

# Risk Factors for Failing Cervical Cancer Screening in Incidental Cervical Carcinoma at Time of Simple Hysterectomy

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# Disclosures

Tara Castellano, MD

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Figure 3. Leading Sites of New Cancer Cases and Deaths – 2018 Estimates

Estimated New Cases	Male					Female		
	Prostate	164,690	19%			Breast	266,120	30%
	Lung & bronchus	121,680	14%			Lung & bronchus	112,350	13%
	Colon & rectum	75,610	9%			Colon & rectum	64,640	7%
	Urinary bladder	62,380	7%			Uterine corpus	63,230	7%
	Melanoma of the skin	55,150	6%			Thyroid	40,900	5%
	Kidney & renal pelvis	42,680	5%			Melanoma of the skin	36,120	4%
	Non-Hodgkin lymphoma	41,730	5%			Non-Hodgkin lymphoma	32,950	4%
	Oral cavity & pharynx	37,160	4%			Pancreas	26,240	3%
	Leukemia	35,030	4%			Leukemia	25,270	3%
Liver & intrahepatic bile duct	30,610	4%	Kidney & renal pelvis	22,660	3%			
<b>All sites</b>	<b>856,370</b>	<b>100%</b>	<b>All sites</b>	<b>878,980</b>	<b>100%</b>			

Estimated Deaths	Male					Female		
	Lung & bronchus	83,550	26%			Lung & bronchus	70,500	25%
	Prostate	29,430	9%			Breast	40,920	14%
	Colon & rectum	27,390	8%			Colon & rectum	23,240	8%
	Pancreas	23,020	7%			Pancreas	21,310	7%
	Liver & intrahepatic bile duct	20,540	6%			Ovary	14,070	5%
	Leukemia	14,270	4%			Uterine corpus	11,350	4%
	Esophagus	12,850	4%			Leukemia	10,100	4%
	Urinary bladder	12,520	4%			Liver & intrahepatic bile duct	9,660	3%
	Non-Hodgkin lymphoma	11,510	4%			Non-Hodgkin lymphoma	8,400	3%
Kidney & renal pelvis	10,010	3%	Brain & other nervous system	7,340	3%			
<b>All sites</b>	<b>323,630</b>	<b>100%</b>	<b>All sites</b>	<b>286,010</b>	<b>100%</b>			

Estimates are rounded to the nearest 10, and cases exclude basal cell and squamous cell skin cancers and in situ carcinoma except urinary bladder. Ranking is based on modeled projections and may differ from the most recent observed data.

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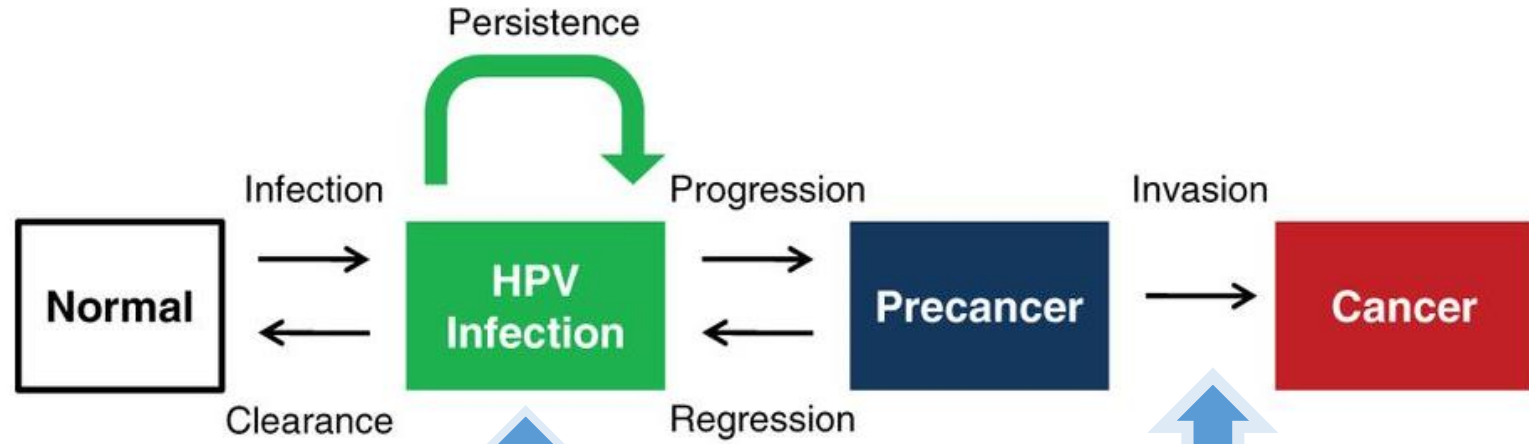
Third most  
common  
gynecologic cancer

