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Radiation Proctitis- Experience at a Tertiary Care Centre of North India



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Abstract

Introduction: Proctitis is a troublesome complication in patients receiving radiotherapy in pelvic malignancies. This is a prospective study done to evaluate the efficacy of 4% formalin in treatment of radiation proctitis and complications associated with it.

Method: Patients with rectal bleeding post radiotherapy for gynaecological malignancy were analysed in our institution from June 2010-May 2011. 50ml of 4% formalin was sprayed through colonoscopy with mucosal contact time of 10 min and observed for 4hrs for any complication.

Results: A total of 22 patients with mean age 57 (range 40-65) years, moderately built with radiation proctitis were subjected to 4% formalin and followed up for 12(range 1-36) months. Mean interval between radiotherapy and presenting symptoms was 15(range 6-24) months. Cessation of bleeding occurred in 77.27% cases after mean of 2(range 1-4) cycles of spray with hemoglobin rise of 2gm% (range 1.1-2.9). 5 patients complained of intense pain relieved with analgesics and one had seizure. There was only one mortality in a patient who got operated for recto-vaginal fistula and expired on 5th post-operative day due to septicemia.

Conclusion: 4% formalin application is a simple, cost-effective, and well-tolerated method of controlling hemorrhagic radiation proctitis with less complications. It can be effectively used in settings with limited resources.

Keywords: Troublesome; Patients; Photon generator; Gamma rays; European organization; Cancer; Oncology; Symptoms; Colonoscopy; Septicemia; Coagulation; Surgery; Bleeding

Introduction

Today, radiation to the pelvis is an important constituent in the treatment of pelvic malignancies and is administered either as neoadjuvant or adjuvant therapy [1]. Radiation therapy can be administered via external beam radiation or radioactive implants termed brachytherapy. External beam radiation is typically administered by an external photon generator with a variety of sources including gamma rays, electron beams, and X-rays via a four-beam approach which results in significant exposure to surrounding organs [2]. Brachytherapy involves the implantation of radioactive pellets (typically iodine 125) into the target tissue with a gradual release over time. This reduces exposure damage to surrounding tissue based on careful positioning of the implanted pellets and consequently lesser colorectal complications [3].

Table 1: Individually attempted to classify the degree of radiation proctitis into a graded system of symptoms progressing from no symptoms to fatal complications.

Grade	Symptoms / Signs	Overall Management
0	No Symptoms	None
1	Occasional urgency and occasional pain; superficial ulceration <1 cm2, occult bleeding, and mild stricture.	Outpatient management; no lifestyle adjustments
2	Intermittent urgency and intermittent pain; superficial ulceration >1 cm2, occasional bleeding, and moderate stricture	Outpatient management; some lifestyle adjustments
3	Persistent urgency and persistent pain; deep ulceration, persistent bleeding, severe stricture	Possible short hospital admission or minor surgical interven- tion; major lifestyle adjustments

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4	Refractory urgency and uncontrollable pain; gross haemorrhage, perforation, fistula, complete obstruction	Long-term hospital admission or major surgical intervention
5	Sepsis, multiorgan failure, and death	Fatal complications

Incidence of radiation proctitis is likely related to the dose of radiation, area of exposure, method of delivery, and the use of cytoprotective agents. The doses generally delivered to the pelvis vary from 45 to 50Gy for adjuvant or neoadjuvant treatment for prostate or anorectal malignancies; up to 90Gy is considered the definite therapy for gynaecological malignancies. Doses above 70Gy cause significant and long-standing injury to the surrounding area [4,5] The Radiation Therapy Oncology Group (RTOG) and the European Organization for Research and Treatment of Cancer (EORTC) individually attempted to classify the degree of radiation proctitis into a graded system of symptoms progressing from no symptoms to fatal complications [6,7] (Table 1).

Radiation proctitis can be managed medically or surgically depending on the severity. Low grade can be managed with the use of nonsteroidal anti-inflammatory drugs, sucralfate, short chain fatty acids, hyperbaric oxygen and anti-oxidants. More severe variety of proctitis may preclude the use of ablative procedures or surgery. Local ablation with formalin is considered to be a safe and effective way to treat radiation proctitis causing significant bleeding. Formalin scleroses and seals fragile neovasculature in radiation damaged tissues to prevent further bleeding through chemical cauterization [8,9]. We have conducted a prospective study to evaluate the efficacy of formalin in managing patients of rectal bleeding with radiation proctitis.

