 3 risk factors:

  - Combination G3 + tumour > 4 cm + age < 40
  - Combination G3 + tumour > 4 cm + L1
  - Combination G3 + tumour > 4 cm + adenocarcinoma
  - Combination G3 + age < 40 + L1
  - Combination G3 + age < 40 + adenocarcinoma
  - Combination G3 + L1 + adenocarcinoma
  - Combination tumour > 4 cm + age < 40 + L1
  - Combination tumour > 4 cm + age < 40 + adenocarcinoma
  - Combination tumour > 4 cm + L1 + adenocarcinoma
  - Combination age < 40 + L1 + adenocarcinoma

  For 4 risk factors:

  - Combination G3 + tumour > 4 cm + age < 40 + L1
  - Combination G3 + tumour > 4 cm + age < 40 + adenocarcinoma
  - Combination G3 + tumour > 4 cm + L1 + adenocarcinoma
  - Combination tumour > 4 cm + age < 40 + L1 + adenocarcinoma
  - Combination G3 + age < 40 + adenocarcinoma + L1

14. Do you as a general rule initiate a neoadjuvant therapy in stage IB2 or IIB prior to a planned radical hysterectomy (please check only one box)?

- Yes, a neoadjuvant chemotherapy
- Yes, a neoadjuvant radiochemotherapy
- No
- I always refer these stages to primary radiochemotherapy.

15. What is your therapy for the majority of patients in FIGO stage IIB (please check only one box)?

- Primary radiochemotherapy
- Surgical staging and subsequent primary radiochemotherapy
- Radial hysterectomy and adjuvant radiochemotherapy
- Others (please specify)

16. Which type of staging do you perform in FIGO stages IIB/IIIB?

- Clinical staging before radiochemotherapy
- Surgical staging before radiochemotherapy

17. What is your therapy in stage IVA (multiple answers possible)?

- Always primary radiochemotherapy
- Always primary exenteration
- Primary exenteration only for urogenital or intestinal fistula
- Individual decision

18. What examinations do you perform on patients after primary radiochemotherapy (without prior radical hysterectomy) in the follow-up period (multiple answers possible)?

- Clinical examination
- Renal ultrasonography
- Vaginal ultrasonography
- Determination of tumour markers
- Pelvic MRI
- PET-CT
- Cervix abrasion
- PAP smear

19. In your opinion when is an operation indicated after primary radiochemotherapy (without radical hysterectomy) (please check only one box)?

- There is no indication because there is no advantage in survival.
- Secondary hysterectomy is always performed.
- A secondary hysterectomy is performed only on suspicion of a persisting tumour (e.g. in the abrasion).
- On suspicion of a local recurrence a secondary exenteration is always performed.

Fig. 1 Questionnaire. (continued)
allows to the groups 1, 2 and 3, respectively, 14 (61%), 6 (26%) and 3 (13%). Three university hospitals did not give an answer to the number of treated patients per year. For medical centres and regional/general hospitals the assignments were as follows: 1 = 94%, 2 = 6%, 3 = 0% and, respectively, 1 = 95.5%, 2 = 4.5% and 3 = 00%. Altogether 91% of all patients with cervical cancer were treated in a hospital that handles less than 26 such patients per year.

**Staging**

In practically all hospitals, staging from stage IB1 was done according to FIGO by bimanual examination (98%). Cystoscopy and rectoscopy were done in 73% and, respectively, 70% of the gynaecological departments. 44% of the hospitals used CT routinely, 52% used MRI, whereas in only 2% PET-CT was used for staging (Fig. 2). Surgical staging was preferred in 48% of the responding hospitals. Here the following approaches for surgical staging were used: open transperitoneal in 41%, open retroperitoneal in 9%, laparoscopic transperitoneal in 47%, laparoscopic extraperitoneal in 1% and robot-assisted transperitoneal in 2%. The sentinel concept was not utilised in 43%; 4% of the hospitals performed sentinel lymphadenectomy in all patients with tumours ≤2 cm, 22% only within studies and 22% only if requested by the patient. In 9 gynaecological departments (4%) the sentinel concept is applied to all tumour stages (Fig. 3). Almost all hospitals (96%) reject the sentinel concept as an addition to conisation/simple hysterectomy for patients with a cervical cancer stage pT1a1.

