Folliculotropic Mycosis Fungoides

Single-Center Study and Systematic Review

Julia S. Lehman, MD; Robert H. Cook-Norris, MD; Brent R. Weed, MD; Roger H. Weenig, MD; Lawrence E. Gibson, MD; Amy L. Weaver, MS; Mark R. Pittelkow, MD

Objectives: To clarify clinicopathologic features and reconcile discrepancies in previous studies of folliculotropic mycosis fungoides (FMF).

Design: A single-center retrospective clinicopathologic study and a systematic review of FMF.

Setting: Tertiary referral center in the midwestern United States.

Patients: Patients with clinical and histopathologic evidence of FMF seen at the tertiary referral center during a 12½-year period.

Main Outcome Measures: Clinicopathologic features of FMF.

Results: Fifty patients (32 male [64%] and 18 female [36%]) met study criteria for the clinicopathologic review. Pruritic patches, plaques, and folliculocentric lesions (milia, cysts, and alopecia) on the head, neck, and trunk were common clinical findings. The mean time to diagnosis of FMF was 5.0 years. Diagnostic latency did not affect risk of death. One-year and 5-year overall survival rates were 96% and 62%, respectively. Frequent microscopic features were follicular mucinosis (74%) and epidermotropism (54%). Systematic review of 186 additional patients confirmed male predominance (ratio of men to women, 3.2:1.0), prevalent pruritus (73%), frequent follicular mucinosis (69%) and epidermotropism (37%) microscopically, and common head, neck, and trunk involvement. Combined data demonstrated that 6% of patients with FMF had concurrent non–mycosis fungoides hematologic malignant neoplasms and that the 5-year overall survival rate was 62% to 64%.

Conclusion: Folliculotropic mycosis fungoides has distinct clinical and microscopic features and is associated with a poor 5-year overall survival rate.

Arch Dermatol. 2010;146(6):607-613

FOLLICULOTROPIC MYCOsis fungoides (FMF) is an uncommon subtype of mycosis fungoides that seems to have a more aggressive natural history than other forms of mycosis fungoides as demonstrated in the results of stage-for-stage comparisons.1,2 The clinical presentation of FMF often differs from the patches and plaques of classic mycosis fungoides and may be associated with decreased clinical suspicion for mycosis fungoides. Moreover, histopathologic findings regarding FMF are varied and may be subtle. These factors may account for a prolonged time to diagnosis.

For editorial comment see page 662

The characteristics of FMF have been published in previous clinical and histopathologic studies and case series.1,4 The objectives of the present study were to corroborate or challenge previous observations of FMF, to reconcile inconsistencies in findings, and to expand existing knowledge regarding clinical, histopathologic, and therapeutic observations and biologic behavior. To achieve these goals, we reviewed the clinical and histopathologic features of FMF in patients who were seen at Mayo Clinic, Rochester, Minnesota, during 12½ years. We also conducted a systematic review and compared our results with those published in the medical literature.
Clinical features at the time of initial presentation are standardized descriptive statistics were used to summarize the data. For survival analysis, duration of follow-up was calculated from the date of the mycosis fungoides diagnosis to the date of the last follow-up or death. Overall survival was estimated using the Kaplan-Meier method. Adverse clinical outcomes were defined as the development of nodules, erythroderma, microscopically confirmed extracutaneous involvement, or death from disease. Time to the first adverse clinical outcome was calculated from the date of onset of skin lesions; for patients without an adverse clinical outcome, the interval was censored at the date of the last clinical follow-up. Separate univariate Cox proportional hazards regression models were fit to evaluate the association between patient characteristics and time to death or time to first adverse clinical outcome. The hazard ratio (HR) that was estimated from the Cox proportional hazards regression model and the corresponding 95% confidence interval (CI) were used to summarize the associations. All calculated P values were 2-sided, and P < .05 was considered statistically significant. Analyses were performed using commercially available statistical software (SAS, version 9.1; SAS Institute, Inc, Cary, North Carolina).

