Optimizing Care and Compliance for the Treatment of Mycosis Fungoides Cutaneous T-Cell Lymphoma With Mechlorethamine Gel

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Background: Mycosis fungoides is the most common form of cutaneous T-cell lymphoma (MF-CTCL). Quality nursing care is necessary for effective diagnosis and treatment of patients with MF-CTCL. Early-stage MF-CTCL (stages Ia and Ib) is most often managed in both dermatology and multidisciplinary settings. These stages can be effectively controlled by skin-directed therapies such as mechlorethamine gel (Valchlor®). Topical mechlorethamine has been used since the 1940s as an alkylating agent; however, compounded formulas have disadvantages that limit patient compliance. In contrast, newly approved mechlorethamine gel has demonstrated an efficacious and well-tolerated profile that has longer stability and is quicker to dry than its compounded counterpart.

Objectives: This article aims to provide recommendations for optimal nursing care of patients who have been diagnosed with stage Ia or Ib MF-CTCL.

Methods: Four real-world patient cases are examined, along with practical considerations for the use of mechlorethamine gel to treat patients with MF-CTCL.

Findings: Nurses can promote patient adherence through specific interventions and strategies, such as education about mechlorethamine gel, its mechanism of action, and safety and efficacy, as well as connecting patients with patient assistance programs or other supportive services.

Cutaneous T-cell lymphomas (CTCLs), primarily skin-affected non-Hodgkin lymphomas, represent about 70%–80% of all cutaneous lymphomas reported in the United States (Bradford, Devesa, Anderson, & Toro, 2009; Imam, Shenoy, Flowers, Phillips, & Lechowicz, 2013). Mycosis fungoides (MF-CTCL) is the most common type of CTCL (50%–70% of CTCL cases) and can present as cutaneous patches, plaques, or tumors, as well as Sézary syndrome (a more aggressive leukemic variant) (Bradford et al., 2009; Criscione & Weinstock, 2007; Imam et al., 2013). An estimated 16,000–20,000 cases of MF-CTCL are reported annually in the United States (Cutaneous Lymphoma Foundation, 2011; Lymphoma Research Foundation, 2012), with about 1,000–1,500 new MF-CTCL cases diagnosed per year (Bradford et al., 2009; Stanford School of Medicine Multidisciplinary Cutaneous Lymphoma Group, 2015). The five-year age-adjusted incidence of MF-CTCL for all stages has been reported as 0.55–0.56 per 100,000 person-years (Imam et al., 2013; Korgavkar, Xiong, & Weinstock, 2013).

