Low-Dose Volumetric Modulated Arc Therapy for a Patient With Head and Neck Involvement of Mycosis Fungoides: A Case Report With a Review of Literature

Naoki Mukumoto 1, Haruo Inokuchi 1, Nobunari Hamaura 1, Mutsumi Yamagishi 1, Mai Sakagami 1, Shogo Matsuda 1, Daisuke Hayashi 2, Daisuke Tsuruta 2, Reiko Shibuya 1

1. Department of Radiation Oncology, Graduate School of Medicine, Osaka Metropolitan University, Osaka, JPN
2. Department of Dermatology, Graduate School of Medicine, Osaka Metropolitan University, Osaka, JPN

Corresponding author: Naoki Mukumoto, nao.muc.1229@gmail.com

Abstract

Mycosis fungoides (MF) is the most common type of cutaneous T-cell lymphoma that slowly progresses over a period of years to decades. In some cases, lesions that spread to the scalp, neck, or facial skin can have a significant impact on cosmetic appearance and a patient’s quality of life. Among the various treatments, radiation therapy is one of the most effective treatment modalities for patients with symptomatic cutaneous lesions. We report on an MF patient who had gradually increasing patches and plaques on the scalp, face, and neck who underwent irradiation with 20 Gy administered in 10 fractions using volumetric modulated arc therapy. After undergoing this highly conformal technique, the patient obtained prolonged local control and significant alleviation of symptoms with acceptable adverse events. This technique constitutes a promising approach for treating a complex target due to its ability to provide homogeneous coverage of irregularly shaped target volumes along with its ability to preserve organs at risk. In addition, we systematically reviewed clinical reports on the management of extensive cutaneous lesions in MF patients undergoing other irradiation techniques.

Introduction

Although mycosis fungoides (MF) is a rare disease with a relatively good prognosis when treated at an early stage, extensive cutaneous lesions can impair a patient’s quality of life (QOL) [1-4]. Radiation therapy is one of the most effective treatment modalities for patients with all stages of MF, including symptomatic cutaneous lesions [5-7]. When skin lesions are solitary or few in number, local radiation therapy using electron or photon beam irradiation is generally indicated, while total skin electron beam therapy (TSEBT) is required when the skin lesions extend over the entire body [5-7]. However, when cutaneous lesions extend widely to areas that are uneven and with curving surfaces, such as the face and neck, these cases require advanced techniques of electron beam irradiation, which pose various challenges such as the heterogeneity of the target volume along with the long treatment time [8-10]. In order to achieve uniformity for the targeted lesions with complex shapes and for subjects with strict organ-at-risk dose constraints, new irradiation techniques have been developed. These new methodologies include intensity-modulated radiotherapy (IMRT) and volumetric modulated arc therapy (VMAT), with these techniques now used during daily clinical practice for cases of advanced head and neck cancer and brain tumors [11, 12]. This report presents data for a successful clinical treatment of extensive MF on the scalp, face, and neck using VMAT with 20 Gy administered in 10 fractions.

Case Presentation

An 82-year-old man presented with a tumor in the posterior auricular region and a three-year history of erythematous patches and plaques on the trunk, extremities, and face. A skin biopsy from a cutaneous chest lesion was performed and a histological evaluation of the specimen revealed the infiltration of atypical lymphocytes into the epidermis, which was positive for CD3, CD4, and CD5, and negative for CD79a, CD8, and CD20 (L26). Atypical lymphocytes were not seen in the peripheral blood. The serum level of the soluble interleukin-2 receptor was elevated at 848 U/mL, and anti-human T-cell leukemia virus 1 and 2 antibodies were not detected.

