



# CANAL ANAL ANATOMY AND ITS DIFFERENT PATHOLOGY

## Intraanal lesions

-Lesion of anal canal from rectal-anal transition area

-**Poorly Differentiated**

-More common in women

## Perianal lesions

- Completely visible

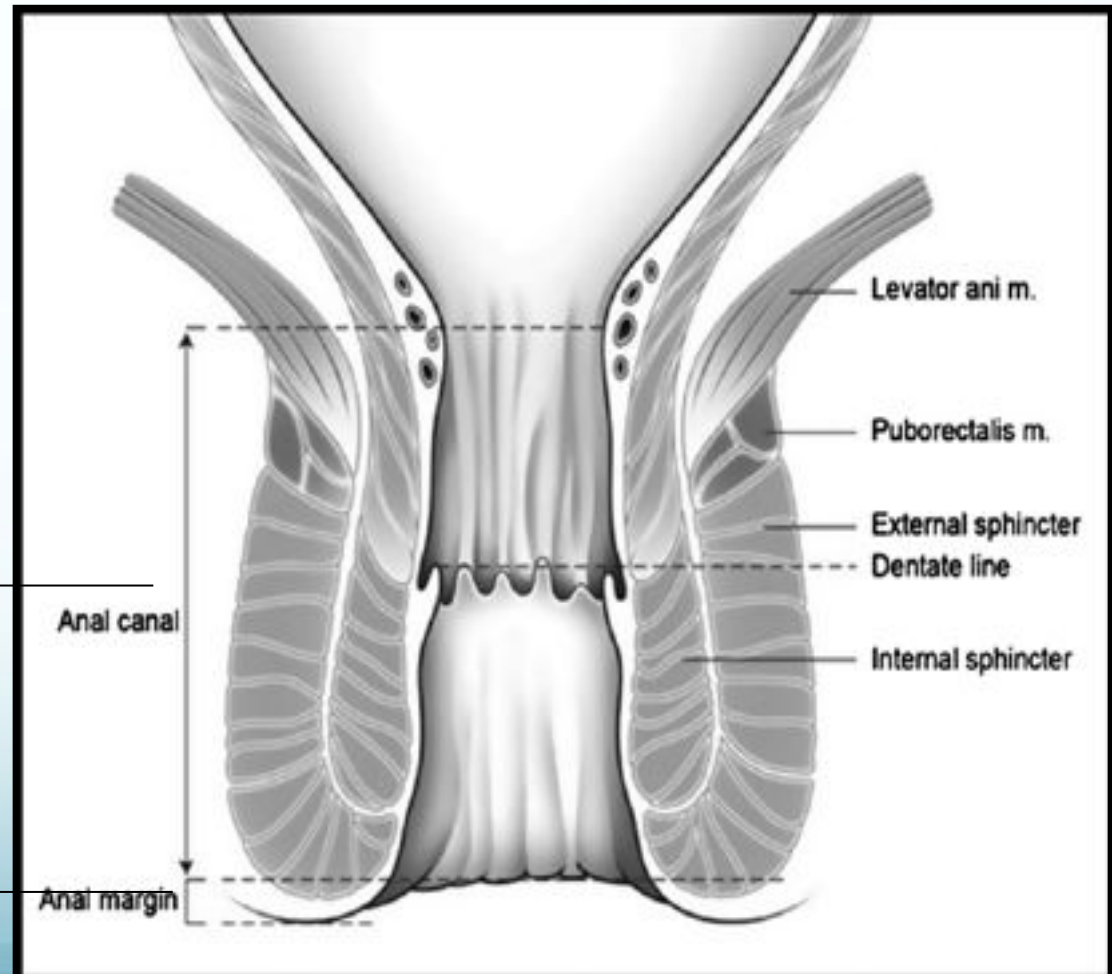
-Within a 5-cm radius of anal orifice

-**Well Differentiated**

-More common in men

## Skin lesions

- Outside of the 5-cm radius



# Anal cancer: ESMO-ESSO-ESTRO clinical practice guidelines for diagnosis, treatment and follow-up <sup>☆</sup>

Robert Glynne-Jones <sup>a</sup>, Per J. Nilsson <sup>b</sup>, Carlo Aschele <sup>c</sup>, Vicky Goh <sup>d</sup>, Didier Peiffert <sup>e</sup>, Andrés Cervantes <sup>f</sup>, Dirk Arnold <sup>g,\*</sup>

Radiotherapy and Oncology 111 (2014) 330–339



TNM staging. American Joint Committee on Cancer/Union for International Cancer Control (AJCC/UICC) seventh edition TNM clinical and pathological classification of anal cancer