**Therapy**

In the great majority of the gynaecological hospitals in Germany the classic open Wertheim operation is used (71.5%). Other procedures for RH with markedly lower usage are represented by laparoscopic-assisted vaginal or vaginal-assisted laparoscopic RH in 4%, total laparoscopic RH in 10%, TMMR in 13%, robot-assisted RH in 1% and robot-assisted TMMR in 0.5%. In 32% of the hospitals several procedures were regularly applied (Fig. 4). An intraoperative immediate section of all resected lymph nodes is offered in 57% of the hospitals, in 26% only for suspiciously enlarged lymph nodes. In 2% of the hospitals – in the sense of a two-stage procedure – a lymphadenectomy is performed first followed by RH only after confirmation of tumour-free lymph nodes (Fig. 5).

In the cases with the identification of positive pelvic lymph nodes in patients with a local operable tumour the RH is discontinued in 16% of the hospitals and, after performance of a paraaortic lymphadenectomy, a primary radiochemotherapy is initiated. 74% of the physicians continue the RH (including a paraaortic lymphadenectomy) and recommend adjuvant RCTX or adjuvant chemotherapy (10%) (Fig. 6). Upon identification of positive lymph nodes.
paraaortic lymph nodes of a locally operable cervical carcinoma, the operation is terminated in 50% of the patients. In 43% the RH is continued in this clinical situation and an adjuvant RCTX is subsequently carried out, in 7% adjuvant chemotherapy (Fig. 7). Young patients with a still unfulfilled desire to have children and a cervical cancer of less than 2 cm in size are referred in 80% to a centre with expertise in radical trachelectomy. 17% of the hospitals perform uterus-sparing operations in house, usually as a radical vaginal trachelectomy (14%). Three percent of the gynaecological departments reject a trachelectomy and always prefer a RH for this constellation.

In FIGO stage IB2 neoadjuvant chemotherapy is applied in 6% of the clinics prior to the planned RH and in 7% adjuvant chemotherapy (Fig. 8). For patients in stage IIIA/IIIB most physicians consider a clinical staging to be sufficient prior to a primary RCTX (69%), merely 31% vote for a surgical staging before commencing therapy. In stage IVA of cervical cancer individual decisions are preferred in most hospitals (Fig. 9).

The indications for adjuvant RCTX after RH form a very heterogeneous pattern. Whereas all hospitals (100%) favour an adjuvant therapy in cases with positive lymph nodes, invasion of the...
parametrium and an R1/R2 resection, the presence of the risk factors stage 3, tumour size > 4 cm, age < 40 years, adenocarcinoma as histological type and invasion of the lymphovascular space (L1) alone or in combination is considered in widely differing ways (Fig. 10 to 13). In the case of one of the above-mentioned risk factors 15-68% of the hospitals recommend an adjuvant RCTX, in the case of 2 factors 42–88%, for 3 factors 68–97% and for 4 factors 90–97%.

Follow-up
In the course of follow-up after primary radiochemotherapy in practically all hospitals a gynaecological examination (99%) and vaginal ultrasonography (99%) are performed. In 90% of the hospitals in addition ultrasonography of the kidneys is performed, a PAP smear in 75%, a cervical abrasion in 28%, a pelvic MRI in 41%, a tumour marker determination in 28% and/or a PET-CT in 4% (Fig. 14). On identification of a local persisting tumour/intrauterine recurrence after primary radiochemotherapy only 28% of the hospitals consider a secondary hysterectomy as not being indicated, since it is not associated with a benefit in terms of survival. In contrast 7% of the hospitals always perform a secondary hysterectomy, 57% do so only when tumour persistence is detected. Eight percent consider a primary exenteration to be indicated in this clinical situation (Fig. 15).
In 2006 the FIGO committee again decided to leave the staging of cervical cancer as a purely clinical process [49]. All German hospitals confer but only in 70% are the cystoscopic and rectoscopic examinations required by FIGO actually carried out. The routine use of CT and MRI in 44% and 52% reflect the widely differing data with regard to the sensitivity and specificity of these two imaging procedures found in the literature [50–54]. PET-CT, the costs of which are not reimbursed in Germany, is of no significance as a staging method in correspondence with its low sensitivity according to Ramirez et al. and LeBlanc et al. [7, 8, 10, 55]. 48% of the hospitals consider surgical staging to be a valid alternative to imaging and to clinical staging, although as yet no randomised clinical trial has been able to confirm the oncological advantages seen in retrospective and prospective studies [56, 57]. The successfully recruiting prospective randomised Uterus-11 study of AGO on the value of surgical staging in patients with cervical cancers of stages IIB–IVA should provide pathbreaking answers to this matter. Why 41% of the hospitals would still choose an open transperitoneal approach for surgical staging is not clear, since it is just the formation of postoperative adhesions before an RCTX that is the main argument against surgical staging.