For our systematic review, the clinical, histopathologic, molecular, and prognostic data from our study were compared with those from previous retrospective clinical and histopathologic studies and case series of FMF, which we identified using PubMed searches and bibliographic cross-referencing. Search terms included folliculotropic mycosis fungoides, pilotropic mycosis fungoides, follicular mycosis fungoides, and adnexotropic mycosis fungoides. Studies describing 3 or more patients with FMF were included. Whenever possible, data were combined by weighted mean.

### RESULTS

Fifty patients met study inclusion criteria (Figure 1). Characteristics of 50 study patients (32 male and 18 female) are summarized in the first column of eTable 1 (http://www.archdermatol.com). The mean interval between onset of symptoms and diagnosis of mycosis fungoides was 3.9 years. The mean age at diagnosis of mycosis fungoides (57.7 years) was similar to the mean age at diagnosis of FMF (58.8 years). For 31 patients, folliculotropic lesions were recognized at or before the time of microscopic diagnosis of mycosis fungoides. The mean age at diagnosis of mycosis fungoides was significantly older for women than for men (63.1 years vs 54.6 years; P = .04, 2-sample t test). Five patients (4 male and 1 female) were younger than 40 years at the time of diagnosis of mycosis fungoides.

Of 41 patients whose medical record included staging data at the time of diagnosis, most patients (30 [73%]) had skin involvement only, without erythroderma. Four patients (10%) had generalized erythroderma. Six patients (15%) had clinical lymphadenopathy but negative microscopic findings from a lymph node biopsy, and only 1 patient (2%) had microscopically confirmed nodal involvement. One patient was found to have visceral involvement at the time of staging.

Clinical features at the time of initial presentation are listed in Table 1, and representative clinical images are shown in Figure 2. Of 47 patients whose symptoms were recorded, 44 (94%) had symptoms that were associated with their skin disease, including pruritus (35 patients [74%]) and pain (13 patients [28%]). Mucinorrhea (ie,
Table 1. Clinical Characteristics of Patients With Folliculotropic Mycosis Fungoides in the Present Study and in Previous Retrospective Studies and Case Series

<table>
<thead>
<tr>
<th>Variable</th>
<th>Present Study (n=50)</th>
<th>van Doorn et al,1 2002 (n=51)</th>
<th>Gerami et al,2 2008 (n=63)</th>
<th>Gerami and Guitart,3 2007 (n=34)</th>
<th>Cerroni et al,4 2002 (n=28)</th>
<th>Hodak et al,5 1999 (n=9)</th>
<th>Vergier et al,6 1996 (n=9)</th>
<th>Flaig et al,7 2001 (n=9)</th>
<th>Pereyo et al,8 1997 (n=3)</th>
<th>Combined Data, %</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td>Source, No./Total No. (%)</td>
<td>Source, No./Total No. (%)</td>
<td>Source, No./Total No. (%)</td>
<td>Source, No./Total No. (%)</td>
<td>Source, No./Total No. (%)</td>
<td>Source, No./Total No. (%)</td>
<td>Source, No./Total No. (%)</td>
<td>Source, No./Total No. (%)</td>
<td>Source, No./Total No. (%)</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Anatomical Area</td>
<td>Morphologic Features of Lesions</td>
<td>Comorbid Malignant Neoplasms</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Head and neck</td>
<td>29 (58)</td>
<td>45 (88)</td>
<td>37 (66)</td>
<td>20/25 (80)</td>
<td>14 (50)</td>
<td>2 (22)</td>
<td>NA</td>
<td>NA</td>
<td>3 (100)</td>
<td>72</td>
</tr>
<tr>
<td>Trunk</td>
<td>30 (60)</td>
<td>NA</td>
<td>41 (95)</td>
<td>NA</td>
<td>23 (82)</td>
<td>8 (89)</td>
<td>NA</td>
<td>NA</td>
<td>3 (100)</td>
<td>79</td>
</tr>
<tr>
<td>Extremities</td>
<td>38 (76)</td>
<td>NA</td>
<td>NA</td>
<td>9 (32)</td>
<td>6 (67)</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>3 (100)</td>
<td>62</td>
</tr>
<tr>
<td>Genitalia and buttocks</td>
<td>10 (20)</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>1 (11)</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>19</td>
</tr>
<tr>
<td>Patches, plaques, or papules</td>
<td>49 (98)</td>
<td>44 (86)</td>
<td>32 (74)</td>
<td>4/27 (15)</td>
<td>26 (93)</td>
<td>7 (78)</td>
<td>7 (78)</td>
<td>NA</td>
<td>2 (67)</td>
<td>78</td>
</tr>
<tr>
<td>Nodules</td>
<td>9 (18)</td>
<td>14 (27)</td>
<td>26 (60)</td>
<td>NA</td>
<td>0</td>
<td>1 (11)</td>
<td>NA</td>
<td>NA</td>
<td>1 (33)</td>
<td>27</td>
</tr>
<tr>
<td>Erythroderma</td>
<td>4 (8)</td>
<td>3 (6)</td>
<td>3 (7)</td>
<td>NA</td>
<td>2 (7)</td>
<td>0</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>7</td>
</tr>
<tr>
<td>Alopecia</td>
<td>19 (38)</td>
<td>NA</td>
<td>28 (65)</td>
<td>NA</td>
<td>NA</td>
<td>7 (78)</td>
<td>3 (33)</td>
<td>NA</td>
<td>NA</td>
<td>51</td>
</tr>
<tr>
<td>Folliculocentric lesions such as milia, cysts, and acneiform lesions</td>
<td>25 (50)</td>
<td>4 (8)</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>9 (100)</td>
<td>9 (100)</td>
<td>NA</td>
<td>3 (100)</td>
<td>41</td>
</tr>
<tr>
<td>Mucinorrhea</td>
<td>6/47 (13)</td>
<td>3 (6)</td>
<td>A few cases</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>9</td>
</tr>
<tr>
<td>Pruritus</td>
<td>35/47 (74)</td>
<td>Most patients</td>
<td>30 (70)</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>NA</td>
<td>3 (100)</td>
<td>73</td>
<td></td>
</tr>
</tbody>
</table>

