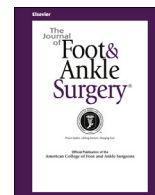




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Case Reports and Series

Squamous Cell Carcinoma From Marjolin's Ulcer of the Foot in a Diabetic Patient: Case Study

Raymond Cavaliere, DPM¹, Danielle M. Mercado, DPM², Malary Mani, MD³¹Podiatric Surgeon, Department of Podiatric Surgery and Medicine, Mount Sinai Beth Israel Medical Center, New York, NY²Podiatric Surgery Resident, Department of Podiatric Surgery and Medicine, Mount Sinai Beth Israel Medical Center, New York, NY³Pathology Fellow, Department of Pathology, Mount Sinai Beth Israel Medical Center, New York, NY

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ABSTRACT

Squamous cell carcinoma (SCC) has been commonly reported by foot and ankle specialists. Marjolin's ulcer is a malignancy that involves a posttraumatic scar or ulceration that can develop into SCC from chronic neuropathic pedal wounds, venous stasis, or decubitus ulcerations. Most Marjolin's ulcers are found in the lower extremity, specifically the feet, and it is twice as common in females as males. Biopsy of the tumor is the reference standard to diagnose SCC, and wide excision of SCC is the most common treatment option. The present case study describes an 83-year-old diabetic wheelchair-bound female who presented to the wound care clinic with a right heel nonhealing pressure ulceration. After biopsy and surgical excision, the patient was found to have SCC. This case was followed up for 5 years in which the patient had successful excision of the tumor with no recurrence. The clinical significance of our case study is to assist in the diagnosis, management, and prognosis of patients with SCC. In addition, this study has shown that adequate excision of the tumor margins and depth is necessary to prevent potential recurrence and metastasis.

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Squamous cell carcinoma (SCC) is the second most common form of cancer on the skin of the feet (1). The incidence of SCC presenting in the lower extremities has been ~1.5%; however, the risk of metastases and recurrence becomes high if not treated (2). SCC is defined as a malignant skin tumor of keratinocytes located in the epidermis, which can begin as small scaly bumps or plaques appearing inflamed and progress to hard, projecting callus-like lesions. The presentation of SCC in the feet can be caused by, but not limited to, the presence of a chronic ulceration, lichen planus, deep mycosis, and lichen simplex chronicus (3). The differential diagnosis for SCC includes keratoacanthoma, basal cell carcinoma, eccrine poroma, sweat gland carcinoma, amelanotic melanoma, pyogenic granuloma, reactive epidermal hyperplasia, an overlying site of infection, cutaneous Hodgkin disease, and mechanical trauma (4).

The term Marjolin's ulcer describes malignant degeneration in any chronic wound. Seventy-one percent of Marjolin's ulcers will develop into SCC, although basal cell carcinoma, melanoma, fibrosarcoma, li-

posarcoma, leiomyosarcoma, osteosarcoma, malignant schwannoma, and mesenchymal tumor have all been identified (5). Marjolin's ulcer is a malignancy that involves a posttraumatic scar or ulceration and can develop into SCC from chronic neuropathic wounds, venous stasis, sinus tracts, osteomyelitis, decubitus ulcerations, warts, burns, or other forms of traumatic injury to the skin (6). Approximately 60% of Marjolin's ulcers are found in the lower extremity, and it is twice as common in females as males (5). Malignant transformation of chronic wounds, regardless of the etiology, can result from the induction of neoplastic cells, radiation, or toxin-induced alterations of epidermal cells into the dermis (7). Currently, it is believed that Marjolin's ulcers develop by a slow healing process and chronic instability of scar tissue (8). When SCC occurs in a Marjolin's ulcer, it will be an aggressive malignancy; however, Marjolin's ulceration with SCC constitutes ~2% of all SCC cases (8).