The significance of the sentinel concept in cervical cancer is evaluated very differently nationwide in Germany. The results of the Uterus-3 trial of AGO published by Altgassen et al. still represent the largest investigation of this topic worldwide. These results are suggestive that SLN can achieve a sufficiently high detection and sensitivity in patients with a tumour size < 2 cm [14, 58]. Merely 9% of all hospitals in Germany employ the sentinel concept routinely for carcinomas ≤ 2 cm. 44% of the hospitals would apply the concept if requested by the patient or in clinical trials. The international trend to define patient collectives who would benefit from less radical surgery with equal oncological efficacy is only implemented in a few hospitals [59]. In addition, the advantages associated with the SLN technique such as the discovery of rare lymph drainage pathways [60] or the detection of micrometastasis [61] are not exploited. Lymph node metastasis in stage IA1 is rare [15, 62]. This supports the opinion of the majority of hospitals in Germany not to use SLN in this stage. It is, however, possible that by this means one could identify with minimal morbidity just those few patients with early lymph node metastasis, whose prognosis is otherwise rather poor [63–65]. The uterus-sparing operation in cases of early invasive cervical cancer is accepted in 97% of the gynaecological departments on account of its excellent oncological and reproductive results [66–68]. In Germany this is almost exclusively performed as rad-

### Discussion

To what extent national guidelines are applied in the therapy for cervical cancer can only be determined on the basis of characteristic reimbursement data or with the help of questionnaires [47, 48]. Whereas the former is only meaningful when a separate character can be coded for every possible treatment modality, the latter is dependent on a high participation. The response rate of 34% achieved with the present questionnaire is adequate and also allows representative conclusions on the composition of the hospitals. The fact that within just a few years 28% of all hospitals have attained certification as oncological centres is indicative of an increased awareness of quality in the treatment of patients with gynaecological cancers. In spite of the gratifying decline in the incidence of these cancers, the number of patients with cervical cancer treated per hospital must be considered very critical to all hospitals in Germany, at most 1000 patients are treated in all stages IA1 to IVA are collected together. If this is extrapolated to all hospitals in Germany, at most 1000 patients are treated in hospitals with extensive expertise and only very few in high-volume hospitals. The great majority of women are treated in hospitals with low case numbers which, in turn, will lead to increasing problems in further training and the establishment of modern therapeutic concepts. The centralisation of therapy for patients with invasive cervical cancer should be a target for the future.

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**Fig. 14** Examinations performed in the course of follow-up (multiple answers possible).

**Fig. 15** Indication for a secondary operation after primary radiochemotherapy for cervical cancer.
cical vaginal trachelectomy in contrast to other countries where abdominal radical trachelectomy also has a high relevance [47]. Various procedures are in use in Germany for radical hysterectomy, in some cases even in one hospital. A positive development is that innovative procedures with excellent oncological outcome, such as TMMR by the Leipzig group [29] or VALRH by the Berlin group [25] are proportionally well represented. Thus we can hope that in the years to come the monocentric data will be confirmed by multicentric results. Less easily understandable are the answers of German hospitals in 74% and 43%, respectively, to continue with RH in the presence of positive pelvic or paraaortic lymph nodes. This could be based on the fact that 43% of the hospitals do not perform intraoperative immediate sections or on economic reasons, especially as practically all hospitals recommend an adjuvant RCTX in the presence of positive lymph nodes. The recommendation of GOG 109 study for an adjuvant therapy in cases of N1, parametrial infiltration and an R1/R2 resection are almost unanimously implemented in German hospitals [40,69]. The comments of the gynaecological departments about an adjuvant therapy after RH in the presence of only one or several intermediate risk factors clearly reflect the, also internationally, much too high rate of trimodal therapy, the oncological utility of which has not yet been clarified for many combinations of these risk factors [41,42]. Marnitz et al. have impressively demonstrated that the rate of adjuvant RCTX can be minimised to 10% with the help of laparoscopic staging and with knowledge of the historical results of preoperative biopsies/conisation; in this way the significantly elevated morbidity of RH and RCTX can be avoided [39]. For this complex of topics, in particular, a revised German guideline with clear unambiguous statements is needed.

