

Trends in prognostic factors and treatment of invasive cervical cancer patients over a 16-year-period (1998-2013): a population-based analysis

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Tumorregister München (TRM)

Introduction

The objective was to identify trends in treatment and outcome of invasive cervical cancer in a population-based setting.

Methods

The Munich Cancer Registry (MCR) is the population based clinical cancer registry of Upper Bavaria and a part of Lower Bavaria (Southern Germany).

Its catchment area has increased from 2.3 million inhabitants to 3.8 million in 2002 and to 4.6 million in 2007 (meanwhile 4.7 million).

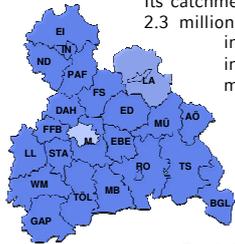


Fig. 1: Catchment area of the Munich Cancer Registry (MCR)

3,246 patients with invasive cervical cancer diagnosed between 1998 and 2013 in the catchment area of the Munich Cancer Registry (MCR) were analysed. Trends in prognostic factors and treatment were examined by comparing patients diagnosed within the years 1998-2008 (n=2,108) and 2009-2013 (n=1,138).

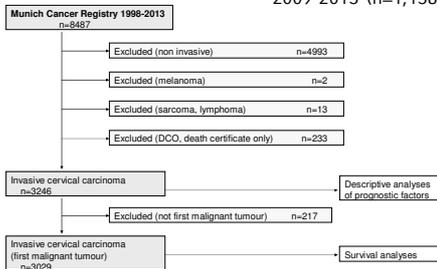


Fig 2.: Flow chart of cervical cancer patients

Results

Prognostic factors

The median age at diagnosis of 50.0 years (p=0.918) and grading with 46.5% G3-tumours (p=0.396) did not change significantly over time.

Table 1: Patients' and tumour characteristics

	1998-2008 n=2108		2009-2013 n=1138		Total n=3246		p
Age	Mean / Median	52.9 / 49.6	52.9 / 50.5	52.9 / 50.0	52.9 / 50.0	0.918	
<40	n	506	246	21.6	752	23.2	0.264
40-49	n	567	26.9	306	26.9	873	26.9
50-59	n	382	18.1	234	20.6	616	19.0
60-69	n	291	13.8	169	14.9	460	14.2
≥70	n	362	17.2	183	16.1	545	16.8
Lymph node involvement	n	%	n	%	n	%	
N0	896	67.0	481	61.1	1377	64.8	0.006
N+	441	33.0	306	38.9	747	35.2	
Missing	771	36.6	351	30.8	1122	34.6	
Grading	n	%	n	%	n	%	
G1	115	6.1	79	7.4	194	6.6	0.396
G2	887	47.2	496	46.5	1383	47.0	
G3/G4	878	46.7	491	46.1	1369	46.5	
Missing / GX	228	10.8	72	6.3	300	9.2	
pT/cT	n	%	n	%	n	%	
1A	363	19.3	205	19.0	568	19.2	<0.001
1B	745	39.6	382	35.4	1127	38.1	
2A	77	4.1	45	4.2	122	4.1	
2B	291	15.5	168	15.6	459	15.5	
3	161	8.6	78	7.2	239	8.1	
4	72	3.8	38	3.5	110	3.7	
M1	173	9.2	162	15.0	335	11.3	
Missing	226	10.7	60	5.3	286	8.8	

Treatment

Treatment approaches did not alter (p=0.951), with 7.1% of the patients having conisation, 64.9% treated by surgery, 21.9% having radiochemotherapy alone. In lymph node surgery there was a slight increase in the use of sentinel lymph node biopsy from 2.4 to 10.1% (p<0.001). For patients with lymph node dissection the median number of examined lymph nodes increased from 28 to 33 (p<0.001).