Patients with early-stage disease (National Comprehensive Cancer Network [NCCN], 2014; Olsen et al., 2007) typically have a more favorable prognosis (median survival of 10–35 years) (Scarisbrick et al., 2014) (see Table 1). In several studies, patients with large cell transformation (a biopsy showing cells...
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<tr>
<th>Clinical Stage</th>
<th>Skin</th>
<th>Lymph Nodes</th>
<th>Viscera</th>
<th>Blood</th>
<th>Description</th>
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| Ia            | T1   | N0          | M0      | B0-1  | • Limited patches, papules, and/or plaques (less than 10% BSA)  
               |      |             |         |       | • No clinically abnormal (palpable; 1.5 cm diameter or greater) peripheral lymph nodes  
               |      |             |         |       | • No visceral organ involvement  
               |      |             |         |       | • Either absence of significant blood involvement (5% or less of peripheral blood lymphocytes are atypical/Sézary cells) or low blood tumor burden (greater than 5% of peripheral blood lymphocytes are atypical/Sézary cells), but does not meet B2 criteria |
| Ib            | T2   | N0          | M0      | B0-1  | • Patches, papules, or plaques covering 10% or greater BSA  
               |      |             |         |       | • No clinically abnormal (palpable; 1.5 cm diameter or greater) peripheral lymph nodes  
               |      |             |         |       | • No visceral organ involvement  
               |      |             |         |       | • Either absence of significant blood involvement (5% or less of peripheral blood lymphocytes are atypical/Sézary cells) or low blood tumor burden (greater than 5% of peripheral blood lymphocytes are atypical/Sézary cells), but does not meet B2 criteria |
| Ila           | T1-2 | N1-2        | M0      | B0-1  | • Limited patches, papules, and/or plaques  
               |      |             |         |       | • Clinically abnormal lymph nodes (histopathology Dutch grade 1 or 2, or NCI LN 0–3)  
               |      |             |         |       | • No visceral organ involvement  
               |      |             |         |       | • Either absence of significant blood involvement (5% or less of peripheral blood lymphocytes are atypical/Sézary cells) or low blood tumor burden (greater than 5% of peripheral blood lymphocytes are atypical/Sézary cells), but does not meet B2 criteria |
| IIb           | T3   | N0-2        | M0      | B0-1  | • One or more tumors at 1 cm or greater in diameter  
               |      |             |         |       | • Either no clinically abnormal (palpable; 1.5 cm diameter or greater) peripheral lymph nodes or clinically abnormal lymph nodes (histopathology Dutch grade 1–2 or NCI LN 0–3)  
               |      |             |         |       | • No visceral organ involvement  
               |      |             |         |       | • Either absence of significant blood involvement (5% or less of peripheral blood lymphocytes are atypical/Sézary cells) or low blood tumor burden (greater than 5% of peripheral blood lymphocytes are atypical/Sézary cells), but does not meet B2 criteria |
| IIIa          | T4   | N0-2        | M0      | B0    | • Generalized erythroderma (80% or greater BSA)  
               |      |             |         |       | • Either no clinically abnormal (palpable; 1.5 cm diameter or greater) peripheral lymph nodes or clinically abnormal lymph nodes (histopathology Dutch grade 1–2 or NCI LN 0–3)  
               |      |             |         |       | • No visceral organ involvement  
               |      |             |         |       | • Absence of significant blood involvement (5% or less of peripheral blood lymphocytes are atypical/Sézary cells) |
| IIIb          | T4   | N0-2        | M0      | B1    | • Generalized erythroderma (80% or greater BSA)  
               |      |             |         |       | • Either no clinically abnormal (palpable; 1.5 cm diameter or greater) peripheral lymph nodes or clinically abnormal lymph nodes (histopathology Dutch grade 1–2 or NCI LN 0–3)  
               |      |             |         |       | • No visceral organ involvement  
               |      |             |         |       | • Low blood tumor burden (greater than 5% of peripheral blood lymphocytes are atypical/Sézary cells), but does not meet B2 criteria |
| IVa           | T1-4 | N0-2        | M0      | B2    | • Any skin involvement (patches, papules, and/or plaques, one or more tumors at 1 cm or greater in diameter, and/or generalized erythroderma [80% or greater BSA])  
               |      |             |         |       | • Either no clinically abnormal (palpable; 1.5 cm diameter or greater) peripheral lymph nodes or clinically abnormal lymph nodes (histopathology Dutch grade 1–2 or NCI LN 0–3)  
               |      |             |         |       | • No visceral organ involvement  
               |      |             |         |       | • High blood tumor burden defined as one of the following: 1,000 Sézary cells/mcl or greater with positive clonal rearrangement of TCR; CD4:CD8 ratio of 10 or greater with positive clone; or CD4+/CD7- cells 40% or greater or CD4+/CD26- cells 30% or greater with positive clone |

BSA—body surface area; LN—lymph nodes; NCI—National Cancer Institute

TABLE 1. Clinical Stages of Mycosis Fungoides Cutaneous T-Cell Lymphoma (Continued)

