# Pelvic Irradiation Side effects, Prevention and Treatment

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# \* Pelvic Irradiation Side Effects

### **RT** is indicated in :

60%: cervical cancer patients,45 % of endometrial cancer patients,35 % of vulvar cancer patients,

100 % of vaginal cancer patients,5 % of patients with ovarian cancer

\* The incidence and severity of RT side effects depend :
 \* Site,

- \* Volume of tissue exposed
- \* Treatment schedule: total dose, dose per fraction, and type of radiation.
- \* Previous surgery,
- \* Concomitant chemotherapy
- \* Comorbid illness :
- \* Smoking history (strong predictor for both bowel and bladder complications)
- \*•Active collagen vascular disease,
- \* Inflammatory bowel disease (IBD)
- \* Vascular disorders diabetes and hypertension
- \* Microvascular disease.

# \*Side effect timing

- \*Acute toxicities :during or shortly after the course of treatment.
- \*Subacute toxicities :4 to 12 weeks after RT pneumonitis or myositis.
- \*Late toxicities :after three months.



### **Acute radiation cystitis:**

- \* Dysuria, frequency, urgency, nocturia, and bladder spasms.
- \* RT-induced bladder inflammation and edema, which can compromise urothelial integrity.

#### IMRT :

\* reduce GU toxicity with adjuvant treatment after <u>hysterectomy</u> for cervical and endometrial cancer

\*

### \*<u>NSAIDS</u> :irritative voiding symptoms

- Anticholinergics and/or antispasmodics
   (<u>oxybutynin</u> or <u>hyoscyamine</u>) for cystitis or bladder spasm
- \*•Cranberry juice or <u>phenazopyridine</u> for dysuria

# \* Late signs and symptoms

\*EBRT Dose <50 Gy : Late GU symptoms only grade 1,2

- \* Urinary urgency, incontinence, and limitations in daily activities due to bladder symptoms (5%)
- \* EBRT + Brachytherapy : D2cc > 101 Gy more late complications ,
- \* Contracture =>> pain syndromes.
- \* Ureteral stricture
- \* Urethrovaginal and vesicovaginal Fistula
- \* Radioation hematuria :Sodium pentosan polysulfate (Cap 100mg), <u>Hyperbaric Oxigenation</u>

# \* GI Toxicity

\*GI toxicity is <u>the most common</u> of both acute / late side effects

### \*Acute radiation injury

Cramping, diarrhea, anorexia, malaise, rectal discomfort, and tenesmus

### Total dose and volume irradiated

the degree of acute toxicity does predict for the development of late effects

loose stool and cramping.

**Small Bowel** :high-volume watery diarrhea.

Large bowel fecal urgency, clustered bowel movements, and tenesmus

<u>Cisplatin</u> with pelvic radiation  $\rightarrow$  >2 fold the risk for grade >3 acute GI side effects .

## \* Early GI Toxicity \* Proctitis and rectal discomfort

- \* Enema with <u>hydrocortisone</u> or cod liver oil, anti-inflammatory suppositories, and a lowresidue diet : no grease, spices, or insoluble fiber.
- \* Two small prospective studies :topical <u>sodium butyrate</u> enemas in the treatment of acute radiation proctitis .Butyrate is a short chain fatty acid.
- \* Ondansetron has been shown to reduce radiotherapy-induced nausea and vomiting.
- \* <u>chlorpromazine</u> and <u>dexamethasone</u>The addition of dexamethasone to ondansetron appears to improve nausea control
- Antidiarrheal medications.
- \* <u>loperamide</u>. When symptoms are refractory, <u>diphenoxylate-atropine</u> and <u>tincture of opium</u> can be effective.
- \* <u>Octreotide</u> is more effective than diphenoxylate-atropine With loperamide-refractory diarrhea, subcutaneous octreotide led to complete resolution for 80 % of patients,
- \* Endometrial cancer undergoing RT +selenium-deficient :<u>Selenium sulfate</u>:reduced rates of grade 2 or higher diarrhea.
- • Probiotics : a preventive strategy for radiation enteritis.