Primary tumour (T)			
TX	Primary tumour cannot be assessed		
T0	No evidence of primary tumour		
Tis	Carcinoma in situ (i.e., Bowen disease, high-grade squamous intraepithelial lesion, and anal intraepithelial neoplasia II–III)		
T1	Tumour ≤2 cm in greatest dimension		
T2	Tumour >2 cm but ≤5 cm in greatest dimension		
T3	Tumour >5 cm in greatest dimension		
T4	Tumour of any size invades adjacent organ(s), e.g., vagina, urethra, and bladder.		
Regional lymph nodes (N)			
NX	Regional lymph nodes cannot be assessed		
N0	No regional lymph node metastasis		
N1	Metastases in perirectal lymph node(s)		
N2	Metastases in unilateral internal iliac and/or inguinal lymph node(s)		
N3	Metastases in perirectal and inguinal lymph nodes and/or bilateral internal iliac and/or inguinal lymph nodes		
Distant metastasis (M)			
M0	No distant metastasis		
M1	Distant metastasis		
Anatomic stage/prognostic groups			
Stage	T	N	M
0	Tis	N0	M0
I	T1	N0	M0
II	T2	N0	M0
	T3	N0	M0
IIIA	T1	N1	M0
	T2	N1	M0
	T3	N1	M0
	T4	N0	M0
IIIB	T4	N1	M0
	Any T	N2	M0
	Any T	N3	M0
IV	Any T	Any N	M1

## FACTORS TO CONSIDER IN TREATMENT DECISION-MAKING FOR AC

Disease-related factors	Patient-related factors	Other
Clinical and radiological TNM stage Site of tumour (margin, canal, rectal) Extent of tumour i.e. involvement of vagina (risk of fistulation) in addition to size Response to treatment (early and at 26 weeks) Need for symptom control	Patient preferences Biological age/renal function/Charlson geriatric assessment Co-morbidities/current medications and performance status  Socio-economic and psychological factors/social support Severity of initial symptoms	Local expertise (brachytherapy etc.) Geriatricians with interest in oncology   Specialist palliative care



## STAGE AND SITE-BASED TREATMENT

<i>Anal canal</i>	
Surgery (radical or local excision) generally contraindicated as primary treatment option	
STAGE I	- Standard dose radiotherapy (RT), infused 5FU and mitomycin (stage group under-represented in randomised studies) - low dose RT, infused FU and mitomycin (no data from randomised studies)
STAGE II-III	- Standard dose RT, infused FU and mitomycin (evidence from multiple randomised studies)
STAGE IV	- 5-FU and cisplatin, carboplatin/taxol, or possibly irinotecan/cetuximab
<i>Anal margin</i>	
STAGE I, well differentiated	- Local excision (re-excision or chemoradiation if involved/close margins)
STAGE II-III	- standard dose RT, infused 5FU and mitomycin C
STAGE IV	- 5-FU and cisplatin, or carboplatin/taxol

# WHEN..... SURGERY?



Impossibile visualizzare l'immagine collegata. È possibile che il file sia stato spostato, rinominato o eliminato. Verificare che il collegamento rimandi al file e al percorso corretti.

## SURGERY AS PRIMARY TREATMENT

Until the mid-1980s, radical surgery was the cornerstone of treatment. However, following publications from the 1970s on combined modality therapy, surgery as the primary therapeutic option has generally been abandoned.

### STILL TODAY:

**LOCAL EXCISION** is recommended in:

- Small lesion < 2cm
- Involving anal margin

**BUT** contraindicated:

- In poorly differentiated
- In lymphnodal involvement

**AND** discussed in canal anal small lesions

If inadequate surgical margins (<5mm), it needs a new local excision

## Chemoradiation schedule and assessment used in the ACT II trial

### Chemoradiation

- **5FU** 1000 mg/m<sup>2</sup> days 1–4 (week 1) and 29–32 (week 5) by continuous 24 h IV infusion.
- **MITOMYCIN** 12 mg/m<sup>2</sup> IV bolus on day 1 (maximum single dose 20 mg)
- **RADIOTHERAPY\***: Total dose 50.4 Gy delivered in 28 daily fractions starting on Day 1.

### Assessment of tumour response

- **Digital examination** at 11, 18 and 26 weeks from the start of the treatment.
- **Abdominopelvic CT** at week 26.
- Confirm residual or recurrent disease by **biopsy** (routine biopsies not recommended).

Complete response

Follow-up

Persistent / recurrent disease

Surgery



### CLINICAL PRESENTATION

Anal canal cancer<sup>a</sup> → Biopsy: squamous cell carcinoma<sup>b</sup>

### WORKUP

- Digital rectal examination (DRE)
- Inguinal lymph node evaluation
  - Biopsy or FNA if suspicious nodes
- Chest x-ray or Chest CT
- Anoscopy
- Abdominal/pelvic CT or MRI
- Consider HIV testing + CD4 level if indicated
- Gynecological exam for women, including screening for cervical cancer
- Consider PET-CT scan<sup>c</sup>

### CLINICAL STAGE

T1-2, N0

T3-T4, N0  
or  
Any T, N+

Metastatic disease

### PRIMARY TREATMENT<sup>g</sup>

Mitomycin/5-FU<sup>d</sup> + RT<sup>e</sup> (45<sup>e</sup>-59 Gy)

Mitomycin/5-FU<sup>d</sup> + RT<sup>e,f,g</sup> (55-59 Gy)

Cisplatin-based chemotherapy<sup>h</sup> ± RT<sup>g</sup>

See Follow-Up Therapy and Surveillance (ANAL-3)

See Follow-Up Therapy and Surveillance (ANAL-3)







### CLINICAL PRESENTATION

Anal margin lesion<sup>1</sup>

Biopsy: squamous cell carcinoma<sup>b</sup>

### WORKUP

- Digital rectal examination (DRE)
- Inguinal lymph node evaluation
  - Biopsy or FNA if suspicious nodes
- Chest x-ray or Chest CT
- Anoscopy
- Abdominal/pelvic CT or MRI
- Consider HIV testing + CD4 level if indicated
- Gynecological exam for women, including screening for cervical cancer