Abbreviation: NA, data were unavailable or were unavailable in a format amenable to direct comparison.

a Whether some of the patients in the study by Gerami et al2 also were reported in the study by Gerami and Guitart3 is unknown. Whether some of the patients in the study by Cerroni et al6 also were reported in the study by Flaig et al8 is unknown.

b Limited anatomic distribution of disease data available for 25 patients.
c Limited disease morphologic data available for 27 patients.
d When mucinorrhea was not addressed, it was assumed to be absent.

fluid expression from follicular ostia) was present in 6 patients (13%), of whom 5 had histopathologic evidence of follicular mucinosis. At presentation, 4 patients had erythroderma. During the follow-up period, 1 patient demonstrated the development of erythroderma and the presence of Sézary cells, which were identified on peripheral smear.

Among 50 patients in the study, the most common dermatologic diagnoses attributed to the eruptions (diagnoses later determined to be FMF) included the following: dermatitis (25 patients [50%]), follicular mucinosis (11 patients [22%]), dermatophyte infection (8 patients [16%]), psoriasis (6 patients [12%]), and folliculitis (4 patients [8%]). Several patients had more than 1 previous dermatologic diagnosis. One patient had a forehead nodule diagnosed previously as unspecified peripheral T-cell lymphoma.

Seven patients (14%) had a concomitant malignant neoplasm, and 1 of these 7 patients had both breast cancer and a B-cell lymphoproliferative disorder (Table 1). Hematologic comorbidities included chronic lymphocytic leukemia (2 patients [4%]) and unspecified B-cell lymphoproliferative disorder (1 patient [2%]). Oncologic comorbidities included breast cancer (2 patients [4%]) and bladder cancer, lung adenocarcinoma, and prostate cancer (1 patient each [2%]). One patient had monoclonal gammopathy of undetermined significance.

SYSTEMIC INVOLVEMENT DATA

Twelve patients (24%) with localized FMF had distant lymph node involvement, which was detected radiographically (9 patients [18%]), histologically (6 patients [12%]), or in both ways. Peripheral smear results for 37 patients indicated that 11 patients (30% of those tested) had unequivocal evidence of circulating convoluted lymphocytes (Sézary cells); 2 of these patients also had generalized erythroderma. Four of 37 patients who had peripheral smears had equivocal evidence of Sézary cells, and 1 of these patients had generalized erythroderma. One of 13 patients who underwent bone marrow biopsy had disease involvement of the marrow.