The classic development of malignancy within Marjolin's ulcers includes nodule formation, induration, and ulceration at the site, which is indicative of the diagnosis. Additional clinical presentations include chronic ulceration present for >3 months, everted wound margins, excessive granulation tissue, purulence, an increase in size, bleeding on contact, crusting over, and pain (9). The reference standard for a definitive diagnosis of SCC is confirmation by biopsy, especially if it presents in chronic, nonhealing ulcerations (10). In the present report, we describe a case of Marjolin's ulceration that was treated and

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Address correspondence to: Danielle M. Mercado, DPM, Department of Podiatric Surgery and Medicine, Mount Sinai Beth Israel Medical Center, 16th Street at 1st Avenue, New York, NY 10003.

E-mail address: Dmercado1027@gmail.com (D.M. Mercado).

followed up for 5 years. In presenting this case, we sought to share the clinical significance of the entity and a strategy for diagnosis, management, and treatment of SCC developing from a Marjolin's ulcer of the foot.

Case Report

An 83-year-old female presented to the general surgery clinic in January 2011 with an ulceration on her right heel. Her medical history consisted of type 2 diabetes mellitus, cerebral vascular accident, residual left hemiparesis, polycythemia, benign hypertension, breast cancer in remission, lumpectomy, and oral chemotherapy for 5 years. Her surgical history consisted of vaginal hysterectomy, tonsillectomy and adenoidectomy, and bilateral cataract extraction with lens implants. The patient admitted to being a former smoker and denied a history of alcohol or substance use. The mechanism of injury was suspected to be neuropathic ulceration, because the right extremity was the primary weightbearing limb in this wheelchair-bound patient.

On initial examination, the wound measured 2.1 cm long, 2.0 cm wide, and 0.1 cm deep. It had a pink base, minimal slough, and was through to the subcutaneous tissue without exposed bone or muscle. No drainage, malodor, or signs of infection were present. The periphery of the wound was healthy. Conservative care included mupirocin (Bactroban) and Adaptic (Acelity, San Antonio, TX) and dry sterile dressing changes daily. The wound exhibited a gradual increase in size and depth until the spring of 2014. The patient refused to consent to a biopsy until May 21, 2014, when a central hypergranular tumor was visible. The soft tissue biopsy was obtained under sterile conditions using a 2-mm punch biopsy. Examination confirmed SCC in situ, with well-differentiated keratinizing features, grade 1, and a neutrophilic-inflamed hemorrhagic crust but could not rule out invasion (Fig. 1). The patient was then referred to the podiatry wound care clinic.

In August 2014, plain films and magnetic resonance imaging (MRI) studies were taken. The plain films of the heel noted osteopenia and heel ulceration, with no evidence of osteomyelitis (Fig. 2). MRI noted a dermal mass of the heel posteriorly corresponding to known SCC, with no evidence of osteomyelitis (Figs. 3 and 4).

In September 2014, the patient was medically cleared for wide excision of the SCC in the right heel. Intraoperatively, the ulcer measured 3.0 cm long, 3.0 cm wide, and 1.5 cm deep (Fig. 5). The SCC had not involved the Achilles tendon. From the surgeon's review of the current data, a recommended 1.0-cm margin was marked about the tumor (Fig. 6). A no. 15 scalpel was used to resect the tumor widely down to the periosteal tissue. The tumor was removed easily and sent for pathologic examination. The fatty tissue beneath the tumor was normal, with healthy borders (Fig. 7). The calcaneus was not exposed and because of the negative results for osteomyelitis on the radiographs and MRI studies, a bone biopsy was not considered. The surgical wound was cauterized using electrosurgery (Fig. 8). The pathologic results confirmed SCC with irregular epidermal hyperplasia, ulceration, and full-thickness atypia of keratinocytes with partial loss of polarity of maturation. The postoperative results were discussed with the pathologist, who confirmed that the SCC was invasive, well-differentiated, grade 1, and ulcerated and that it had been completely excised, with negative surgical margins (Fig. 9).