13,240 new cases  
4,170 deaths

Incidence

US: 7.5 per 100,00  
Oklahoma: 9.2 per 100,00

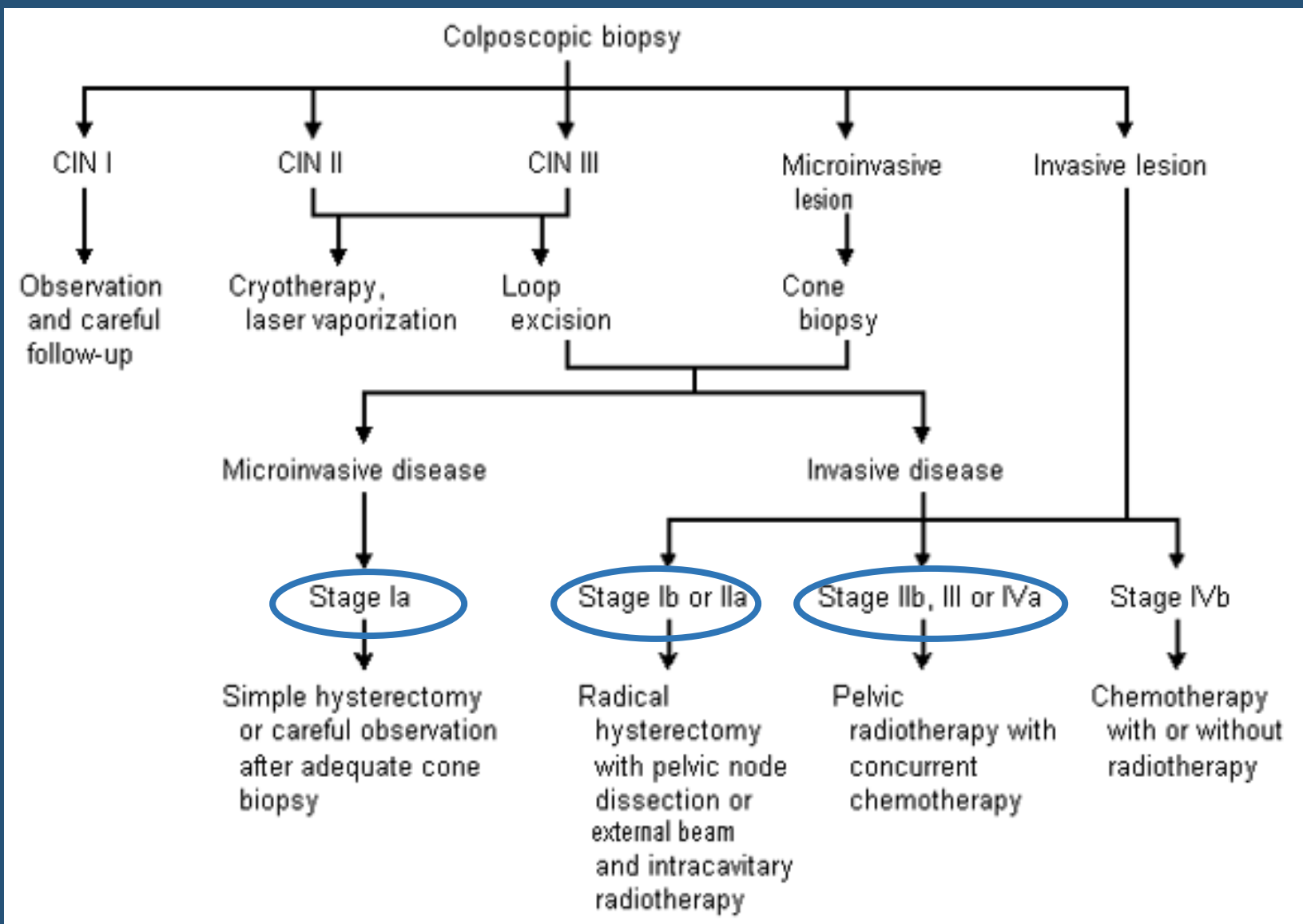
Natural history  
model



HPV implicated  
99% of all  
cervical  
cancers

HPV vaccine  
primary  
prevention

Screening  
secondary  
prevention



# Indications for Simple Hysterectomy

- Benign indications<sup>[4]</sup>:
  - Uterine fibroids (51%)
  - AUB (42%)
  - Pain/Endometriosis (30%)
  - Pelvic Prolapse (18%)
- Premalignant indications:
  - Persistent cervical dysplasia
  - Endometrial hyperplasia



# ASCCP Guidelines<sup>[6,7]</sup>

Table 1. Screening Guidelines for Cervical Cancer