The patient was diagnosed with MF T3N0M0B0 Stage2B according to the International Society for Cutaneous Lymphomas/European Organization of Research and Treatment of Cancer (ISCL/EORTC) revised classification system. Although he was treated with chemotherapy (bexarotene) and skin-directed therapies such as topical agents and phototherapy, the cutaneous lesions became resistant to these therapies and the side effects such as renal impairment and hypertriglyceridemia made it difficult to administer adequate
doses of chemotherapy. Patches and plaques gradually increased, most prominently on the facial skin and scalp, and this appearance of cutaneous lesions such as erythema and alopecia impaired the QOL of the patient (Figure 1A, 1B). The patient had no previous history of radiation therapy and was switched to radiation therapy using VMAT.

FIGURE 1: Clinical images
Prior to the treatment, observed patches and plaques were found to have spread all over the scalp and on the skin of the face to the neck (A, B). Although at one week after treatment grade 1 radiation dermatitis, grade 2 alopecia, and mild dilation of the subcutaneous veins were observed, there was a tendency for the cutaneous lesions to be resolved (C, D). At four months after the treatment, there was complete response observed for the cutaneous lesions in addition to the resolution of the alopecia (E, F).

Computed tomography (CT) simulation was performed in the supine position at a slice thickness of 2 mm. After immobilization of the subject using a type S shell (Toyo Medic), a 0.5 cm bolus was placed to fit the patient, with a second type S shell then immobilized over the bolus (Figure 2). VMAT plan was created using the Pinnacle treatment planning system version 9.10 (Philips). The clinical target volume (CTV) included a 4 mm area ranging from the skin surface of the entire scalp and from the face to the neck. The planning target volume (PTV) was defined as the CTV plus a 5 mm margin. The PTV was cropped by a 3 mm margin from the skin surface to form PTV evaluate (PTVeval) structure, which was used for dose normalization. A dose of 20 Gy in 10 fractions was prescribed to the mean dose of the PTVeval using a 6 MV photon beam from Versa HD (Elekta). VMAT plan was designed by 3 arc rotation fields to deliver a highly conformal dose distribution to the target with complex shapes, while minimizing the dose delivered to the organs at risk such as brain and parotid glands. Lenses and lacrimal glands were not spared due to eyelids involvement. Figure 3 shows the dose distribution and dose-volume histogram of the VMAT plan. The irradiation time was 555 seconds, with a total time starting from the time the patient entered the treatment room until leaving, of approximately 20 minutes. Toxicity evaluation was defined with the CTCAE v5 scale. Radiation therapy was performed in combination with the administration of low-dose bexarotene to control cutaneous lesions outside the irradiated area, with no adverse events observed during the treatment period. At one week after completion...
of the radiation therapy, the patient presented with grade 1 radiation dermatitis, grade 2 alopecia, and mild
dilation of the subcutaneous veins, with cutaneous lesions observed to be less erythematous along with the
absence of any bone marrow suppression (Figure 1C, 1D). Two months after completion of the radiation
therapy, the cutaneous lesions had almost completely disappeared, although there was slight redness
remaining in some areas. Four months after completion of the radiation therapy, complete response (CR)
was observed for the cutaneous lesions and the patient’s normal hair had grown back (Figure 1E, 1F). At 16
months of completion of the radiation therapy, CR remained for the lesions within the irradiated area, and
there were no observed late adverse events resulting from the administered irradiation.

**FIGURE 2: Patient immobilization device**

After being immobilized with a type S shell (Toyo Medic), a 0.5 cm bolus was placed to fit the patient, with a
second type S shell then immobilized over the bolus (A, B). Patient positioning at CT simulation (C).
Discussion

