

Retrospective review of adverse events after treatment of non-melanoma skin cancer with image-guided superficial radiation therapy

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Background

- Basal cell carcinoma (BCC) and squamous cell carcinoma (SCC) are the most common types of non-melanoma skin cancer.¹⁻³
- Treatment options include wide local excision, electrodesiccation and curettage, topical therapies, oral anti-neoplastic medications, and Mohs.^{1,3,4}
- Superficial radiation therapy (SRT) is another treatment option as both a definitive and adjunct therapy.^{1,3-6}
- Although superficial radiation therapy has shown favorable outcomes with a low risk of complications, it still poses a risk for adverse events.
- The frequency and severity of these complications are not well defined with newer SRT devices that have image-guided high-frequency ultrasound.

Objective

This study aimed to evaluate the complications of treating non-melanoma skin cancer with Image-guided superficial radiation therapy at an academic outpatient dermatology clinic.

Methods

Retrospective chart review on patients treated with image-guided superficial radiation therapy (IG-SRT) at the Health Science Center Dermatology Clinic

Retrospective chart review

Study period: December 12/2018 to 12/2022

Follow up through: 12/31/2023

Adverse events collected during the treatment and follow-up

Primary endpoint

Any cutaneous adverse event that required treatment or diagnostic studies beyond the 2-week follow-up visit

Results

Table 1. Adverse events

Characteristic s	Patient 1	Patient 2	Patient 3	Patient 4	Patient 5	Patient 6
Age (years)	82	84	64	64	59	83
Gender	Female	Female	Male	Male	Female	Male
Location	Right lower leg	Left lower leg	Vertex scalp	Center forehead	Right naris	Right lower leg
Adverse event	Chronic non-healing wound	Chronic non-healing wound	Discoid lupus	Geometric Favre-Racouchot syndrome	Geometric Favre-Racouchot syndrome	Radiation recall
Histological subtype	SCC, Invasive	SCC in-situ	SCC in-situ	BCC, nodular	BCC, nodular	SCC, invasive
Follow-up	Dermatology & wound care ~ 2 years	Dermatology & wound care ~ 4 years	Topical Tacrolimus 0.1% ointment & photo-protection	Topical adapalene 0.1% gel → lost to follow-up	Topical adapalene 0.1% gel → shave removal & curettage	Topical triamcinolone 0.1% ointment

Table 2. Rate of Adverse Events with different treatment modalities

Treatment modality	Adverse event rate	Adverse events
IG-SRT	1.1% (6/566)	Chronic non-healing wound, discoid lupus, geometric Favre-Racouchot syndrome, radiation recall
Mohs surgery ⁷	0.5% - 2%	Infection, hematoma, dehiscence, necrosis
Vismodegib ⁸	12% - 74%	Muscle spasm, dysgeusia, alopecia, weight loss, fatigue, diarrhea, nausea, headache
Wide local excision ^{9,10}	3.8% - 20%	Bleeding, infection, flap necrosis
Electrodesiccation & Curettage ¹¹	-----	Keloid, hypertrophic scar
PD-1 Inhibitors ¹²	Up to 100%	Diarrhea, fatigue, nausea, constipation, rash, anemia, hypothyroidism, pneumonitis, death

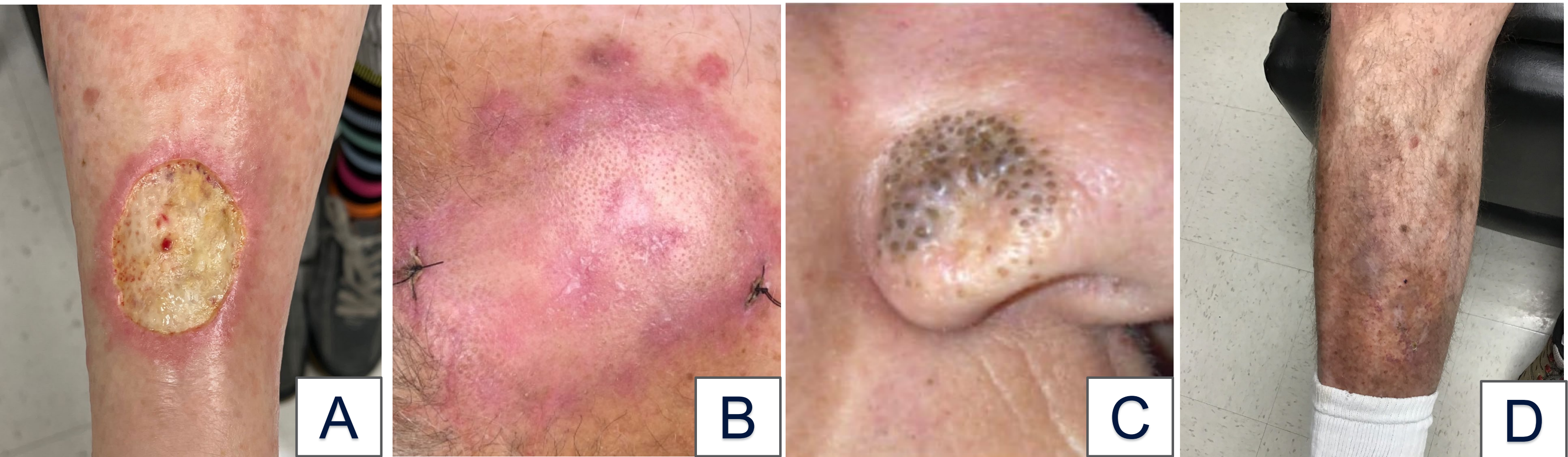


Figure 1. A) Well-demarcated ulcerated plaque with central granulation tissue and an erythematous border in the field of prior radiation therapy- chronic non-healing wound B) Pruritic, annular, scaly plaque with accentuated borders in area of prior radiation therapy- discoid lupus C) Well-demarcated plaque at site of prior radiation treatment with comedones around a central scar at site of malignancy- geometric Favre-Racouchot syndrome D) Edematous hyperpigmented plaque with focal linear hemorrhagic crust – radiation recall