<table>
<thead>
<tr>
<th>Clinical Stage</th>
<th>Skin</th>
<th>Lymph Nodes</th>
<th>Viscera</th>
<th>Blood</th>
<th>Description</th>
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<tbody>
<tr>
<td>IVa2</td>
<td>T1-4</td>
<td>N3</td>
<td>M0</td>
<td>B0-2</td>
<td>Any skin involvement (patches, papules, and/or plaques, one or more tumors at 1 cm or greater in diameter, and/or generalized erythoderma [80% or greater BSA])</td>
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<td></td>
<td>Clinically abnormal lymph nodes (histopathology Dutch grade 3–4 or NCI LN 4; clone positive or negative)</td>
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<td>Either absence of significant blood involvement (5% or less of peripheral blood lymphocytes are atypical/Sézary cells) or presence of cancerous cells in the blood (low blood tumor burden: greater than 5% of peripheral blood lymphocytes are atypical/Sézary cells, or high blood tumor burden defined as one of the following: 1,000 Sézary cells/ml or greater with positive clonal rearrangement of TCR; CD4:CD8 ratio of 10 or greater with positive clone; or CD4+/CD7- cells 40% or greater or CD4+/CD26- cells 30% or greater with positive clone)</td>
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<tr>
<td>IVb</td>
<td>T1-4</td>
<td>N0-3</td>
<td>M1</td>
<td>B0-2</td>
<td>Any skin involvement (patches, papules, and/or plaques, one or more tumors at 1 cm or greater in diameter, and/or generalized erythoderma [80% or greater BSA])</td>
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<td></td>
<td>Either no clinically abnormal (palpable; 1.5 cm or greater in diameter) peripheral lymph nodes or abnormal lymph nodes (histopathology Dutch grade 1–4 or NCI LN 0–4; clone positive or negative)</td>
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<td></td>
<td></td>
<td>Visceral involvement (pathology confirmation of specific organ involved)</td>
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<td>Either absence of significant blood involvement (5% or less of peripheral blood lymphocytes are atypical/Sézary cells) or presence of cancerous cells in the blood (low blood tumor burden: greater than 5% of peripheral blood lymphocytes are atypical/Sézary cells, or high blood tumor burden defined as one of the following: 1,000 Sézary cells/ml or greater with positive clonal rearrangement of TCR; CD4:CD8 ratio of 10 or greater with positive clone; or CD4+/CD7- cells 40% or greater or CD4+/CD26- cells 30% or greater with positive clone)</td>
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BSA—body surface area; LN—lymph nodes; NCI—National Cancer Institute


four times larger or greater than a small lymphocyte in 25% or more of dermal infiltrate (Salhany et al., 1988) have shown poorer outcomes (Scarisbrick et al., 2014). Decreases in overall survival, overall disease-specific survival, and median survival have been seen with relative risk for disease-related death (21.6 times more likely in patients with stage IV compared with stage Ia) (Agar et al., 2010).

Delayed diagnosis or misdiagnosis leading to under-reporting and incorrect treatment may occur because MF-CTCL can present similarly as other types of skin conditions such as psoriasis, vitiligo, chronic contact dermatitis, eczema, folliculitis, pityriasis lichenoides chronica, pigmented purpuric dermatoses, and pityriasis lichenoides et varioliformis acuta (Girardi, Heald, & Wilson, 2004; Sarantopoulos et al., 2013; Smedby et al., 2008; Zackheim, & McCalmon, 2002). To diagnose MF-CTCL, the NCCN (2014) guidelines recommend biopsy and subsequent dermatopathology, immunohistochemistry, and molecular analysis (T-cell receptor [TCR] gene rearrangement) of suspicious skin sites. Physical examination, laboratory tests (complete blood count with manual differential [CBC diff], lactic acid dehydrogenase [LDH], TCR gene rearrangement, Sézary flow cytometry), and, if more advanced disease is suspected, human T-cell lymphotropic virus serology and bone marrow or lymph node biopsy also are important factors in diagnosis.

Treatment for MF-CTCL is contingent on disease stage and patient response. The goals of treatment are to relieve symptoms and to achieve remission and delay progression while minimizing the emergent side effects related to treatment. Therapeutic options for early-stage MF-CTCL include topical treatments such as corticosteroids, mechlorethamine (Mustargen®), and carmustine (BCNU®); localized radiation; retinoids; phototherapy, such as psoralen plus ultraviolet A (PUVA) or ultraviolet B; and total skin electron beam. Systemic treatments such as oral retinoids, interferons (INFα, INF-γ), histone deacetylase inhibitors, extracorporeal photopheresis, or single- or multi-agent chemotherapy regimens also may be used if blood involvement is positive (Girardi et al., 2004; NCCN, 2014; Reavely & Wilson, 2004).

In advanced disease, treatments that reduce tumor burden or delay disease progression are implemented (Jawed, Mykowskki, Horwitz, Moskowitz, & Querfeld, 2014). Patients with advanced disease may receive more aggressive single-agent or multimodal strategies such as radiation, systemic first-line therapies (gemcitabine [Gemzar®], liposomal doxorubicin [Adriamycin®]), or second-line therapies (pentostatin [Nipent®], chlorambucil [Leukeran®], temozolomide [Temodar®], cyclophosphamide [Cytoxan®], etoposide [VePesid®], higher doses of methotrexate [Rheumatrex®], or low-dose plaralatrex [Folotyn®]) (Jawed et al., 2014; NCCN, 2014).