### CLINICAL STAGE

T1, N0 Well differentiated

T2-T4, N0 or Any T, N+

Metastatic disease

### PRIMARY TREATMENT<sup>9</sup>

Local excision

Adequate margins

Inadequate margins

Observe

Re-excision (preferred) or Consider local RT<sup>e</sup> ± 5-FU-based chemotherapy<sup>d</sup>

Mitomycin/5-FU<sup>c</sup> + RT<sup>e,f,g</sup> (55-59 Gy)

See Follow-up Therapy and Surveillance (ANAL-3)

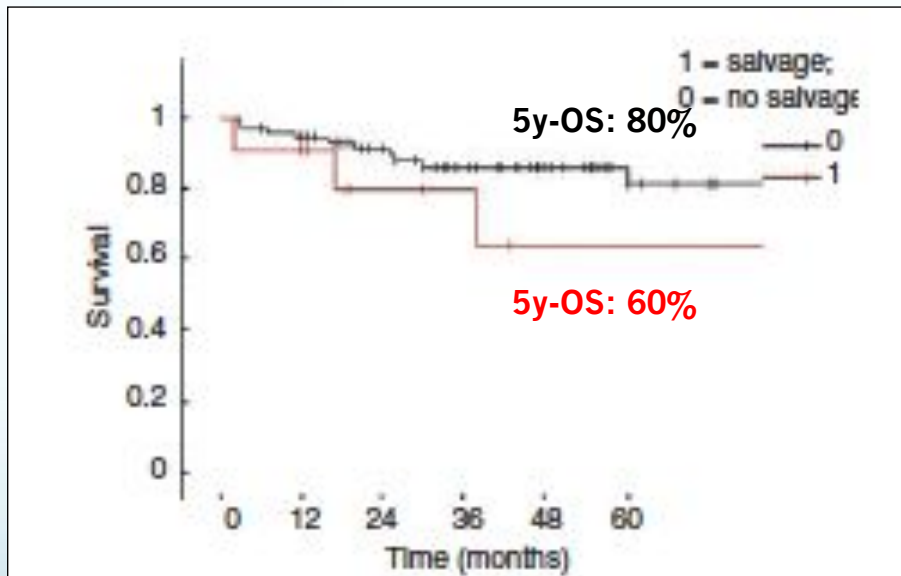
Cisplatin-based chemotherapy<sup>h</sup> ± RT<sup>e</sup>



# Outcome of salvage surgery for anal squamous cell carcinoma

D. A. Harris\*, J. Williamson\*, M. Davies\*, M. D. Evans\*, P. Drew† and J. Beynon\* on behalf of the Swansea Pelvic Oncology Group

2013, Association of colonproctology of Great Britain



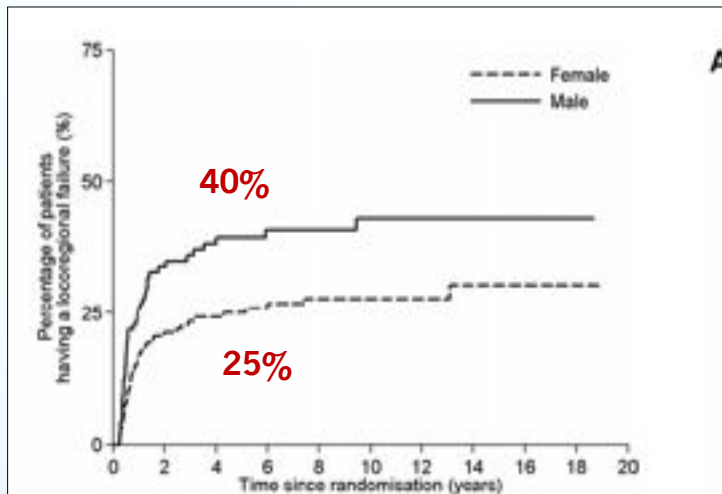
OS of persistent/recurrent disease treated with only ChRt or plus Salvage Surgery

# Prognostic Factors for Recurrence and Survival in Anal Cancer

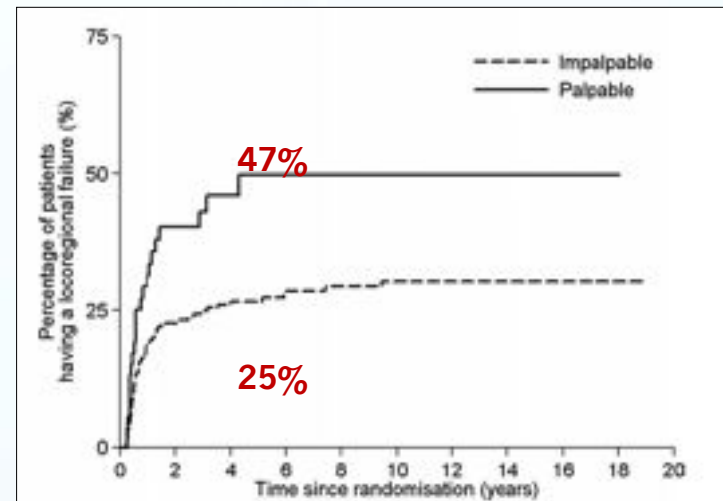
Generating Hypotheses From the Mature Outcomes of the First United Kingdom Coordinating Committee on Cancer Research Anal Cancer Trial (ACT I)