HISTOPATHOLOGIC CHARACTERISTICS

Histopathologic features are summarized in eTable 2, and representative photomicrographs are shown in Figure 3. Prominent syringotropism of atypical lymphocytes without folliculotropism was noted in 1 biopsy specimen, and prominent syringotropism was present in another biopsy specimen in which no hair follicle could be found in numerous sections. Both patients had clinical evidence of follicular involvement (associated alopecia) and were included in the analysis. Immunohistochemical data were available for biopsy specimens from 42 patients (84%), with staining of neoplastic lymphoid cells in all specimens indicating a CD3−CD4+CD8− immunophenotype. The presence of tissue eosinophilia had low sensitivity (19%) but much higher specificity (70%) for predicting pruritus.

MOLECULAR DATA

Flow cytometric immunotyping was performed on peripheral blood of 31 patients (62%), resulting in the identification of an unequivocal circulating clone in the specimens from 7 patients (23% of those tested). For 3 patients, the circulating clone was attributed to a lymphoproliferative malignant neoplasm other than mycosis fungoides. To detect TCR gene rearrangement clones, polymerase chain reaction or flow cytometry studies were performed on the peripheral blood of 17 patients (34%). A clonal population of T cells was identified in the peripheral blood of 8 patients (47% of those tested), one of whom had another lymphoproliferative malignant neoplasm. Polymerase chain reaction or flow cytometry studies were performed on the lesional skin of 21 patients, revealing a clone in the specimens from 15 patients (71%). Of 9 patients whose peripheral blood and lesional skin were tested for TCR gene rearrangement, 6 had a clone detected in both tissues. Of 4 patients in whom the peripheral blood clone was compared with that of the skin, 2 had identical clones. Of patients who had both skin and peripheral blood specimens tested for TCR gene rearrangement clones, all were found to have positive clones in lesional skin tissue.

OVERALL FOLLOW-UP AND SURVIVAL

General follow-up data were available for 47 patients (94%), and treatment data were available for 43 patients (86%). Among 15 patients known to be deceased, the
mean follow-up period from mycosis fungoides diagnosis to death was 3.3 years (range, 0-20.4 years). Among 32 patients alive at last follow-up, the median duration from diagnosis of mycosis fungoides to last follow-up was 2.9 years (range, 4 days to 14.7 years). The overall survival rates at 1, 2, and 5 years after diagnosis were 96%, 90%, and 62%, respectively (Table 2 and Figure 4). The cause-specific survival rate was not calculated because only 3 patients died of mycosis fungoides. The interval between onset of skin lesions and diagnosis did not affect risk of death (HR per 1-year increase in time to diagnosis, 1.02; P = .77).

TREATMENT AND FOLLOW-UP

Initial treatment data are given in Table 2. The mean number of individual treatments per patient was 4.2 (range, 1-10; median, 4.0). Fifteen patients (30% of all 50 patients) were hospitalized for management of their cutaneous disease at some point in the follow-up period. At the time of their last clinical assessment, 21 of 43 patients (49%) for whom treatment follow-up data were available had improved partially or completely. Five of 43 patients (12%) had complete or near-complete resolution of skin lesions at least once in the follow-up period; 1 patient had disease remission throughout the follow-up period. These 5 patients were treated using topical corticosteroids with l-lysine hydrochloride, psoralen–UV-A photopheresis, topical or oral retinoids, and no therapy.

PROGNOSTIC ASSOCIATIONS

Among 50 patients identified for the study, 15 (30%) had an adverse clinical outcome. The median time from onset of skin lesions to adverse clinical outcome was 3.4 years (range, 0.4-16.2 years; mean, 5.1 years). Development of an adverse outcome was not associated with male sex (HR, 1.02; 95% CI, 0.36-2.88; P = .97) or with older age at onset (HR per 10-year increase in age, 0.89; 0.58-1.36; P = .58). Adverse clinical outcomes did not seem to be associated with histopathologic findings of follicular mucinosis (HR, 0.55; 95% CI, 0.18-1.68; P = .28), neutrophilic inflammation (0.29; 0.07-1.29; P = .10), eosinophilic inflammation (0.39; 0.14-1.11; P = .08), or granulomatous inflammation (0.17; 0.02-1.30; P = .09). No association was noted between adverse clinical outcome and age at onset, sex, tumor thickness, or tumor stage.