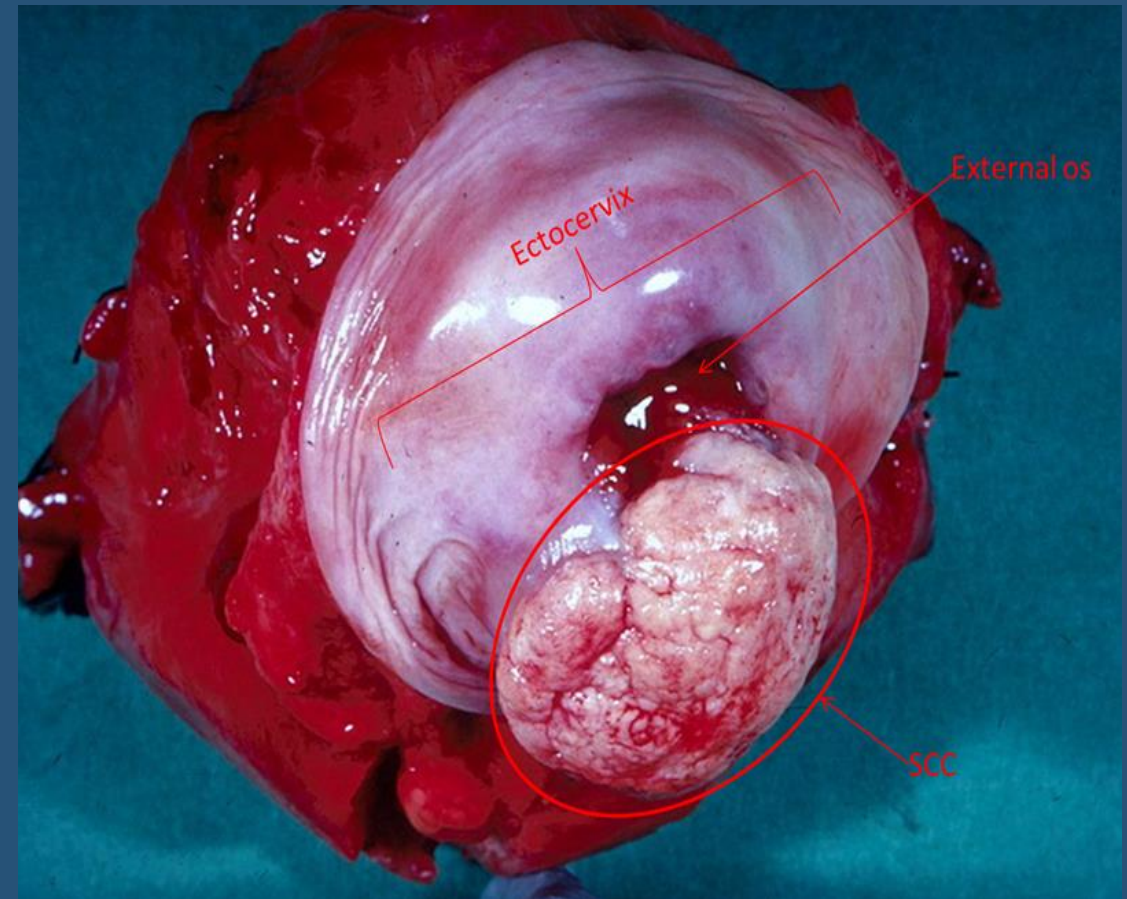
Age Group	Recommendation
<21 y	No screening recommended
21-29 y	Cytology (Pap smear) every 3 years Abnormal Pap tests may be followed up with HPV testing
30-65 y	Cytology + HPV testing every 5 years (the preferred recommendation) or Cytology every 3 years
>65 y	No cervical cancer screening required if the woman has had regular screening in the previous 10 years and no serious precancers in the last 20 years or Continue testing if an abnormality was found in the last 20 years

HPV: human papillomavirus. Source: References 9-13.