Robert Glynne-Jones, MD, FRCR<sup>1</sup>; David Sebag-Montefiore, MD, FRCR<sup>2</sup>; Richard Adams, MD, FRCR<sup>3</sup>; Simon Gollins, MD, FRCR<sup>4</sup>; Mark Harrison, MD, FRCR<sup>1</sup>; Helen M. Meadows, MSc<sup>5</sup>; Mark Jitlal, MSc<sup>5</sup>; for the United Kingdom Coordinating Committee on Cancer Research Anal Cancer Trial Working Party

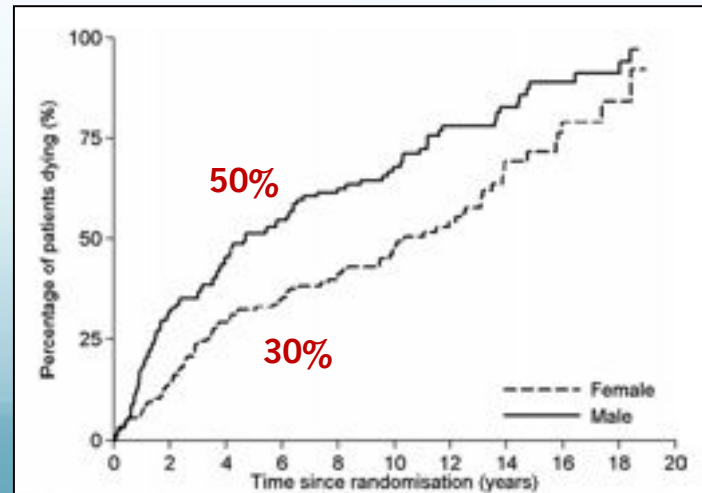
## 5y-LR-RECURRENCE



## 5y-LYMPHNODAL-RECURRENCE

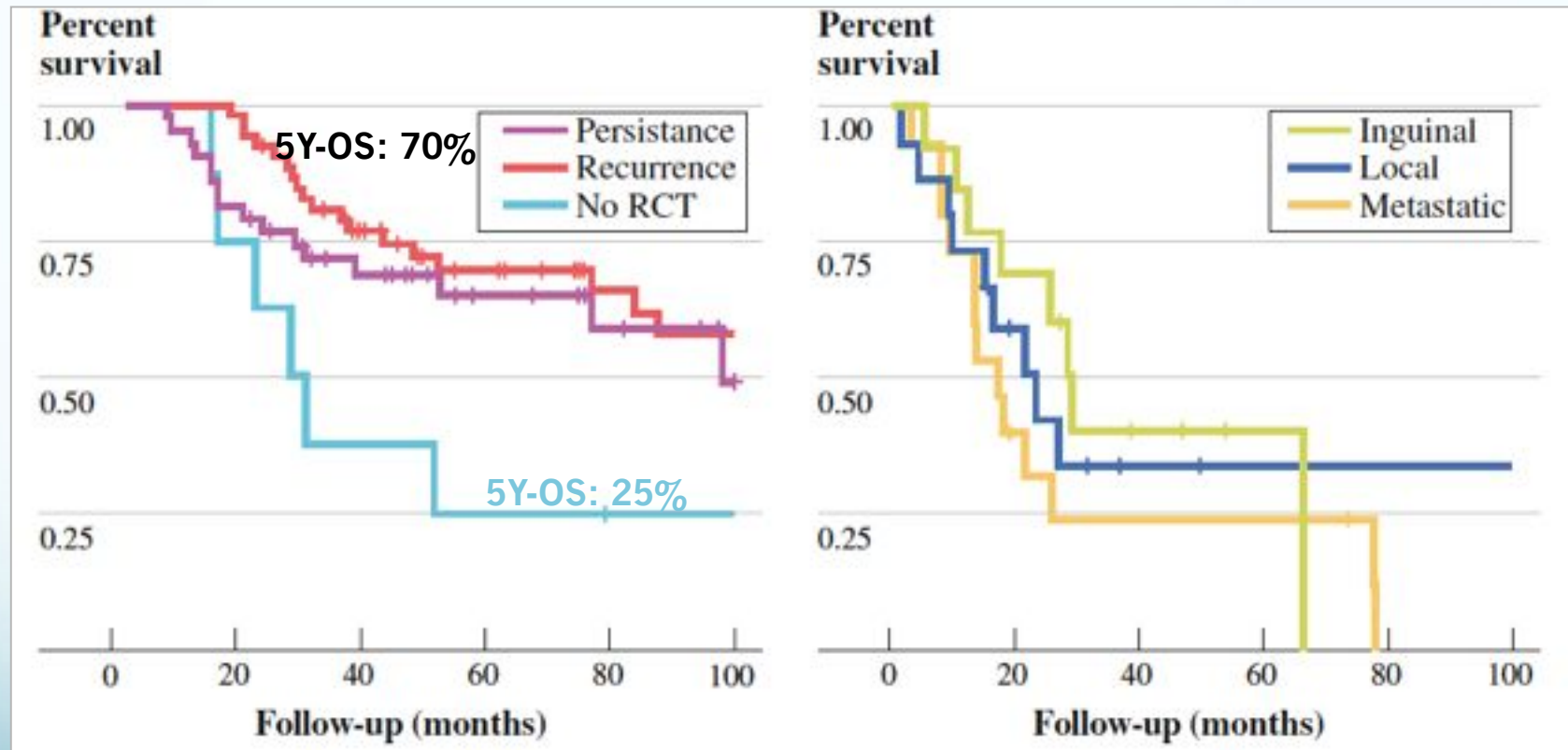