Wound vacuum-assisted closure (VAC) was applied to the right heel ulceration on postoperative day 1. The patient was referred to an oncologist who decided radiation was not indicated owing to the negative margins. The wound VAC was continued until postoperative day 45 when the patient returned to the operating room for debridement of the ulceration and application of Apligraf (Organogenesis, Canton, MA; Fig. 10). The wound VAC was reapplied postoperatively until the surgical site had healed. At the final follow-up examination in September

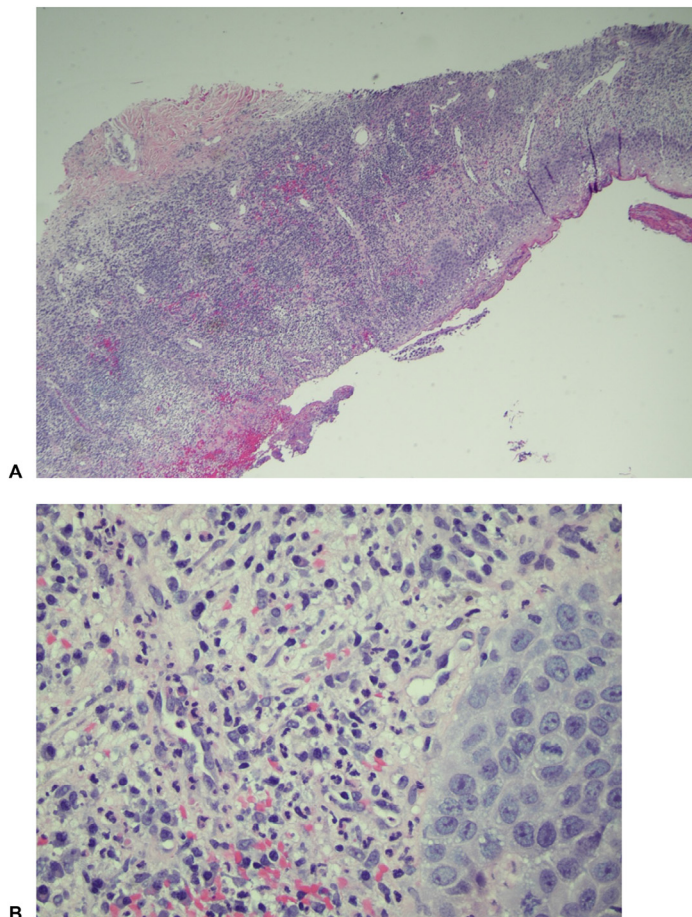


Fig. 1. (A) Skin biopsy showing epidermal ulceration (original magnification $\times 40$; hematoxylin and eosin stain) composed of (B) acute and chronic inflammatory cells and organizing granulation tissue, with reactive degenerative changes (original magnification $\times 400$; hematoxylin and eosin stain).

2015, complete wound closure had been achieved without complications (Fig. 11). In July 2016, 22 months following the wide excision of the SCC, the patient died of vascular complications; however, the right heel had no evidence of recurrence or reulceration.

Discussion

SCC found in chronic wounds typically arises from poorly managed acute traumatic wounds, in addition to wounds that form because of vascular insufficiency, diabetic neuropathy, pressure, or hemoglobinopathy (7). SCC forming from a Marjolin's ulceration that initially developed from a pressure wound has typically been found to be more aggressive than other carcinomas developing in association with chronic wounds (6). Marjolin's ulcers have the ability to develop at any anatomic location; however, their incidence is greatest in the lower extremities, and 71% of Marjolin's ulcerations can progress to SCC (8).

Marjolin's ulcerations are associated with 2 common physical presentations. One presentation is typically a shallow, well-defined ulceration with a periphery consisting of nodular elevations, which is indicative of SCC located at the margins. The second presentation is more aggressive in growth, representing an exophytic tumor consisting of papillary granulations (6). SCC forming from a Marjolin's ulceration that is initially from a pressure wound has typically been found to be more aggressive than other carcinomas associated with

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