# Incidental Invasive Cervical Cancer

- Incidence difficult to define. Most studies are majority microscopic disease and non-US populations<sup>[8-12]</sup>
- Terminology not uniform:
  - “Occult” ~ microscopic disease
  - “Cut through” ~ macroscopic disease
  - “Incidental” ~ any



# Why it matters

- Adjuvant radiotherapy or radical reoperation is mandatory after inadequate operation.
- Those with residual disease after simple hysterectomy (gross tumors or microscopic positive surgical margins) or IIb disease are at higher risk for locoregional recurrence following treatment and have a worse 5 year OS
  - 38-50% vs 60-70% in those treated appropriately <sup>[14]</sup>
- Evidence for higher toxicity associated with post hysterectomy radiation therapy <sup>[15]</sup>

# Objectives

- Review patient cases with incidental cervical carcinoma at time of simple hysterectomy.
- Determine risk factors associated with inappropriate execution of cervical cancer screening guidelines and/or risks factors for screening failures.

# Demographics

- N= 59 subjects
  - Median age 44 years (range 30-82)
  - Caucasian (93%, n=55)
- Indication:
  - AUB (57%, n=34)
  - Dysplasia (41%, n=24)
  - Pain (24%, n=14)
  - Fibroids (10%, n=6)
- Insurance status:
  - Under- or uninsured: 23.7%, n=14
- Histology:
  - SCC (63%, n=37)
  - Adenocarcinoma (30%, n=18)
  - other (7%, n=4)
- Stage:
  - Ia1, Ia2, Ib1 (65%, n=38)
  - Ib2, IIa (7%, n=4)
  - IIb, III, IV (15%, n=9)
- Referring provider
  - ObGyn (71%, n=42)
  - Unknown (22%, n=13)
  - GenSurg/Other (7%, n=4)

# Results

- Cervical dysplasia was a listed indication for surgery in 41% (n=24) of subjects.
- Inappropriate screening was found in 71% (n=42) of the referrals.
- False negative screening occurred in 22% (n=13) subjects.
- Incomplete data or documentation in 6% (n=4)

# Results: Mistakes in Screening

- Inappropriate screening was found in 71%, n=42 of the referrals.
  - 38% (n=16) of the there was no documented pap smear
  - 21% (n=9) of the time the pap wasn't triaged appropriately
  - 17% (n=7) received pap and colposcopy but not conization
  - 23% (n=10) had a LEEP that wasn't managed appropriately.

# Results: Mistakes in Screening

N=43	No pap (n=17)		Pap but failed colposcopy (n=9)		Colpo but failed treatment (n=7)		Excised but failed referral (n=10)	
<b>Age:</b> n=40	</=45 (n=10) >45 (n=7)	p= 0.50	</=45 (n=6) >45 (n=3)	p=0.91	</=45 (n=3) >45 (n=2)	p=0.80	</=45 (n=7) >45 (n=2)	p=0.70
<b>Histology:</b> <b>SCC vs AC</b> n=43	SCC (n=10) Other (n=7)	p= 0.48	SCC (n=7) Other (n=2)	p= 0.37	SCC (n=4) Other (n=3)	p=0.75	SCC (n=7) Other (n=2)	p= 0.37
<b>Insurance</b> <b>Yes vs No</b> N=39	Yes (n=13) No (n = 4)	p=0.95	Yes (n=7) No (n=2)	p= 0.94	Yes (n=7) No (n=2)	p=0.34	Yes (n=7) No (n=1)	p=0.43

# Results: Screening Failures

- False negative screening occurred in 24% (n=14) of 59 subjects.
  - There was no significant association with age, insurance status, or histology to risk of false negative screening.