## 5y- DEATH PERCENTAGE



## Abdominoperineal Resection for Squamous Cell Anal Carcinoma: Survival and Risk Factors for Recurrence

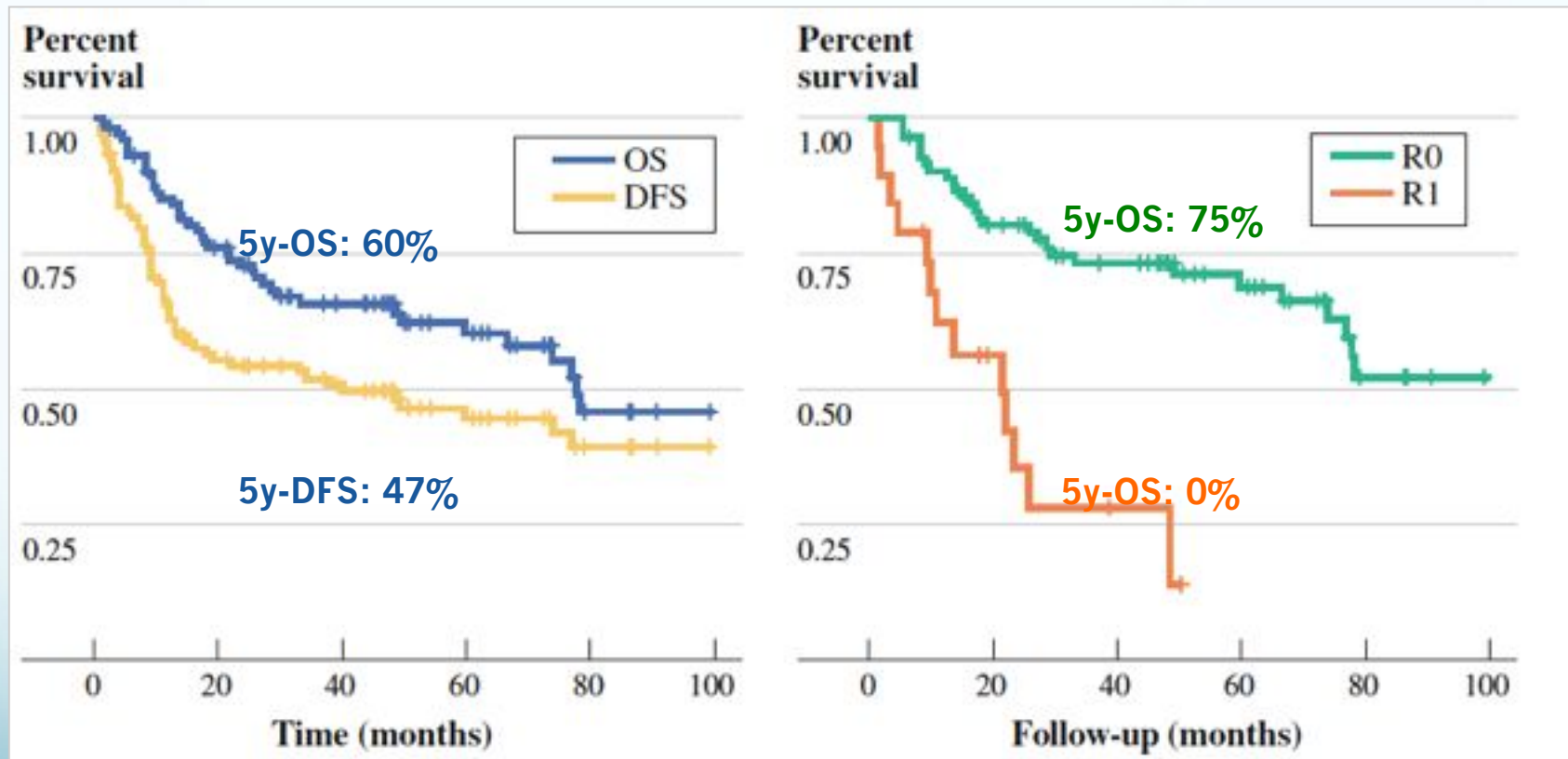
Jérémie H. Lefèvre, MD<sup>1</sup>, Hélène Corte, MD<sup>1</sup>, Emmanuel Tiret, MD<sup>1</sup>, David Boccara, MD<sup>2</sup>, Marc Chaouat, MD<sup>2</sup>, Emmanuel Touboul, MD<sup>3</sup>, Magali Svrcek, MD, PhD<sup>4</sup>, Magalie Lefrancois, MD<sup>1</sup>, Conor Shields, MD<sup>1</sup>, and Yann Parc, MD, PhD<sup>1</sup>