Abbreviations: FMF, folliculotropic mycosis fungoides; NA, data were unavailable or were unavailable in a format amenable to direct comparison; NC, not calculated because fewer than 10 patients still at risk; PUVA, psoralen–UV-A.

<table>
<thead>
<tr>
<th>Variable</th>
<th>Present Study (n=50)</th>
<th>van Doorn et al.1, 2002 (n=51)</th>
<th>Gerami et al.2, 2008 (n=43)</th>
<th>Gerami and Guiltart3, 2007 (n=34)</th>
<th>Cerroni et al.6, 2002 (n=28)</th>
<th>Hodak et al.7, 1999 (n=9)</th>
<th>Vergier et al.4, 1996 (n=9)</th>
<th>Flaig et al.5, 2001 (n=9)</th>
<th>Pereyo et al.8, 1997 (n=3)</th>
<th>Combined Data, %</th>
</tr>
</thead>
<tbody>
<tr>
<td>TREATMENT AND FOLLOW-UP</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Duration from diagnosis of mycosis fungoides to death</td>
<td>3.3 years (range, 0.4-16.2 years; mean, 5.1 years)</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Overall survival rate at 1, 2, and 5 years after diagnosis</td>
<td>96%, 90%, and 62%, respectively</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Cause-specific survival rate</td>
<td>Not calculated</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Adverse clinical outcomes</td>
<td>15 (30%)</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Table 2. Treatment and Prognostic Data from the Present Study and From Previous Retrospective Studies and Case Series

The study by Cerroni et al.6 also were reported in the study by Flaig et al.8 is unknown.
comes and the presence of Sézary cells on peripheral smear (2.17; 0.63-7.52; P = .22).

**SYSTEMATIC REVIEW**

We identified 8 previously published studies that each reported the clinical features of FMF in 3 or more patients; FMF has been described in 236 patients overall, including our study. Authors of 2 of the histopathologic studies also published clinical analyses that were included in our systematic review; therefore, the same patient data may have been referenced twice in the systematic review analysis. Patient characteristics and diagnostic data are summarized in eTable 1, clinical data in Table 1, histopathologic data in eTable 2, and therapeutic and prognostic data in Table 2.

**COMMENT**

Also termed pilottropic or follicular mycosis fungoides, FMF is a mycosis fungoides variant characterized by hair follicle epithelium infiltration by neoplastic lymphoid cells. It is more common in men than in women. Compared with that in men, the mean age at disease onset in women was significantly older, coinciding with postmenopausal years. Estrogen could serve as a protective factor (or testosterone as a predisposing factor) in patients otherwise prone to FMF.

Mycosis fungoides has numerous clinical manifestations, and the folliculotropic variant is no exception. Although head and neck involvement is frequent in FMF, these areas generally are spared in classic mycosis fungoides. The clinical index of suspicion for mycosis fungoides may be reduced, leading to a delay in diagnosis.

Several patients with FMF in our series had comorbid hematologic malignant neoplasms. Although composite cutaneous lymphomas have been documented in patients with mycosis fungoides and B-cell chronic lymphocytic leukemia, we observed no neoplastic B cells in cutaneous biopsy specimens from our cohort.

The implications of having circulating Sézary cells without erythroderma (observed among 9 patients in our study) are undetermined. One group uses the term mycosis fungoides with leukemic involvement to describe mycosis fungoides with circulating Sézary cells in patients not otherwise meeting diagnostic criteria for Sézary syndrome. In our study, 1 patient had FMF that progressed to Sézary syndrome.

Just as the clinical features of FMF are varied, so are the histopathologic findings (Table 2). Distinguishing between the clinical entity of follicular mucinosis (alopecia mucinosa) and FMF can be challenging histologically. Some authors have proposed that mycosis fungoides, follicular mucinosis, and FMF represent conditions along a continuous spectrum. Accurate diagnosis is important because follicular mucinosis unassociated with mycosis fungoides, especially in young patients, tends to have a persistent but benign disease course, in contrast to FMF.

