Supplementary data

Supplementary Table 1. Time to recurrence according to recurrent pattern (n = 91).

Pattern of recurrence	n	Time to recurrence, months
		Median (range)
Vagina only	8	9.0 (4.2 – 51.4)
Pelvis	3	13.3 (5.8 – 71.5)
Distant only	58	17.1 (4.8 – 97.2)
Pelvis-distant	22	11.0 (3.2 – 74.4)
Total	91	13.3 (3.2 – 97.2)

Supplementary Table 2	. Time to failure for	patients with extravaginal	recurrences (n=83).
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Pattern of	S	urvivors	Noi	a volvo	
recurrence	No. of Pts	TTF	No. of Pts	TTF	<i>p</i> value
Pelvis	2	42.4 (13.3 – 71.5)	1	5.8	1.000
Distant	18	21.9 (5.1 – 97.2)	40	12.4 (4.8 – 77.9)	0.156
Pelvis-distant	4	13.7 (5.9 - 74.4)	18	11.0(3.2 - 35.1)	1.000
Total	24	19.7 (5.1 – 97.2)	59	12.2 (3.2 – 77.9)	0.025

TTF, Time to failure; pts, patients.

Supplementary Table 3. The relation between time to recurrence and SAR in patients with extra-vaginal recurrence (n = 83).

Time to recurrence	N	5-year SAR	p value
≤6 months	13	15.4	0.033
> 6 months	70	29.8	
≤9 months	21	9.5	< 0.001
> 9 months	62	33.7	
≤ 12 months	33	12.8	0.005
> 12 months	50	38.2	
≤ 15 months	44	14.6	0.004
> 15 months	39	42.2	
≤ 18 months	52	18.1	0.025
> 18 months	31	44.6	
\leq 24 months	62	25.8	0.272
> 24 months	21	33.6	
Total	83		
SAR, survival after recur	rence.		

Supplementary Table 4. Multivariate analysis of survival after recurrence (n = 91).

	HR of death (95% CI)	p value
Cervical invasion (+ vs -)	3.75 (1.87 – 7.52)	< 0.001
Histology (type 2 vs 1)	3.21 (1.57 – 6.55)	< 0.001
Cytology (+ vs -)	2.22 (1.01 – 4.91)	0.048
Recurrent site (multiple vs single)	1.84 (0.92 – 3.69)	0.084
CA-125 at recurrence ($> 35 \text{ U/mL}$ vs $\leq 35 \text{ U/mL}$)	2.06 (1.07 – 3.97)	0.030
Grade (3 vs 1-2)	1.44 (0.68 – 3.06)	0.344
ER and PR (ER(-) or PR(-) vs ER(+) & PR(+))	1.24 (0.60 – 2.58)	0.558
In-field failure of previous RT (yes vs no)	2.60 (0.97 – 6.95)	0.057

ER, estrogen receptor; PR, progesterone receptor; CI, confidence interval; RT, radiotherapy; HR, hazard ratio.

Alternative model with cervical invasion replaces FIGO stage

	Pre-	opPR+ and PR+ at	rec	Pre-opP	Pre-opPR+ and PR- at rec			Pre-opPR- and PR- at rec		
										value
	No. of Pts	5-y SAR	p value	No. of Pt	5-y SAR	p value	No. of Pts	5-y SAR	p value	
	(%)	(%)		(%)	(%)		(%)	(%)		
	17(100%)	94.1		16(100%)	42.2		18(100%)	23.8		0.001
Number of recurrent sites										
Solitary	11(65%)	100.0	0.176	10(63%)	36.0	0.873	6(33%)	55.6	0.008	0.070
Multiple	6(35%)	83.3		6(37%)	50.0		12(67%)	8.3		0.028
Pattern of recurrence										
Vagina	6(35%)	100.0	0.277	0	NA	0.054	0	NA	0.256	NA
Pelvis	2(12%)	All alive (20.2,		0	NA		0	NA		NA
		31.6 M)								
Distant	5(29%)	100.0		12(75%)	47.6		9(50%)	37.0		0.258
Pelvis-distant	4(24%)	75.0		4(25%)	25.0		9(50%)	11.1		0.102

Rec, recurrence; SAR, survival after recurrence; f/u, follow-up; pts, patients; NA, not applicable.