- **RECURRENCES: 40%**
- **SIDE OF RECURRENCES DIDN' T INFLUENCE OS**
- **SURVIVAL BENEFIT FROM CHRT**

## Abdominoperineal Resection for Squamous Cell Anal Carcinoma: Survival and Risk Factors for Recurrence

Jérémie H. Lefèvre, MD<sup>1</sup>, Hélène Corte, MD<sup>1</sup>, Emmanuel Tiret, MD<sup>1</sup>, David Boccara, MD<sup>2</sup>, Marc Chaouat, MD<sup>2</sup>, Emmanuel Touboul, MD<sup>3</sup>, Magali Svrcek, MD, PhD<sup>4</sup>, Magalie Lefrancois, MD<sup>1</sup>, Conor Shields, MD<sup>1</sup>, and Yann Parc, MD, PhD<sup>1</sup>

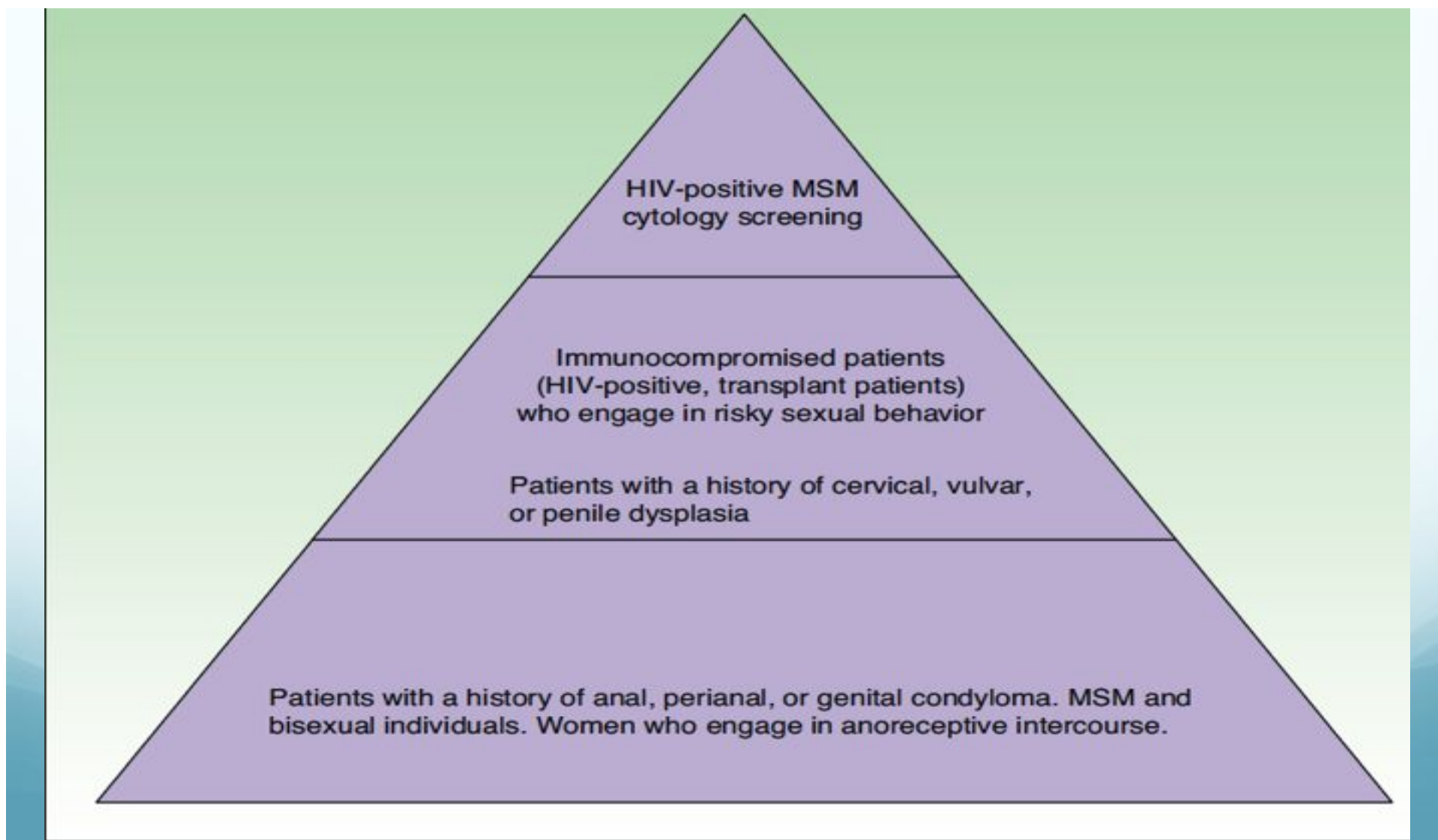


THE MAJOR PROGNOSTIC FACTOR OF OS, AFTER SALVAGE SURGERY, IS  
**THE POSITIVITY OF SURGICAL MARGINS**  
THE NEED OF AN **EXTENSIVE SURGERY**