MF is a peripheral T-cell lymphoma that primarily affects the skin and which is often treated using local electron beam irradiation or TSEBT [5-7]. However, with recent advances in irradiation techniques, the efficacy of total scalp irradiation using IMRT, VMAT, or total skin helical tomotherapy (TSHT) has been reported [13-17]. Moreover, these newer irradiation techniques have made it possible to deliver uniform doses to complexly shaped skin surfaces, thereby sparing the adjacent normal tissue and critical structures [14-17]. In addition, these treatments require less treatment time and can be performed in a supine position, which can help to reduce patient burden as compared to that associated with the use of complex electron beam irradiation.
Furthermore, some authors have reported good outcomes for cutaneous T-cell lymphoma of the scalp when using IMRT treatments, in conjunction with clinically acceptable adverse events [13-15]. While the efficacy of TSHT has also been reported in several cases, photon beams are more likely to cause severe bone marrow suppression as compared to electron beams due to the increased dose that is administered to the deeper parts of the body [16-19]. Death due to bone marrow suppression associated with TSHT has been previously reported [18].

Furthermore, it has also been reported that MF patients developed bone marrow suppression including grade 4 thrombocytopenia after TSHT despite receiving bone marrow mean doses (arms not included) that were as low as 1.66 and 2.3 Gy, respectively [19]. In our patient, although the lesion had spread to the trunk as well, we decided to only use VMAT to treat the skin on the head, face, and neck due to the consideration that irradiation to a wider area could potentially cause bone marrow suppression, with increases in the lesions on the scalp and face also reducing the QOL. Even at the previously reported lower dose of 20 Gy, we were able to obtain a relatively long-term local control, and furthermore, with the additional application of VMAT to the local lesions, there were minimal adverse events along with high patient satisfaction [5-7].

The advantages of treating our current case with VMAT were the shorter treatment times, the less laborious setup, and the high dose homogeneity to the target. These results suggest that this approach can be utilized for treating patients with other cutaneous malignancies that include the extensive spread of lesions on the scalp and face.

**Conclusions**

MF is a highly radiosensitive disease and radiation therapy plays an important role in the management of patients with all stages of MF. VMAT can deliver a highly conformal dose to the target with complex shapes, while minimizing the dose delivered to the organs at risk. For MF with extensive lesions that ranged over an area from the skin of the scalp to the face and neck, low-dose radiation therapy with VMAT was found to be very useful and led to excellent local control with little toxicity.

**Additional Information**

**Disclosures**

**Human subjects:** Consent was obtained or waived by all participants in this study. Ethical Committee of Osaka Metropolitan University Graduate School of Medicine issued approval 2021-275. **Conflicts of interest:** In compliance with the ICMJE uniform disclosure form, all authors declare the following: **Payment/services info:** All authors have declared that no financial support was received from any financial institutions that submitted work. **Financial relationships:** All authors have declared that they have no financial relationships or activities that could appear to have influenced the submitted work.

