Cancer of the Anal Canal A Multidisciplinary Perspective

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# All Talks need clear Objectives



#### **Overview and Objectives**

- Verarching: Try to focus on (a) what is relevant for surgeons and (b) what is new
- Underlying anatomy/histology (not new)
- 2. Staging Work-up (a little new)
- 3. Treatment
  - Early: T1-T2 : A little bit of everything?
  - Advanced: T4 or N2/3 : Radiation evolving
  - Follow-up guidelines

Background Anatomy/Histology



# Histology

Mucosa lined Anal Canal

- Begins at junction of puborectalis portion of levator ani and the external anal sphincter
- Ends at Anal verge
- Divided by the dentate line (transition from glandular or columnar, to squamous mucosa)
- Epidermis lined anal margin
- \* This begins at introitus of the anal orifice
- Transition from squamous mucosa to epidermis lined peri-anal skin

# Terminology

 <u>Adenocarcinomas</u> – glandular elements in the anal tract/low rectum. Treated like rectal cancer (although LN drainage may be different)

 <u>Anal cancers</u> – tumors that develop from the mucosa (SCC, Basaloid, Nonkeratinized above dentate, keratinizing below dentate)
<u>Perianal (SKIN) cancers</u> – tumors that are distal to squamous mucocutaneous junction (e.g. hair) Staging Where are the risks?

# Anal Cancer Staging: T stage

- T1 <2 cm
- T2 2-5 cm
- T3 >5 cm
- T4 Invades adjacent organ

# Anal Cancer Nodal staging

- N0 No nodal mets
- N1 Perirectal Lymphnodes
- N2 Unilateral internal iliac or inguinal lymphnodes
- N3 Perirectal and iliac/inguinal OR Bilateral iliac/inguinal



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# Patterns of Recurrence



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# Prognostic groups - Staging

- I T1N0
- II T2-T3N0
- IIIA T1-3N1 or T4N0
- IIIB T4N1 or N2 or N3 disease
- IV M1
- 5 yr overall survival going from:
- I 85+% II 70-85%
- IIIA 50-60+% IIIB 40-50%

#### Work Up

1 Hx - Px2. CT A/P (+/- MRI) 3 Chest imaging PET scan (changes RT 13% of time\*) Other: Nodal biopsy Trans-rectal U/S (N1 staging)

\*Int J Radiat Oncol Biol Phys. 2012 Sep;84(1):66-72. Epub 2012 May 15.

## Anal Cancer Treatment



"The doctor's on strike, so I had his stitches done by the vet"

#### One slide about an Anal Cancer Trial

> UK consortium (Lancet 1996; 348: 1049–54) Eligibility T1-T4 (any N) – no mets Randomly assigned 585 patients RT alone 45 Gy + 15 Gy boost Same RT with infusional 5FU and Mitomycin C Chemo reduced LOCAL FAILURE and Cause specific mortality

# Trials are good for those in the middle



### Middle of the road tumors: T3 or N1 tumors

- These guys are the bulk of patients we see
  - They were the bulk of the trial
- They get treated based on that (and other) Randomized trial evidence
  - Nothing new here
  - Chemotherapy is 5FU and Mitomycin C
  - Radiation is anywhere from 50-60 Gy

# Early Anal Cancers Treatment in Evolution



#### Early Anal Cancers

These patients seem to present in a variety of ways

- "Regular" Bleeding anal lesions
- Hemorrhoids
- Skin tags
- Warts
- SCC of Anal canal may or may not have been the top of the differential

#### T1-T2 after local resection

Bulk of tumor should be removed (R1 or at most 1-2 cm of tumor remaining) Low radiation dose (30 Gy) and only 1 cycle of chemo \* (Half the radiation and half the chemo) Institutions have been giving varying volumes of radiation I would radiate the inguinal and low rectal nodes as well as the primary (little sideeffects vs. risk of recurrence)

### Radiation Volumes – Low dose



#### What to do with T4/N2-3 Tumors?

\* Various places do various things

- Induction chemo has NOT worked (RCT)
- More radiation?
  - Higher doses
  - Boost the primary site
  - Brachytherapy

No good (randomized) evidence to do anything different than standard But, only at most 50% with local recurrence are salvagable

# Patterns of Recurrence



Posterior-anterior field

Lateral field

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# This volume is not tolerated to a high dose



# IMRT – Intensity Modulated Radiation Therapy



# Local Control is better with more conformal radiation



**r.** 2011 Aug 1;117(15):3342-51.

#### Survival may also be better



Cancer. 2011 Aug 1;117(15):3342-51.

#### BC Data: "Freedom from any Recurrence"



5 yr Outcomes worse with treatment breaks:

Continuous course 69.5%

Split course 56.1%

P=0.001

Data from Dr. John Hay

# Post Treatment Recommendations (NCCN)

- Hx and Px
- > DRE
- Inguinal exam
- > Anoscopy
- Yearly CT chest/abdo/pelvis for those with locally advanced disease (look for mets)

We can salvage local recurrence with APR? Benefit of finding a distal recurrence?

#### Summary

- Work Up: PET scanning should be added to the work-up algorithm
- T1-T2 Tumors: Evolving role for local resection and low dose chemo and XRT. Especially important for those that may not tolerate higher doses
- T3 and N+: Radiation treatments are evolving aiming to spare normal structures and increase dose to tumors
- Need to follow to detect early local recurrence.
  - \* ?Utility of detection of early metastatic disease?

# The End

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"I only got up for a drink of water, and a queue's formed next to my bed."