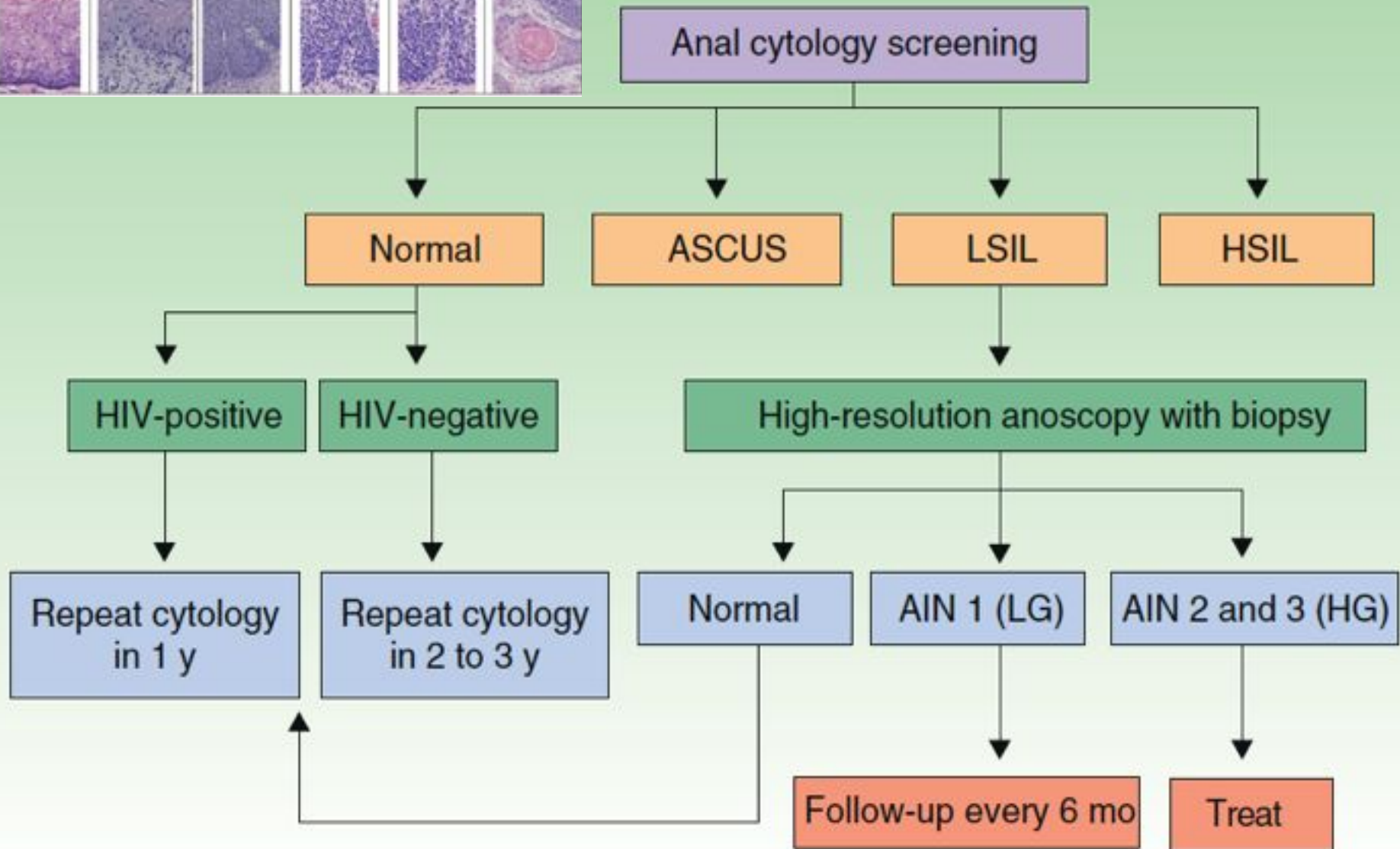
## REVIEW

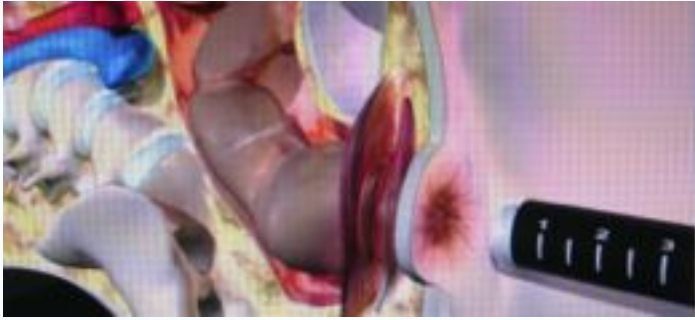
# Early Detection of Anal Intraepithelial Neoplasia in High-Risk Patients ☆