The optimal therapy for patients in FIGO stages IB2 and IIB has been a subject of controversial discussion for many years, but is still not clear as is reflected in the results of this and earlier questionnaires [98]. The spectrum of nationally as well as internationally applied therapies encompasses primary radical hysterectomy ± adjuvant radiochemotherapy, neoadjuvant (radio)chemotherapy followed by radical hysterectomy, primary radiochemotherapy (RCTX) or TMMR. In the not yet closed prospective randomised international EORTC study 55994, neoadjuvant chemotherapy (RCTX) or TMMR. In the not yet closed prospective randomised international EORTC study 55994, neoadjuvant chemotherapy followed by radical hysterectomy is being compared in the presence of high metabolic activity and as is demonstrated by Balleyguier et al. [84], accordingly the moderate use in 41% of the hospitals is justified. PET-CT is associated with a high rate of false negative and false positive findings in the diagnosis of recurrences as has been demonstrated by Chung et al. in 276 patients [85], whereas in other studies it has revealed a good correlation between complete metabolic response and survival [86]. Further studies on the value of PET-CT in the follow-up are needed. The determination of the tumour markers SCC and CEA is undertaken in 28% of the German hospitals [46,87], even when the available data suggest that the determination of these markers is not reliable [46,87].

Routine taken cytological samples – as is done in 75% of the gynaecological departments – should no longer be considered as the sole follow-up examination on account of the low detection rate for recurrences of between 0 and 17% [88–91,97]. According to Nijhuis et al. the combination of examinations under anaesthesia with biopsy sampling/cervical abrasion can detect or exclude a local persisting tumour after primary RCTX with a high probability [91]. This method should thus be employed more often than its current implementation in merely 28% of the hospitals. The wide spectrum of answers with regard to secondary operation in patients after primary RCTX for cervical cancer reflects the lack of evidence on this topic. While Motton et al. did not observe any increased rate of complications, Classe et al. and Colombo et al. recorded complications in 26–48% of their secondary operations. To what extent a simple hysterectomy is oncologically more meaningful than a radical hysterectomy or an exenteration also remains to be clarified. None of the studies could show any advantages with regard to overall survival, merely local control or, respectively, PFS were improved [91–97]. A randomised study of surgical strategies for persisting tumours is ethically not possible. Of course, the results presented here have their limitations. The design and length of the questionnaire did not allow the inclusion of all interesting questions and all possible answers. Also the fact that only a good third of the hospitals responded means that the results are still representative, although a higher participation would have been highly desirable.

**Conclusion**

The results of this questionnaire on the therapy for cervical cancer in Germany in 2012 reveal on the one hand that innovative concepts (laparoscopic procedures for RH, TMMR, surgical staging) have found acceptance in German hospitals while, on the other hand, long established treatment concepts (RH in case of
positive pelvic and/or paraaortic lymph nodes, therapy in FIGO stage IIB, recommendation for adjuvant therapy) are being retained. Randomised prospective studies should be put into practice. The lack of evidence for many questions leaves room for a broad spectrum of opinions and treatment pathways. The future new version of the S3 guidelines on the diagnostics and therapy for patients with cervical cancer should thus include unambiguous statements for many of these aspects.

Acknowledgements ▼
The authors are extremely grateful to all colleagues who returned the questionnaires and provided helpful suggestions and support.

Conflicts of Interest ▼
None.

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