Supplementa	ry Table 5I	3. Pattern	of recurre	ence and nu	mber of re	current sit	te according	to ER/PR	status at i	nitial diagn	osis and re	currence	
	Pre-opE	R+ and ER-	at rec	Pre-opER+ and ER- at rec			Pre-opER- and ER- at rec			Pre-opER- and ER+ at rec			p value
	No. of Pts	5-y	p value	No. of Pt	5-y SAR	p value	No. of Pts	5-y SAR	p value	No. of Pts	5-y SAR	p value	
	(%)	SAR		(%)	(%)		(%)	(%)		(%)	(%)		
		(%)											
	17(100%)	88.2		12(100%)	41.7		17(100%)	21.2		5(100%)	75.0		0.011
Number of recurrent sites													
Solitary	12(71%)	91.7	0.315	7(58%)	42.9	0.752	6(35%)	41.7	0.014	2(40%)	100.0	0.564	0.274
Multiple	5(29%)	80.0		5(42%)	40.0		11(65%)	9.1		3(60%)	66.7		0.121
Pattern of recurrence													
Vagina	6(35%)	All alive	0.426	0	NA	0.133	0	NA	0.164	0	NA	0.317	NA
		(34.5-1											
		01.9M)											
Pelvis	2(12%)	All alive		0	NA		0	NA		0	NA		NA
		(20.2,											
		31.6 M)											
Distant	5(29%)	80.0		10(83%)	50.0		8(47%)	31.3		3(60%)	100.0		0.540
Pelvis-distant	4(24%)	75.0		2(17%)	0.0		9(53%)	11.1		2(40%)	50.0		0.018

Rec, recurrence; SAR, survival after recurrence; f/u, follow-up; pts, patients; NA, not applicable.

Supplementary Table 6A. Failure pattern and survival of patients with stage III/IV according to adjuvant therapy (n = 53).

	No. with isolated vaginal failure (5-y SAR)	No. with pelvic failure (5-y SAR)	No. with distant failure (5-y SAR)	No. with pelvic+distant failure (5-y SAR)
Surgery alone (n = 0)	0 (NA)	0 (NA)	0 (NA)	0 (NA)
Surgery + CT (n = 17)	0 (NA)	1 (DOD at 2 months)	7 (14.3%)	9 (33.3%)
Surgery $+RT (n = 8)$	0 (NA)	0 (NA)	7 (38.1%)	1 (DOD at 9.1 months)
Surgery +CT+RT $(n = 28)$	0 (NA)	1 (Alive at 31.6 months)	23 (16.4%)	4 (0.0%)

SAR, survival after recurrence; CT, chemotherapy; RT, radiotherapy; DOD, died of disease; NA, not applicable.

	No. with isolated vaginal	No. with pelvic failure	No. with distant failure	No. with pelvic+distant	
	failure (5-y SAR)	(5-y SAR)	(5-y SAR)	failure (5-y SAR)	
Surgery alone (n = 6)	1 (Alive at 42.6 months)	1 (Alive at 20.2 months)	3 (33.3%)	1 (DOD at 14.1 months)	
Surgery $+CT (n = 1)$	0 (NA)	0 (NA)	0 (NA)	1 (DOD at 3.2 months)	
Surgery $+RT (n = 3)$	0 (NA)	0 (NA)	3 (0.0%)	0 (NA)	
Surgery + $CT+RT$ (n = 1)	0 (NA)	0 (NA)	0 (NA)	1 (DOD at 32.9 months)	

Supplementary Table 7. Failure pattern and survival of patients with positive cytology according to adjuvant chemotherapy (n=18).

	No. with isolated vaginal failure (5-y SAR)	No. with pelvic failure (5-y SAR)	No. with distant failure (5-y SAR)	No. with pelvic+distant failure (5-y SAR)
Adjuvant chemother	apy			
Yes	0 (NA)	0 (NA)	11 (0.0%)	4 (0.0%)
No	0 (NA)	0 (NA)	2 (1 DOD at 12.6 months, 1 Alive at 52.9 months)	1 (Alive at 72.6 months)

SAR, survival after recurrence; DOD, died of disease; NA, not applicable.

	Simple hysto	erectomy $(n = 29)$)		RH-PLND (n =	= 13)		
	No. with isolated vaginal failure (5-y SAR)	No. with pelvic failure (5-y SAR)	No. with distant failure (5-y SAR)	No. with pelvic+distant failure (5-y SAR)	No. with isolated vaginal failure (5-y SAR)	No. with pelvic failure (5-y SAR)	No. with distant failure (5-y SAR)	No. with pelvic+distant failure (5-y SAR)
Adjuvant pelvic RT	<u> </u>							
Yes	0 (NA)	0 (NA)	14 (0.0%)	4 (0.0%)	0 (NA)	0 (NA)	5 (0.0%)	0 (NA)
No	0 (80.0%)	2 (1DOD at 2 months, 1 Alive at 20.2 months)	2 (0.0%)	7 (0.0%)	1 (Alive at 42.6 months)	0 (NA)	5 (40.0%)	2 (50.0%)
5-year SAR*	0%				38.5%			

SAR, survival after recurrence; RT, radiotherapy; RH-PLND, radical hysterectomy and pelvic lymph node dissection; DOD, died of disease; NA, not applicable.

p = 0.034