Topical mechlorethamine has been used to treat leukemic diseases since the 1940s (Goodman et al., 1946) and has been recommended as a primary treatment option for MF-CTCL (NCCN, 2014; Trautinger et al., 2006). Mechlorethamine is an
alkylating agent that inhibits quickly proliferating cells and, possibly via immune mechanisms, affects keratinocyte-Langerhans cell–T-cell interactions (Kim et al., 2005). Initially prescribed as compounded aqueous- or petrolatum-based ointment, these mechlorethamine formulations had disadvantages (e.g., greasy, less stable, more difficult to apply) that limited patient adherence (Kim, Martinez, Varghese, & Hoppe, 2003; Ramsay, Parnes, & Dubin, 1984). The U.S. Food and Drug Administration approved a topical mechlorethamine gel (Valchlor®) formulation in 2013 for patients diagnosed with stage Ia and Ib MF-CTCL who have received prior skin-directed therapy. Overall clinical experience (Kim, Duvic, Guitart, & Lessin, 2014; Lessin et al., 2013) has shown that mechlorethamine gel is generally well tolerated and similar in effectiveness to compounded mechlorethamine ointment. The most common adverse effects seen in the clinical trials were mild to moderate in severity and manageable and included pruritus, dermatitis, and hyperpigmentation. Following topical mechlorethamine use, nonmelanoma skin cancers have been reported in patients who received treatments known to cause nonmelanoma skin cancer, such as long-term phototherapy (particularly PUVA), radiation, and immunosuppressive chemotherapy; however, topical mechlorethamine did not show an increased risk of secondary cancers or comorbidities (Lindahl, Fenger-Grøn, & Iversen, 2014). Patients should be monitored for redness, inflammation, swelling, itchiness, blisters, secondary skin infections, ulceration, and nonmelanoma skin cancers during and following treatment with mechlorethamine gel. Contrary to mechlorethamine compounded in ointment, mechlorethamine gel has longer stability and is a quick-drying, greaseless gel (Actelion Pharmaceuticals Inc., 2013).

Although guidelines are available to help stage MF-CTCL, institutions may follow their own protocol for MF-CTCL diagnosis, treatment, patient follow-up, and maintenance plans. Real-world patient cases along with practical management tips are presented in this article to help healthcare providers recognize the signs and symptoms of the disease and educate nurses about topical mechlorethamine applications for use in the MF-CTCL treatment plan.

Case Reports

Case Study 1

A 60-year-old Caucasian man, Fitzpatrick skin type 2 (a numerical classification schema for human skin color), body mass index (BMI) 27 kg/m², presented with a persistent, non-pruritic rash (see Figure 1). Past medical history was remarkable for nine years of atopic dermatitis with generalized eczematous eruption treated by over-the-counter moisturizers. Physical examination in August 2013 revealed 1–3 cm atrophic erythematous patches with a body surface area (BSA) of 5%–8%. The lesions were thin, scaly, and annular/arciform, located at upper inner thighs and buttocks. The patient described the eruption as having a waxing and waning clinical course, improving after sun exposure and worsening after shower. The patient was bothered by the physical appearance of the rash and had irritation and discom- fort when wearing clothing.

In August 2013, the patient was diagnosed with MF-CTCL stage Ia (T1N0M0B0). Skin biopsy of a representative patch revealed infiltrate of small-sized lymphocytes with irregular nuclear contours, condensed chromatin, scant cytoplasm, indistinct nucleoli present at the dermal-epidermal junction (tagging), focally extending into the epidermis and adnexal structures. A superficial dermal and perivascular lymphoid infiltrate also was present. Immunoperoxidase studies demonstrated that the lymphoid infiltrate was composed primarily of CD3-positive T-cells with intact expression of CD2 and CD5, and CD7 loss. The majority of the T-cells, including those at the dermal-epidermal junction and extending into the epidermis, expressed CD4 (CD4:CD8 ratio of about 8:1).

Relevant laboratory tests, including CBC diff and platelet count, liver function tests (LFT), and LDH were all found to be within normal limits. Flow cytometry (CD4+/CD7- or CD4+/CD26-) revealed no evidence of blood involvement.

Following the diagnosis of MF-CTCL, a three-month trial of topical corticosteroids twice per day achieved only a partial response and a few new sites of emerging disease appeared. Topical corticosteroid treatment was discontinued and the patient was switched to mechlorethamine gel applied nightly. The patient was instructed to apply a thin film of mechlorethamine gel nightly to affected areas of the skin and allow the skin to dry completely for at least four hours before or 30 minutes after a shower or washing. The patient was informed to continue treatment for three months, unless any adverse reactions occurred such as redness, itching, swelling, blisters, ulcers, and/or skin infections, which would require a return to the clinic for follow-up. The patient was informed that a positive response may include pigment changes (darker or lighter), a smoother texture, and/or no new patches emerging. No concomitant therapy was given.