**References**

1. Maguire A, Puellés J, Raboisson P, Chavda R, Gabriel S, Thornton S: Early-stage mycosis fungoides: epidemiology and prognosis. Acta Derm Venereol. 2020, 100:adv00013. 10.2540/00015555-5367
2. Talpur R, Singh L, Daulat S, et al.: Long-term outcomes of 1,263 patients with mycosis fungoides and Sézary syndrome from 1982 to 2009. Clin Cancer Res. 2012, 18:5051-60. 10.1158/1078-0432.CCR-12-0604
3. Demirer MF, Gan S, Jones J, Miller DR: Significant impact of cutaneous T-cell lymphoma on patients’ quality of life: results of a 2005 National Cutaneous Lymphoma Foundation Survey. Cancer. 2006, 107:2504-11. 10.1002/cncr.22252
4. Molloy K, Jonak C, Woiw A-Jin FJ, et al.: Characteristics associated with significantly worse quality of life in mycosis fungoides/Sézary syndrome from the Prospective Cutaneous Lymphoma International Prognostic Index (PROCLIP) study. Br J Dermatol. 2020, 182:770-9. 10.1111/bjd.18089
5. Sager O, Gamsiz H, Dinçoglu F, Demiral S, Uysal B, Beyzaedeoglu M: Radiotherapy for mycosis fungoides: review of the literature. J Dermatol Res Ther. 2017, 3:45. 10.23957/2469-5750/1510045
6. Tandberg DJ, Cricaciucescu O, Kelsey CR: Radiation therapy for cutaneous T-cell lymphomas. Dermatol Clin. 2015, 33:703-13. 10.1016/j.det.2015.05.006
7. Specht L, Dabaja B, Illidge T, Wilson LD, Hoppe RT: Modern radiation therapy for primary cutaneous lymphomas: field and dose guidelines from the International Lymphoma Radiation Oncology Group. Int J Radiat Oncol Biol Phys. 2015, 92:32-9. 10.1016/j.ijrobp.2015.01.008
8. Song IH, Jung JY, Park JW, Lee GW, Chae SM, Kay CS, Son SH: Dosimetric comparison of three different treatment modalities for total scalp irradiation: the conventional lateral photon-electron technique, helical tomotherapy, and volumetric-modulated arc therapy. J Radiat Res. 2015, 56:717-26. 10.1093/jrr/rru049
9. Able CM, Mills MD, McNeese MD, Hogstrom KR: Evaluation of a total scalp electron irradiation technique. Int J Radiat Oncol Biol Phys. 1991, 21:1063-72. 10.1016/0360-3016(91)90751-o
10. Lozano F, Perez N, Iglesias A, et al.: Volumetric arc therapy for total scalp irradiation: case report for a recurrent basal cell carcinoma of the scalp. Ecaneremedicalscience. 2017, 11:737. 10.3332/ecancer.2017.737
11. Wang X, Eshbruch A: IMRT for head and neck cancer: reducing xerostomia and dysphagia. J Radiat Res. 2016, 57 Suppl 1:s69-75. 10.1093/jrr/rrv047
12. Bas Ayata H, Ceylan C, Kölcü A, Güden M, Engin K: Comparison of multiple treatment planning techniques for high-grade glioma tumors near to critical organs. Oncol Res Treat. 2018, 41:514-9. 10.1159/000487642
13. Samant RS, Fox GW, Gerig LH, Montgomery LA, Allan DS: Total scalp radiation using image-guided IMRT for progressive cutaneous T cell lymphoma. Br J Radiol. 2009, 82:e122-5. 10.1259/bjr/6138056

14. Stang K, Allte F, Steber J, Emami B, Surucu M: Leukemia cutis of the face, scalp, and neck treated with non-coplanar split field volumetric modulated arc therapy: a case report. Cureus. 2015, 7:e430. 10.7759/cureus.450

15. Ostheimer C, Janich M, Hübsch P, Gerlach R, Vordermark D: The treatment of extensive scalp lesions using coplanar and non-coplanar photon IMRT: a single institution experience. Radiat Oncol. 2014, 9:82. 10.1186/1748-717X-9-82

16. Haraldsson A, Engleson J, Bäck SÅJ, Engelholm S, Engström PE: A Helical tomotherapy as a robust low-dose treatment alternative for total skin irradiation. J Appl Clin Med Phys. 2019, 20:44-54. 10.1002/acm2.12579

17. Yonekura K, Ichiki M, Takeda K, Uchiyama N, Nishida H, Dokiya T: Successful treatment of tumor stage mycosis fungoides with total skin helical tomotherapy. J Dermatol. 2022, 49:289-93. 10.1111/1346-8138.16246

18. Kitaguchi M, Yamashita H, Takenaka R, Okuma K, Nawa K, Nakagawa K: Helical skin radiation therapy including total skin radiation therapy using tomotherapy for primary cutaneous lymphoma with bone marrow suppression as a related adverse event. Pract Radiat Oncol. 2021, 11:e508-21. 10.1016/j.prro.2020.11.007

19. Schaff EM, Rosenberg SA, Olson SJ, Howard SP, Bradley KA: Bone marrow suppression as a complication of total skin helical tomotherapy in the treatment of mycosis fungoides. Radiat Oncol. 2018, 15:67. 10.1186/s13014-018-1013-2