Table 2: Treatment options based on time of diagnosis

	1998-2008 n=2108		2009-2013 n=1138		Total n=3246		p
Therapy	n	%	n	%	n	%	
Conisation	142	7.0	78	7.2	220	7.1	0.951
Surgery	821	40.7	430	39.9	1251	40.4	
Surgery+radiotherapy	495	24.5	263	24.4	758	24.5	
Radiotherapy	442	21.9	237	22.0	679	21.9	
Others	119	5.9	71	6.6	190	6.1	
Missing	89	4.2	59	5.2	148	4.6	
Surgery	n	%	n	%	n	%	
Residual tumour	n	%	n	%	n	%	
R0	840	85.7	594	91.7	1434	88.1	<0.001
R1	140	14.3	54	8.3	194	11.9	
Missing	336	25.5	45	6.5	381	19.0	
Sentinel surgery	n	%	n	%	n	%	
yes	32	2.4	70	10.1	102	5.1	<0.001
thereof positive	4	12.5	10	14.3	14	13.7	0.808
LN surgery (incl. sentinel)	n	%	n	%	n	%	
yes	965	73.3	547	78.9	1512	75.3	0.006
no	351	26.7	146	21.1	497	24.7	
Type of LN surgery (incl. sentinel)	n	%	n	%	n	%	
sentinel alone	6	0.5	32	4.6	38	1.9	<0.001
pelvic	451	34.3	194	28.0	645	32.1	
pelvic+paraort.	271	20.6	268	38.7	539	26.8	
LND nos.	237	18.0	53	7.7	290	14.4	
no LN surgery	351	26.7	146	21.1	497	24.7	
LND (without SLNB)	n=959	n=515	n=1474				
dissected LNs	Mean / Median	30.4 / 28.0	34.8 / 33.0	32.0 / 30.0			<0.001
n	n	%	n	%	n	%	
1 - 24	333	37.4	141	27.8	474	33.9	<0.001
> 24	558	62.6	366	72.2	924	66.1	
Missing	68	7.1	8	1.6	76	5.2	
positive LNs	n	%	n	%	n	%	
0	649	72.8	361	71.2	1010	72.3	0.602
1 - 2	123	13.8	80	15.8	203	14.5	
> 2	119	13.4	66	13.0	185	13.2	
Missing	68	7.1	8	1.6	76	5.2	

In surgery of the primary tumour, the percentage of patients with no residual tumour (R0) increased from 85.7% to 91.7% (p<0.001), and was 98.5% in pT/cT1a and 97.6% in pT/cT1b patients diagnosed 2009-2013 (not shown).

Table 3: Treatment based on pT/cT stage

pTN/cTN M	1998-2008 n=2108						2009-2013 n=1138					
	n	%	Coni*	Surg*	R(CT)	Surg* + R(CT)	n	%	Coni*	Surg*	R(CT)	Surg* + R(CT)
1A NO MO	110	5.8	6.4	85.5	1.8	0.9	44	4.1	6.8	88.6	4.6	0.0
1A N+ MO	2	0.1	0.0	0.0	50.5	50.5	1	0.1	0.0	0.0	100	0.0
1A NX MO	251	13.3	39.8	53.0	0.8	0.0	160	14.8	39.4	56.3	0.0	0.0
1B NO MO	558	29.7	2.3	70.6	21.9	1.6	293	27.2	0.7	71.7	22.5	1.4
1B N+ MO	111	5.9	0.0	21.6	73.0	4.5	57	5.3	0.0	24.6	66.7	8.8
1B NX MO	76	4.0	10.5	35.5	23.7	21.1	32	3.0	21.9	40.6	6.3	9.4
2A NO MO	36	1.9	0.0	50.0	47.2	0.0	23	2.1	0.0	39.1	47.8	4.4
2A N+ MO	34	1.8	0.0	26.5	70.6	2.9	18	1.7	0.0	11.1	66.7	22.2
2B NO MO	118	6.3	0.9	28.0	54.2	16.1	78	7.2	1.3	21.8	61.5	14.1
2B N+ MO	120	6.4	0.0	14.6	18.8	62.5	27	2.5	0.0	7.4	12.2	68.3
3 NO MO	26	1.4	0.0	7.7	7.7	80.8	26	2.4	0.0	0.0	26.9	73.1
3 N+ MO	48	2.6	0.0	14.6	18.8	62.5	27	2.5	0.0	7.4	12.2	68.3
2-3 NX MO	147	7.8	0.7	6.1	11.6	68.0	41	3.8	0.0	7.3	12.2	68.3
4 NO MO	72	3.8	0.0	6.9	11.1	68.1	38	3.5	0.0	7.9	10.5	73.7
M1	173	9.2	1.2	5.2	20.2	46.8	162	15.0	0.6	11.1	6.8	50.6
TX / Missing	226	10.7	4.4	15.5	8.4	38.1	60	5.3	1.7	5.0	5.0	28.3
Total	1882	100	7.0	41.8	25.3	18.9	1078	100	7.1	39.6	24.1	20.4