N=59	Age: n=54 </= 45 (n=34) > 45 (n=20)		Histology: n=57 SCC (n=37) Other (n=20)		Insurance: n=53 Yes (n=41) No (n=12)	
False negative screen (n=14)	</=45 (n=8)	p=0.60	SCC (n=7)	p=0.31	Yes (n=11)	p=0.90
Mistake in screening (n=45)	45 yo (n=6)		Other (n=7)		No (n=3)	

# Conclusions

- Failure to adhere to screening guidelines was most common reason for ICC at the time of simple hysterectomy
- Less frequently, false negative screening was the cause.
- No independent risk factor in either group was able to be identified in this small single institutional study.

# Future Directions

- Continue patient and provider education regarding adherence to pre-operative cervical cancer screening
- Continue to collect data on the best management of incidentally found cervical cancer, especially in bulkier disease and presence of positive margins.

# References:

- [1] <https://seer.cancer.gov/statfacts/html/cervix.html> Accessed 1/2018
- [2] Walboomers JM, Jacobs MV, Manos MM, et al. Human papillomavirus is a necessary cause of invasive cervical cancer worldwide. J Pathol 1999; 189:12.
- [3] [https://www.nccn.org/professionals/physician\\_gls/pdf/cervical.pdf](https://www.nccn.org/professionals/physician_gls/pdf/cervical.pdf). Accessed 1/2018.
- [4] ACOG practice Bulletin “Choosing the Route of Hysterectomy for Benign Disease” July 2017
- [5] Daley E, Perrin K, Vamos C, Hernandez N, Anstey E, Baker E, Kolar S, Ebbert J. Confusion about Pap smears: lack of knowledge among high-risk women. J Women's Health (Larchmt). 2013;22:67–74.
- [6] Massad LS, Einstein MH, Huh WK, et al. 2012 updated consensus guidelines for the management of abnormal cervical cancer screening tests and cancer precursors. J Low Genit Tract Dis 2013; 17:S1.
- [7] <http://www.asccp.org/asccp-guidelines>. Accessed 1/2018
- [8] Srisomboon J, Pantusart A, Phongnarisorn C, Suprasert P. Reasons for improper simple hysterectomy in patients with invasive cervical cancer in the northern region of Thailand, J Obstet Gynaecol Res, 2000, vol. 26 (pg. 175-80)Google
- [9] Behtash N, Mousavi A, Mohit M, Modares M, Khanafshar N, Hanjani P. Simple hysterectomy in the presence of invasive cervical cancer in Iran, Int J Gynecol Cancer , 2003, vol. 13 (pg. 177-81)
- [10] Ayhan A, Otegen U, Guven S, Kucukali T. Radical reoperation for invasive cervical cancer found in simple hysterectomy, J Surg Oncol, 2006, vol. 94 (pg. 28-34)
- [11] Koh HK, Jeon W, Kim HJ, Wu HG, Kim K, Chie EK, et al. Outcome analysis of salvage radiotherapy for occult cervical cancer found after simple hysterectomy. Jpn J Clin Oncol 2013;43:1226–1232
- [12] Chapman JA, Mannel RS, DiSaia PJ, Walker JL, Berman ML. Surgical treatment of unexpected invasive cervical cancer found at total hysterectomy, Obstet Gynecol, 1992, vol. 80 (pg. 931-4)
- [13] Kinney WK, Egorshin EV, Ballard DJ, Podratz KC. Long-term survival and sequelae after surgical management of invasive cervical carcinoma diagnosed at the time of simple hysterectomy, Gynecol Oncol, 1992, vol. 44 (pg. 24-7)
- [14] Hsu W, Sheung PW, Jen YM, et al. Long-term treatment results of invasive cervical cancer patients undergoing inadvertent hysterectomy followed by salvage radiotherapy. Int. J. Radiation Oncology Biol. Phys., Vol. 59, No. 2, pp. 521–527, 2004.
- [15] Smith KB, Amdur RJ, Yeung AR, Morris CG, et al. Postoperative Radiotherapy for Cervix Cancer Incidentally Discovered After a Simple Hysterectomy for Either Benign Conditions or Noninvasive Pathology. *American Journal of Clinical Oncology* . June 2010, Vol 33.3 (p229-232).