At the time of this writing, the patient has tolerated treatment for five months with no dermatitis. The patches have faded, smoothed, have a mild-brawny hyperpigmentation, and

FIGURE 1. Case Study 1 Patient
Note. Photograph courtesy of Dana-Farber/Brigham and Women’s Cancer Center. Used with permission.
the involved BSA has been reduced from 5%–8% to 2%, indicating stable-to-improved disease. The patient was educated that a mild degree of hyperpigmentation may be a positive response to treatment. The patient’s quality of life has improved because clothing is more comfortable to wear and the patient is less distressed about the appearance of the patches. The treatment plan is to continue the mechlorethamine gel treatment until patches are cleared (complete response).

Case Study 2

A 45-year-old Hispanic man, Fitzpatrick skin type 6, BMI of 29 kg/m², presented with hyperpigmented patches and moderate pruritus. Past medical history included a diagnosis of dermatitis 15 years prior, having a waxing and waning clinical course, but worsening during the past two years. Physical examination revealed mildly xerotic, darkly hyperpigmented patches on the upper inner arms, abdomen, back, buttocks, and upper thighs with a 25%–30% BSA (see Figure 2). Lesions were distributed over axilla, upper inner arms, lower abdomen, lower back, buttocks, and hips. The patient described his skin as inflamed and tender to the touch during the past six months, resulting in disrupted sleep and difficulty in working (the patient worked many hours as a manual laborer). His wife attends his clinical visits and she also aids in applying topical treatments.

Topical corticosteroids applied twice per week as pulse therapy (two weeks on treatment/two weeks off treatment) and an oral antihistamine provided relief of pruritus initially, but more recent response was poor and the patient discontinued the corticosteroids. Compounded mechlorethamine ointment (10 mg%) was used three years ago but was discontinued because of dermatitis.

In February 2014, the patient was diagnosed with MF-CTCL stage Ib (T2N0M0B0). A 4 mm skin punch biopsy taken from two patches presenting on his back and thigh revealed infiltrate of small-sized lymphocytes with irregular nuclear contours, indistinct nucleoli, condensed chromatin, and scant cytoplasm present at the dermal-epidermal junction (tagging) and focally extending into the epidermis and adnexal structures. Superficial dermal and perivascular lymphoid infiltrate also was present. Immunoperoxidase studies demonstrated that the lymphoid infiltrate was composed primarily of CD3-positive T-cells with intact expression of CD2, CD5, and CD7. The majority of the T-cells, including those at the dermal-epidermal junction and extending into the epidermis, expressed CD4 (CD4/CD8 ratio of about 8:1). The TCR gene rearrangement studies identified a clonal T-cell receptor gamma chain gene rearrangement. Relevant laboratory tests revealed no elevation of white blood cells and no evidence of blood involvement.

Following the diagnosis, phototherapy was considered but was not a viable option because of geographic distance for the patient. Mechlorethamine gel was prescribed. Because of the previous dermatitis reaction following compounded mechlorethamine ointment use, the patient was instructed to patch test the mechlorethamine gel; the patient was instructed to test a small area on one patch by applying mechlorethamine gel nightly for 10 days. The patient reported no adverse reaction. After the successful patch test, the patient was instructed to apply mechlorethamine gel regionally to all affected areas nightly and a lighter amount to skin folds. He did not receive other concomitant therapy. The patient’s wife was instructed to wear gloves when applying to the patient’s difficult-to-reach lesions.

At the three-month follow-up visit, the patient reported a burning sensation at the treatment site, had moderate erythema and increased hyperpigmentation that was diagnosed as severe dermatitis. Mechlorethamine gel was discontinued and class one topical corticosteroids were applied twice per day for two weeks. The dermatitis resolved and the majority of lesions cleared. For the remaining lesions, topical corticosteroids were continued for an additional two weeks (applied once daily in the morning) and concomitant mechlorethamine gel was reinitiated (applied every other night).