Discussion

- Both patients with chronic non-healing wounds had venous stasis with chronic edema. A chronic inflammatory state with pre-existing cutaneous changes in addition to the radiation may have an increased risk of non-healing wounds¹³
- Patients are questioned about risk factors for radiation complications including a history of cutaneous lupus prior to superficial radiation treatment
- Both patients with Favre-Racouchot Syndrome were active tobacco users throughout treatment and both occurred on the face
- Other treatment sites of patient 4 that were not on the face did not cause Favre-Racouchot Syndrome
- The radiation recall occurred at a site of a previous superficial radiation therapy treatment field. He had approximately 10 prior radiation sites, but only one developed radiation recall
- The multiple courses of radiation treatments were likely a risk factor for having an incident of radiation recall
- IG-SRT shows favorable treatment outcomes with low adverse event rates compared to other common treatment modalities (Table 2)
- Limitations of the study include a single clinic population, a shorter follow-up period of 1-5 years, and a small patient sample size

Conclusion

- IG-SRT is a safe and effective treatment modality for the treatment of basal cell carcinoma and squamous cell carcinoma
- Patients with pre-existing co-morbid conditions may have an increased risk of developing certain adverse events
- Prospective studies investigating the frequency, risk factors for adverse events, and prevention strategies would enhance patients' ability to choose between the different treatments for non-melanoma skin cancer.

References

- McGregor S, Minni J, Herold D. Superficial Radiation Therapy for the Treatment of Nonmelanoma Skin Cancers. J Clin Aesthet Dermatol. Dec 2015;8(12):12-4.
- Dundar Y, Cannon RB, Hunt JP, Monroe M, Suneja G, Hitchcock YJ. Radiotherapy regimens in patients with nonmelanoma head and neck skin cancers. Int J Dermatol. Apr 2018;57(4):441-448. doi:10.1111/ijd.13879
- Han H, Gade A, Cecil FM, Lawson A, Auerbach S, Nestor MS. Superficial radiation therapy for nonmelanoma skin cancer: A review. Dermatological Reviews. 2023;3(6):409-417. doi:10.1002/der2.174
- Nestor MS, Berman B, Goldberg D, et al. Consensus Guidelines on the Use of Superficial Radiation Therapy for Treating Nonmelanoma Skin Cancers and Keloids. Journal of Clinical and Aesthetic Dermatology. 2019;12(2):12-18.
- Pashazadeh A, Boese A, Friebe M. Radiation therapy techniques in the treatment of skin cancer: an overview of the current status and outlook. J Dermatolog Treat. Dec 2019;30(8):831-839. doi:10.1080/09546634.2019.1573310
- Waldman A, Schmults C. Cutaneous Squamous Cell Carcinoma. Hematol Oncol Clin North Am. Feb 2019;33(1):1-12. doi:10.1016/j.hoc.2018.08.001
- Alam M, Ibrahim O, Nodzenski M, et al. Adverse events associated with mohs micrographic surgery: multicenter prospective cohort study of 20,821 cases at 23 centers. JAMA Dermatol. Dec 2013;149(12):1378-85. doi:10.1001/jamadermatol.2013.6255
- Verkouteren BJA, Wakkee M, Reyniers AKL, et al. Eight years of experience with vismodegib for advanced and multiple basal cell carcinoma patients in the Netherlands: a retrospective cohort study. Br J Cancer. Mar 2021;124(7):1199-1206. doi:10.1038/s41416-020-01220-w
- Bordeaux JS, Martires KJ, Goldberg D, Patten SF, Fu P, Maloney ME. Prospective evaluation of dermatologic surgery complications including patients on multiple antiplatelet and anticoagulant medications. J Am Acad Dermatol. Sep 2011;65(3):576-583. doi:10.1016/j.jaad.2011.02.012
- Bouhassira J, Bosc R, Grela L, et al. Factors associated with postoperative complications in elderly patients with skin cancer: A retrospective study of 241 patients. J Geriatr Oncol. Jan 2016;7(1):10-4. doi:10.1016/j.jgo.2015.11.004
- Knox JM, Lyles TW, Shapiro EM, Martin RD. Curettage and electrodesiccation in the treatment of skin cancer. Arch Dermatol. Aug 1960;82:197-204. doi:10.1001/archderm.1960.01580020039006
- Verkouteren BJA, Wakkee M, Reyniers AKL, et al. Eight years of experience with vismodegib for advanced and multiple basal cell carcinoma patients in the Netherlands: a retrospective cohort study. Br J Cancer. Mar 2021;124(7):1199-1206. doi:10.1038/s41416-020-01220-w
- Jacobson LK, Johnson MB, Dedhia RD, Niknam-Bienia S, Wong AK. Impaired wound healing after radiation therapy: A systematic review of pathogenesis and treatment. JPRAS Open. 2017;13:92-105. doi:10.1016/j.jpra.2017.04.001



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