# \*Late GI Toxicity

#### \* Mucosal atrophy and loss of mucin-producing goblet cells

- \* The symptoms of late GI toxicity :
- \* •Chronic diarrhea -. Ongoing antidiarrheal medications are often necessary.
- \* •Malabsorption This can be related to RT involving the distal ileum. vitamin B12 deficiency occurs in 12 to 20 percent. Cholestyramine when bile salt malabsorption

#### \* Fibrosis

- \* Recurrent ileus or obstruction
- \* Prolonged chronic radiation enteritis -> Malnutrition: perioperative nutritional therapy
- \* Mucosal telangiectasias or ulcerations
- \* The signs :painless hematochezia, tenesmus, or pain.
- \* Median time to onset is 14 months and symptoms typically appear within three years
- \* Image-guided brachytherapy has the potential to further reduce its incidence].
   \* IMRT
- \* Rectal proctopathy:
- \* Avoiding constipation may limit episodes of bleeding.
- \* <u>Sucralfate</u> enemas, aluminum hydroxide
- \* In refractory cases: endoscopic intervention
- \* Topical formalin , argon plasma coagulation at controlling bleeding
- \* topical butyrate does not reduce symptoms of chronic proctitis
- \* Surgical management for :transfusion-dependent bleeding, refractory pain, and fistula,

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	ANTIDIARRHEAL INTERVENTIONS	
SCREENING	INTERVENTION <sup>j</sup>	
<ul> <li>Provide immediate antidiarri therapy indicated by grade</li> </ul>	heal	
GRADE 1	<ul> <li>If chemotherapy induced, decrease or delay the next dose of ch</li> <li>Provide oral hydration and electrolyte replacement</li> <li>Initiate antidiarrheal (eg, diphenoxylate/atropine) if patient not a</li> </ul>	nemotherapy already on opioids
GRADE 2	<ul> <li>Provide IV fluids if patient is unable to tolerate oral fluids</li> <li>Initiate/continue antidiarrheal—as above</li> <li>Consider anticholinergic agents</li> <li>If non-C.diff infection-related: Treat with appropriate antibiotics</li> <li>If C. diff infection-related: Administer antibiotics and probiotics</li> <li>If chemotherapy-induced, decrease or delay the next dose of cf</li> <li>If immunotherapy-mediated diarrhea, consider</li> <li>Corticosteroids</li> <li>Infliximab</li> <li>Probiotics</li> <li>See Management of Immunotherapy-Related Toxicity Guidelin immunotherapy-related diarrhea</li> </ul>	as appropriate nemotherapy n <u>es</u> for
GRADES 3/4	<ul> <li>Inpatient hospitalization (intensive care for Grade 4 if consisten</li> <li>For GVHD diarrhea, consider limiting diet, steroids, and IV nutri</li> <li>Provide IV fluids and use antidiarrheal agents and anticholinerg</li> <li>Consider somatostatin analog</li> <li>Consider parenteral hydration in home setting</li> </ul>	nt with goals) ition Activate W gics as mentioned above Go to Settings



## \*Acute vaginal mucositis

\* <u>Vaginal douches</u>:local anesthetic+anti-inflammatory agent, <u>benzydamine</u>

\*vaginal estrogen

- \*Vaginal stenosis : <u>the most common</u> late vaginal side effect in EBRT and/or BT
- \*Vaginismus
- \*sexual dysfunction (dysparunia)
- \* <u>Topical estrogen</u> (one to three times weekly) applied in the first six months following radiation reduces dyspareunia and improves vaginal caliber
- \*Fistulas Rectovaginal and vesicovaginal fistulas are a rare but serious. The need for <u>interstitial brachytherapy</u> may increase this risk compared with intracavitary brachytherapy
- \*As with vaginal necrosis: <u>hyperbaric oxygen</u> and <u>pentoxifylline</u>

## \*Ovaries Skin And Bone Marrow \*Infertility or premature menopause

- \*premature ovarian failure (POM) The dose is 16.5 Gy at 20 Y/O and 14.3 Gy at 30 Y/O
- \*laparoscopic ovarian transposition
- \*at least 3 cm from the radiation field border+clips.
- \*High rates of preservation (80 to 88 %)
- \*BONE AND BONE MARROW
- \*SKIN : erythema and soreness to moist desquamation, AND ulceration
- \*skin hygiene, water-based creams, or ointments such as <u>lanolin</u>
- \*Moist desquamation :<u>Silvadene cream</u>
- Radiation-induced fibrosis: oral pentoxifylline + vitamin E