The patient has tolerated treatment with mechlorethamine gel for three months with an improved disease response and no return of the dermatitis. The patches are reduced in size and are less inflamed, painful, and itchy with BSA affected by disease reduced from 25%–30% to 5%. The patient’s quality of life has improved; he is sleeping better and is able to work the long hours he would normally work. The plan is to continue mechlorethamine gel treatment until lesions are cleared (complete response). Topical corticosteroids will be used as needed for any early signs of dermatitis.

Case Study 3

An 87-year-old Caucasian man, Fitzpatrick skin type 2, BMI 24.5 kg/m², with a long-standing, stable 15-year history of MF-CTCL stage Ib presented with chronic scattered patches and plaques (see Figure 3). During this history, treatment with narrowband ultraviolet B phototherapy produced a significant reduction in patches and plaques (partial response). About two years ago, MF-CTCL was remarkable for tumors 1 cm or greater in diameter (erythematous raised nodules). Biopsy revealed
large cell transformation (epidermotropism by large atypical lymphoid, CD30+ cells). In light of the patient’s large cell transformation and multidisciplinary care (collaboration with the oncology department), the patient was re-staged to MF-CTCL stage IIb (T3N0M0B0; no lymph node or blood involvement). Localized electron beam treatment was employed to address the tumor stage manifestation on the back and he was treated with an oral retinoid per NCCN (2014) guidelines, producing a strong partial response. The tumors resolved; however, the patient continued to present with persistent plaques in a limited distribution on his forearms. The patient is a retired engineer and is independent in all aspects of daily living.

Physical examination in January 2013 revealed erythematous, scaly, annular (2–3 cm) plaques on his forearms at BSA of 3%–5%. No pruritus was noted. Following skin biopsy of multiple lesions, immunostains demonstrated a predominance of CD3+ epidermotropic atypical lymphoid cells. Rare scattered CD20+ B cells were noted. The CD4:CD8 ratio was markedly increased within the epidermis. There was a loss of staining with pan T-cell markers CD5 and CD7. Importantly, clusters and aggregates of large atypical lymphoid cells showed positive staining with antibodies to CD30. The CD30 staining was particularly remarkable.

Relevant laboratory tests revealed no elevation of white blood cells and no evidence of blood involvement. Computerized tomography and positron-emission tomography of the chest, abdomen, and pelvis were negative for lymphadenopathy and visceral involvement.

To address the several lingering plaques, mechlorethamine gel applied nightly was added to supplement the oral retinoid therapy. Six months after mechlorethamine gel was added to the treatment plan, the patient had tolerated treatment with no dermatitis or hyperpigmentation. Lesions were completely resolved. Mechlorethamine gel was discontinued and oral retinoid was continued as maintenance therapy to control the documented transformed (tumor stage) MF-CTCL.

At the time of this writing, the patient continues with long-term oral retinoid maintenance and the plaques have not returned to the affected sites. If the patient presents with new lesions, mechlorethamine gel will be reinitiated.

**Case Study 4**

A 49-year-old African American woman, Fitzpatrick skin type 5, BMI of 38.1 kg/m², with a three-year history of MF-CTCL stage Ib presented with scaling patches (see Figure 4) and mild pruritus. Co-morbidities include severe osteoarthritis, type 2 diabetes, and hypercholesteremia. Physical examination revealed confluent, hyperpigmented, mildly xerotic patches and plaques affecting 40% of the BSA. Lesions were distributed on the patient’s breasts, abdomen, back, and buttocks. Prior treatments included tropical corticosteroids and compounded mechlorethamine ointment. Because of financial constraints and cost of compounded medication, the patient had poor adherence with the compounded mechlorethamine ointment (10 mg%) and had discontinued therapy for several months. The patient’s daughter lives with the patient and helps her by applying the topical treatments to difficult-to-reach areas.

In July 2011, the patient was diagnosed with MF-CTCL stage Ib (T2N0M0B0); a skin punch biopsy from a patch on her
buttock revealed that the epidermis had an infiltration of small abnormal lymphoid cells with markedly irregular or convoluted nuclei. The cells lined up at the basal epidermis. Immunostains showed that the atypical lymphocytes were positive for CD2 and CD3, with partial loss of CD4, CD5, and CD7. Relevant laboratory tests revealed no elevation of white blood cells and no evidence of blood involvement.