In our opinion, development of intrafollicular mucin deposits is a nonspecific secondary phenomenon that may occur when neoplastic or nonneoplastic lymphoid cells infiltrate the follicular epithelium. Clinical variables and close observation over time may be required to establish the diagnosis with certainty. Presence of a prominent eosinophilic infiltrate identified in our patients and in others was reasonably specific for associated pruritus.

Although clonal T-cell populations were detected in the biopsy specimens of most patients in our study and in others, detection of a T-cell clone is not specific for cutaneous malignant neoplasms. However, in the appropriate clinical context, the presence of a clone with TCR gene rearrangement may serve as an independent predictor of disease progression. Blood T-cell clones also are nonspecific and should not be interpreted to mean that the patient has systemic involvement. Identical T-cell clones were detected in the blood and skin of 2 patients in our study, which implies systemic disease involvement and may be a marker of poor prognosis. Data in our study were insufficient to evaluate this possibility.

Although limited by their retrospective nature and the inability to standardize or directly compare treatments, other studies and ours found that FMF is less responsive to treatment than classic mycosis fungoides. Some investigators propose that the deep extension of lymphocytes into the hair follicles limits the response to superficial therapies such as UV-B phototherapy and topical corticosteroids. Other investigators have hypothesized that the lymphocyte biologic characteristics that determine pilottropism also contribute to therapy resistance.

We particularly aimed to calculate 5-year overall survival rates given apparent discrepancies among survival data of previous large retrospective studies. The present study and the study by van Doorn et al determined 5-year overall survival by measuring the time from diagnosis of mycosis fungoides to death, and these rates were similar (62% and 64%, respectively). Gerami et al used the duration between the time of lesion onset to death. After accounting for a mean time to diagnosis of 4.5 years, the survival rate reported by Gerami et al is similar (67%). Cause-specific mortality rates and time to disease progression could not be calculated because of few cause-specific deaths and our inability to gather accurate data on the timing of disease progression.

Our data may have overestimated the severity of FMF because of the tertiary referral nature of our practice and...
the likely possibility that early cases of FMF may not have been recognized. Given the potentially subtle clinical and histopathologic findings in FMF, patients with early stages of the disease may have been unrecognized.

The pathogenesis of FMF is not well understood. Heightened expression of intercellular adhesion molecule 1 in perifollicular keratinocytes in FMF, which may be induced by the neoplastic T lymphocytes, may lead to obstruction of the follicle orifice by neoplastic T cells.4,5,23,24

Skin infection is known to exacerbate mycosis fungoides.25 In the presence of Staphylococcus aureus superantigens, nonmalignant T lymphocytes stimulate the proliferation of malignant T lymphocytes by molecular cross-talk.26 Indeed, several patients in our series with evidence of impetiginization had temporary improvement of their FMF after a course of systemic antibiotics. The finding that bacterial toxins worsen mycosis fungoides most likely has implications for FMF, as the follicular ostia are reservoirs of bacteria and pruritus (a stimulus for scratching) and subsequent impetiginization is a common clinical feature of FMF.

Accepted for Publication: November 18, 2009.

Correspondence: Roger H. Weenig, MD, Associated Skin Care Specialists, PA, 7205 University Ave NE, Fridley, MN 55432 (rogerweenig@gmail.com).

Author Contributions: Dr Weenig had full access to all the data in the study and takes responsibility for the integrity of the data and the accuracy of the data analysis.

Study concept and design: Lehman, Cook-Norris, Weenig, Gibson, Weaver, and Pittelkow. Acquisition of data: Lehman, Cook-Norris, Weed, Weenig, and Pittelkow. Analysis and interpretation of data: Lehman, Weenig, Gibson, Weaver, and Pittelkow. Drafting of the manuscript: Lehman, Cook-Norris, Weed, Weenig, and Pittelkow. Critical revision of the manuscript for important intellectual content: Lehman, Cook-Norris, Weenig, Gibson, Weaver, and Pittelkow. Statistical analysis: Weaver. Administrative, technical, or material support: Weenig, Gibson, and Pittelkow. Study supervision: Lehman, Weed, Weenig, Gibson, and Pittelkow.

Financial Disclosure: None reported.