Material and Methods

The present study was conducted at our prime institute, PGIMS, Rohtak from June 2010 to May 2011. Thirty patients of age ranging from 40-65 years with mean age 57 years with symptoms of rectal bleeding, post-radiotherapy for gynecological malignancies were included in this study. Complete Haemogram was done in every patient. Mean hemoglobin of 8.7% (range -3.5-11gm%) was noted with microcytic hypo chromic anemia. Platelets and total leukocyte count was normal. Mean interval between radiotherapy and presenting symptoms- 15 months with range of 6 months-2 years. 8 patients were excluded and 22 patients out of which, 16 patients had carcinoma cervix and only 6 patients had carcinoma endometrium, underwent colonoscopy. 50 ml of 4% formalin was sprayed through colonoscopy with a contact period of 10 minutes. Patients were observed for 4 hours for any complications and then followed up weekly. If there is no relief after 1 week, repeat session was done.

Observation

After formalin spray, out of 22 patients, rectal bleeding stopped in 17 patients (77.2%) with mean hemoglobin rise of 2gm%. Patients required 1-4 cycles of spray with mean of 2 cycles. 5 patients remained refractory even after 4 cycles and were treated with argon plasma coagulation. 5 patients required blood transfusion with mean of 3 units (range 1-5 units). 5 patients had intense pain relieved after injectable pain killers. Only 1 patient

had seizure. There was only 1 death due to septicemia in a patient operated for recto vaginal fistula.

Discussion

In radiation proctitis, vascular telangiectasia and nonhealing mucosal ulceration, perhaps due to an underlying obliterative arteritis, may lead to severe recurrent haemorrhage. These patients can be managed conservatively with anti-inflammatory agents, sucralfate, short chain fatty acids, antioxidants, ablation procedures with formalin or argon plasma coagulation and in severe cases may require surgery. Formalin application has been an important method for controlling haemorrhage in patients with proctitis. Formalin scleroses and seals fragile neovasculature in radiation damaged tissues to prevent further bleeding through chemical cauterization [10]. Formalin has been historically used as Intravesical instillation in treatment of radiation cystitis [11,12]. This concept was first used by Robinson et al in 1986 as a method of treating radiation proctitis in a 71-year-old male with severe bleeding [13].

Two methods of formalin application have been described so far, which includes rectal instillation of 4% formalin solution or direct topical application of a 10% formalin solution [14]. Topical application is generally applied through rigid proctoscope or directly applying with formalin-soaked gauge with contact period of about 2-3 minutes. The perianal skin needs to be protected during the procedure to prevent stricture and skin damage. Rectal instillation of 4% formalin has been successfully employed for treatment of radiation proctitis.

In 1993, Seow-Choen et al. published the first case series involving eight patients treated for intractable rectal bleeding from radiation proctitis with instillation of 4 percent formalin [15]. The bleeding ceased in all patients; however, one developed a worsening stricture requiring Hegar's dilation. Use of formalin in refractory cases of radiation proctitis has been studied by Scalarides et al., [16] in 16 patients. Bleeding stopped in 12 patients after single application and occurred sporadically in 3 patients. Only 1 patient required 3 applications before bleeding stopped and 4 patients had developed significant anal pain and tenesmus after treatment. In 2003, Parikh et al. [17] described the application of cotton pledget soaked in 4 percent formalin applied to the proctitis through a rigid sigmoid scope or operating anoscope. Of 33 patients, 29 (88 percent) had improved or complete resolution of bleeding after a mean of 3.4 treatments and at a follow-up of 18 months.

We have reported a case study of 22 patients of gynaecological malignancy who presented with rectal bleeding post radiation proctitis. 4% formalin was instilled with a mean contact time for 10 minutes. Out of 22 patients, bleeding stopped in 17 patients (77.7%) after a mean of 2 sessions. On follow up, these patients

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had improved haemoglobin with an average rise of 2mg%. 5 patients were refractory to the treatment and were treated with argon plasma coagulation. 5 patients had rectal pain which was managed with pain killers and no major complication was reported in our study.

Conclusion

Radiation proctitis can be a therapeutic challenge, even in the most experienced hands. Majority of patients can be treated with topical modalities, while surgery may offer the only chance of relief from life-threatening symptoms. Treatment with local formalin instillation is simple, safe, inexpensive and well tolerated modality and may be used as first line treatment in a poor resource country.

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