Following the initial diagnosis in 2011, the patient received a three-month course of topical corticosteroids with no clearing of patches or plaques. Despite the previous intermittent use

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**Dermatitis**

- Dermatitis can present as dry, scaly patches that are pruritic and warm to the touch. Dermatitis may be a common occurrence in patients with MF-CTCL; therefore, patients should not be alarmed if dermatitis is experienced. Intense hyperpigmentation (particularly in patients with darker skin), erythema (redness), swelling, increased sensitivity, blisters, ulceration, pain, and burning also may accompany the dermatitis and treatment should be stopped or temporarily discontinued until symptoms have resolved.
- Symptoms can be alleviated by suggesting the patient soak in cool water for 10–20 minutes and then apply emollients, cold compresses for localized itch, oatmeal baths, mentholated cooling topicals, wet wraps, cool compresses, or ice packs. A daily moisturizer is recommended because dry skin tends to itch more.
- Patients may wish to wash their clothes in fragrance-free laundry detergent. Some healthcare providers suggest a bleach bath for their patients (2 tsp of bleach per gallon of water), which helps to reduce bacteria load on the skin and prevents infections. For patients with severe dermatitis, patients could apply topical corticosteroids or cover with plastic wrap or sauna suit.
- In some patients with MF-CTCL who have used mechlorethamine, dermatitis has been observed within the first couple of weeks followed by a clearing of the lesion. If dermatitis is seen, suggest stopping or tapering mechlorethamine use and apply a topical corticosteroid until dermatitis is resolved.
- Corticosteroid treatment could be stopped or given intermittently with mechlorethamine (corticosteroids in the morning and mechlorethamine at night). Corticosteroid treatment could be applied for 2–3 months continuously or as pulse therapy (e.g., 2–3 weeks on treatment, 2–3 weeks off treatment). Mechlorethamine could then be reintiated at the healthcare provider’s discretion, which could be at a reduced frequency (every other night or twice a week).

**Pruritus**

- Localized or generalized itching can be relieved by addressing the underlying disease process. Over-the-counter preparations (e.g., emollients, anti-pruritic, and anti-inflammatory) and/or systemic antihistamines, steroids, doxepin (Silenor®), gabapentin (Neurontin®), aprepitant (Emend®), mirtazapine (Remeron®), selective serotonin reuptake inhibitors, or naltrexone (Contravé®) could help alleviate symptoms.

**Travel**

- Patient travel can be a factor for refrigeration and storage of topical agents such as mechlorethamine gel. If taking a shorter trip, the patient could take an ice cooler with ice for carrying the mechlorethamine gel and coordinate refrigeration at the destination. The healthcare provider may need to write a note to the airline or Transportation Security Administration to allow the patient to bring the mechlorethamine gel on the plane.
- If the patient is traveling for a longer period of time and refrigeration is not available, possibly prescribe a topical corticosteroid instead for the duration of the trip and then switch back to mechlorethamine gel when the patient returns home.

**Quality-of-life issues (emotional distress, financial concerns)**


**Number of tubes to prescribe per percent of BSA**

- In a randomized, observer-blind, multicenter, noninferiority, active-controlled clinical trial of 260 patients with MF-CTCL to study the efficacy of mechlorethamine gel, the mean daily use of mechlorethamine gel was 2.8 g (1–2 tubes per month); maximum daily use was 10.5 g (5–6 tubes per month). Calculating the number of tubes per BSA, this equates to about one tube per 10% BSA per month.

**Timeline (response)**

- Once disease clears and, barring no adverse reactions, the patient should be treated with mechlorethamine for an additional month. A gradual decrease in dosing may be possible. Reinitiate treatment if disease reoccurs.

**Timeline (management)**

- Initial follow-up visit may be scheduled at 2–3 months with subsequent visits every 3–6 months if the patient is responding to treatment.
- In a clinical study of patients with MF-CTCL, patients who applied topical mechlorethamine as a maintenance treatment had a longer enduring response during maintenance therapy than those who did not apply topical mechlorethamine as a maintenance treatment. Patients may benefit from adding topical mechlorethamine as part of a long-term maintenance regimen. Patients also may respond well to topical mechlorethamine after relapse following more aggressive therapies.

**Differences between compounded mechlorethamine ointment and mechlorethamine gel**

- Mechlorethamine gel formulation is quick-drying, easy to apply, and does not stain clothing; however, it does require refrigeration. Patients also have been helped by patient assistance programs offered for mechlorethamine gel.

**Unpredictability of the disease**

- Patients may be concerned about the unpredictable nature of the disease (i.e., patients do not know when the disease will return). Patients should notify their healthcare provider about any changes in disease appearance or side effects. It also is important that the patient return for regularly scheduled follow-up visits.