*Coni: Conisation, Surg: Surgery, includes trachelectomy, hysterectomy, wertheim surg. and exenteration R(CT): Radio(chemo)therapy Patients with other treatment options were excluded from the analysis

In pT/cT2 the number of patients who underwent surgery alone decreased, while surgery with adjuvant radiochemotherapy increased. The percentage of radiochemotherapy in patients with positive lymph node status increased from 23.6% to 29.6% (p=0.035) (not shown).

Survival analysis

Relative 5-year survival rates differed from 98.6% in stage pT/cT 1A to 11.2% in patients with a metastasis at time of diagnosis.

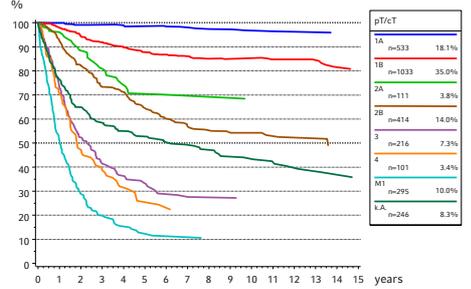


Fig 3.: Relative survival based on pT/cT stage
There was no difference in relative survival, time to locoregional recurrence or time to metastasis between patients diagnosed 1998-2009 and patients diagnosed 2009-2013.

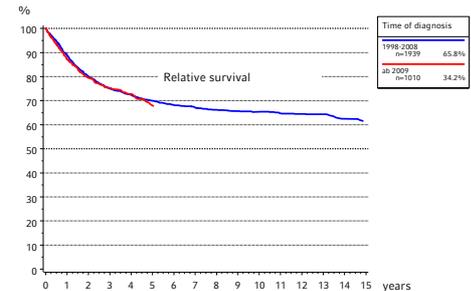


Fig 4.: Relative survival based on time of diagnosis

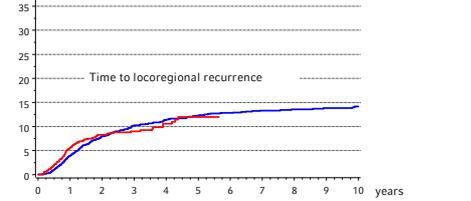


Fig 5.: Time to locoregional recurrence based on time of diagnosis (cumulative incidence)

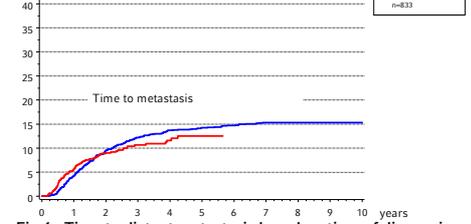


Fig 6.: Time to distant metastasis based on time of diagnosis (cumulative incidence)

Conclusions

There is a trend towards the increased use of radiochemotherapy, especially in advanced stages. In surgery there is a slight increase in the use of sentinel lymph node biopsy and an increase in R0-resections. Relative survival and time to locoregional recurrence and metastasis did not change over time.