E. Sendagorta,<sup>a,\*</sup> P. Herranz,<sup>a</sup> H. Guadalajara,<sup>c</sup> F.X. Zamora<sup>b</sup>

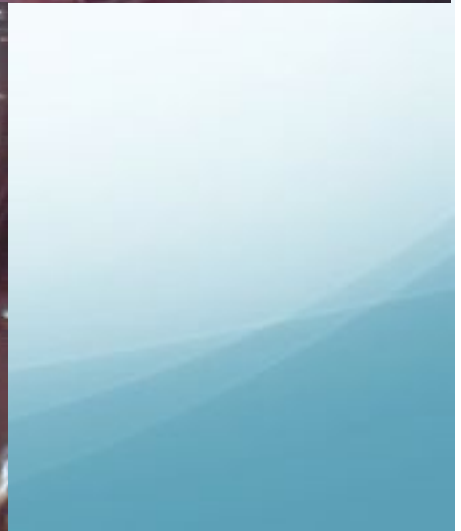
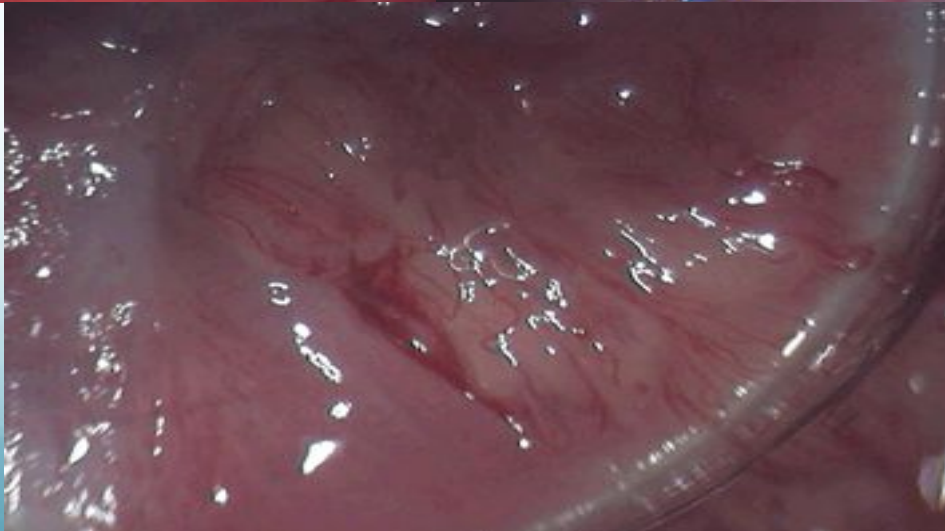
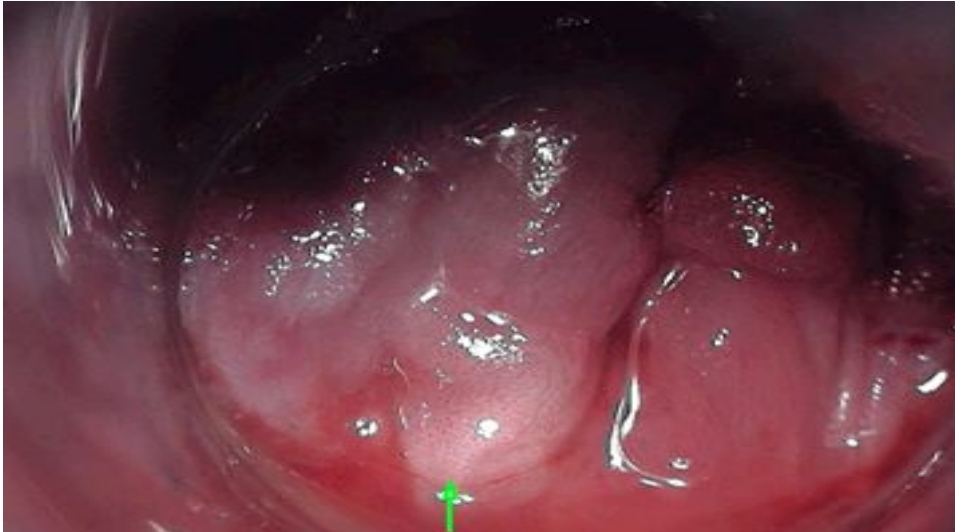


Non-Dysplastic Epithelium	LSIL		HSIL		Micro-Invasion
	AIN1		AIN3		
	Mild Dysplasia	Moderate Dysplasia	Severe Dysplasia	Carcinoma In Situ	





## DIGITAL VIDEOPROCTOSCOPY





# Will local ablation of high grade anal intraepithelial neoplasia prevent invasive anal cancer?



Alessia Dalla Pria and Mark Bower

*AIDS* 2013, 27:1185–1186

Yes, treatment causes regression of AIN. However whether this prevents the evolution of invasive cancer remains uncertain. In one study of anal cancer in people living with HIV, seven of 74 patients had been enrolled on

29% [11]. It is also important to note that the rates of AIN2/3 were higher in women (7%) than in MSM (5%) and heterosexual men (1%). Both these findings have important relevance to screening programs and health economic studies of these projects.

- A GOOD PROGRAM OF SCREENING IS NECESSARY TO PREVENT PRE-CANCEROUS LESIONS
- BUT THE TREATMENT OF THESE LESIONS REMAINS DEBATED FOR THE PREVENTION OF ANAL CANCER

# CONCLUSIONS

- AC NEEDS A MULTIDISCIPLINARY TEAM AND TREATMENT
- THE SURGEON, IN LOCALLY ADVANCED AC , FOLLOWS THE ONCOLOGIST/RADIOTHERAPIST IN PERSISTENT/RECURRENT DESEASE
- IN EARLY STAGES, ESPECIALLY IN ANAL MARGIN CANCER, SURGERY IS ESSENTIAL
- A PATHOLOGICAL SCORE COULD BE HELPFUL TO DISCRIMINATE A SUBGROUP OF PATIENTS AT HIGH RISK OF RECURRENCE OR PERSISTENT DESEASE
- HYMMUNODEFICIENT PATIENTS REQUIRE A STRONG SCREENING PROGRAM
- DIGITAL PROCTOSCOPY COULD BE PROMISING FOR AN EARLY DIAGNOSIS OF AC IN HIGH RISK PATIENTS