**Patch test, treatment induction, and titration**

- Particularly if the patient has had a hypersensitive reaction to previous compounded mechlorethamine ointment, it may be prudent for the patient to test a small area initially. Patients could apply mechlorethamine gel nightly on a small area for as many as 10 days prior to applying to all areas. If the mechlorethamine gel appears to be tolerated, then treatment could be titrated (initially applied three times per week and add an additional day each subsequent week until applying every day).

**Cannot access area on the body**

- Apply using a back applicator or ask a caregiver to apply. Caregivers should wear disposable nitrile gloves when applying.

**When to apply**

- Some patients would rather apply daytime use or overnight. This depends on what the patient prefers with lifestyle and home situation.

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**FIGURE 5. Topics and Recommendations for Mycosis Fungoides Treatment With Mechlorethamine Gel**

of compounded mechlorethamine ointment, a strong partial response was seen: involved BSA was reduced to less than 10% BSA. The patient was restarted on therapy and was switched from compounded mechlorethamine ointment to the gel. She was advised to initially apply the gel three times per week and to add an additional day of treatment each subsequent week until applying every day, and in sparing amounts twice weekly to skin fold areas. The patient was instructed to apply regionally because of the generalized and confluent nature of the disease. She did not receive concomitant therapy. The daughter was educated to wear gloves when she applies the mechlorethamine gel to the difficult-to-reach areas.

The patient has tolerated treatment for nine months with no dermatitis or hyperpigmentation. The BSA of the lesions has been reduced from 40% to 10%, with resolution of the pruritus. The patient exhibits gradual improvement; disease regression is noted by skin smoothing, reduction in patch size, and no new areas of disease. The patient prefers the gel to the compounded ointment because it dries quicker, is easier to apply, does not stain her clothing, and the tube can be easily disposed. The plan is to continue the mechlorethamine gel treatment until lesions are cleared (complete response).

### Discussion

MF-CTCL is rare and potentially life-threatening. Given the chronicity, long-term nature, and quality-of-life impact of MF-CTCL, correct diagnosis and consistent compliance with treatments are necessary to achieve optimal clinical outcomes. Success in treating patients with stage Ia or Ib MF-CTCL is largely based on patient and caregiver education regarding MF-CTCL and proper treatment.

MF-CTCL is a spectrum disorder; therefore, a multidisciplinary team of dermatology nurses, nurse practitioners, oncology nurses, and physicians may be needed to ensure accurate diagnosis, develop individualized treatment and maintenance plans to prevent disease progression, and to treat the range of symptoms associated with MF-CTCL and its therapies. Early-stage MF-CTCL is most often successfully managed in ambulatory dermatology practices. Later stage or recalcitrant MF-CTCL may be managed in multidisciplinary settings. As part of the multidisciplinary team, nurses educate patients about MF-CTCL diagnosis and staging and provide reassurance about disease progress, symptom management, and response to treatment.

Photograph documentation of the disease before and after treatment can serve as a visual learning tool in the electronic health record and help track a patient’s recovery.

Nursing care measures can promote good patient adherence through specific interventions or strategies, such as educating the patient about topical agents (e.g., steroids, mechlorethamine), educating about the mechanism of action and correct application, and assessing the safety and efficacy of treatments. The overall clinical experience with the gel formulation of mechlorethamine has demonstrated a generally well-tolerated profile with most side effects being manageable and mild to moderate in severity. Nurses can connect patients with patient assistance programs offered for these treatments and/or other supportive services for managing MF-CTCL.

### Implications for Practice

- Educate patients about mechlorethamine gel (Valchlor®), its mechanism of action, safety, and efficacy, and connect patients with patient assistance programs or other supportive services.
- Empower patients to be responsible for self-care by addressing manifestations of their disease when in the home setting.
- Integrate mechlorethamine gel as part of a skin-directed treatment strategy for patients with stage Ia or Ib mycosis fungoides cutaneous T-cell lymphoma.

### Conclusion

Nurses’ education of patients regarding topical agents may lead to positive outcomes, such as empowering the patient to be responsible for self-care. Topical agents allow patients to address manifestations of their disease at home, which may reduce travel time to the clinic or phototherapy treatment center and possibly improve patient adherence. Nurses’ assessment of the effectiveness of treatment for patients with MF-CTCL helps guide future decision making and improves patients’ independence and quality of life.

### References


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