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CASE REPORT

# Diabetic ulcer with cutaneous squamous cell carcinoma: A case report

Yun Luo, Chen-Ying Li, Yu-Qing Wang, Sheng-Min Xiang, Cheng Zhao

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### Abstract

### BACKGROUND

Chronic skin ulcers are a risk factor for the development of skin tumors. In patients with diabetes, chronic refractory ulcers may also contribute to higher susceptibility to skin tumors. Timely surgical removal of chronic and nonhealing diabetic foot ulcers can effectively prevent progression to squamous cell carcinoma. Such cases may be misdiagnosed owing to currently insufficient clinical evidence. However, in cases of chronic ulcer wounds, it is crucial to enhance clinical awareness regarding their potential progression into malignant lesions.

An 84-year-old male patient with diabetes presented with a significantly ulcerated area on his foot. The ulcer had been present to varying degrees since 1996. Between 2012 and July 2019, even after receiving treatments such as herbal medicines or heat clearance and detoxification complete healing of the wound was not achieved. In July 2020, histopathological analysis confirmed a well-differentiated cutaneous squamous cell carcinoma. After the treatments, the ulcer wound healed slowly and did not expand.

### **CONCLUSION**

Potentially malignant lesions in chronic ulcer wounds should be identified and treated in a timely manner to prevent their progression.

Key Words: Diabetic ulcer; Chronic skin ulcers; Skin cancer; Squamous cell carcinoma;

Case report

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Core Tip: Chronic skin ulcers are considered a risk factor for developing skin tumors. In people with diabetes, chronic refractory ulcers may also increase susceptibility to skin tumors. From a clinical perspective, raising awareness of their potential progression to malignant lesions is essential.

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### INTRODUCTION

Chronic skin ulcers are considered a risk factor for developing skin tumors. In patients with diabetes, chronic refractory ulcers may contribute to increased susceptibility to skin tumors[1]. Patients with diabetes are at risk of developing ulcers, with the main contributing factors being loss of protective sensation, peripheral artery disease, foot deformities, and a history of lower-limb amputation[2,3]. Poor blood sugar control and circulatory disorders further increase the risk of ulcer development in patients with diabetes, leading to high recurrence rates or chronic nonhealing of skin ulcers[4]. Repeated diabetic ulcers may transform into malignant lesions such as cutaneous squamous cell carcinoma (cSCC)[5,6]. Therefore, routine skin tissue biopsy can help avoid misdiagnosing local skin wounds that exhibit infiltrative or exophytic growth of granulation tissue. Timely surgical debridement and treatment of recurrent diabetic ulcers may prevent their development and progression to cSCC[6].

### CASE PRESENTATION

### Chief complaints

An 84-year-old male patient (born in October 1939) with diabetes presented with a significantly ulcerated area on his foot.

### History of present illness

The ulcer had been present to varying degrees since 1996.

### History of past illness

The patient was diagnosed with diabetes in 1990. Between 1996–1998, he experienced the breakdown of the ulceration on the posterior aspect of his right foot. Anti-infection treatment and traditional Chinese medicine for heat were then administered. The main herbal medicine used was Chen Lanhua Granule (main ingredients: Yinchen, Zehlan, and Chonglou), taken orally. The external medicine used was Jiangjun Powder (main ingredients: Dahuang, Mangxiao, and Gancao). After the dampness was cleared, the wound healed after a local operation to purify and clear the muscles, followed by wound-dressing. However, in 2012, the original wound ruptured on the dorsal aspect of the patient's right foot. Between 2012-2018, despite improvements in the recurrent ulcers on his foot with treatment interventions, complete healing was not achieved. The wound exhibited noticeable granulation, edema, and prominence. By August 2018, there was gradual enlargement of the ulcerated area on his foot with an irregular shape resembling a rotten vegetable pattern accompanied by soybean curd secretions-as well as increased blood and liquid oozing, and a strong foul odor (Figure 1). Pathology results obtained on December 30, 2018 indicated partial hyperplasia and keratosis of the squamous epithelium at the site (right dorsum of foot), resulting in a pathological diagnosis of squamous epithelial hyperplasia. Further pathological examination conducted on February 21st, 2019 revealed pseudo-epitheliomatous hyperplasia of the squamous epithelium (right foot) with partial atypical hyperplasia (Figure 2). Between 2012 and July 2019, several corrective surgeries were performed for the chronic ulcers present on his right foot at our hospital. However, even after receiving treatments such as the herbal medicines Chen Lanhua Granule, Dahuang, Mangxiao, Gancao, and others for heat clearance and detoxification—which did result in sufficient symptom improvement to discharge the patient from hospital care – complete healing of the wound was not achieved.

### Personal and family history

The patient had a history of diabetes and denied other family history.



Figure 1 The foot ulcer area in 2018, showing an irregular shape resembling a rotten vegetable pattern, accompanied by soybean curd secretions as well as significant blood and liquid oozing.

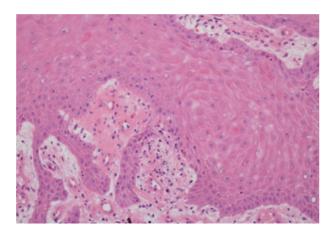


Figure 2 Pathological examination conducted on February 21st, 2019, revealed pseudo-epitheliomatous hyperplasia of the squamous epithelium (right foot) with partial atypical hyperplasia.

### Physical examination

The ulcer had an irregular shape resembling a rotten vegetable pattern accompanied by soybean curd secretions – as well as increased blood and liquid oozing, and a strong foul odor.

### Laboratory examinations

In July 2020, histopathological analysis confirmed a well-differentiated cSCC within a tissue sample from the wound (Figure 3).

### Imaging examinations

On February 13, 2020, radiographic examination revealed misalignment of the fifth metatarsophalangeal joint in the right foot, accompanied by osteoporosis and soft tissue edema in the same region.

### **FINAL DIAGNOSIS**

The patient was diagnosed with well-differentiated cSCC of the foot.

### **TREATMENT**

Local tissue excision surgery was performed, followed by appropriate cleaning. Nano-silver excipients were used to promote ulcer wound healing, in conjunction with Chinese herbal decoctions to clear heat, detoxify, and remove saprophytic muscle. Simiao and Tuoli Xiaodu powders were used as the main prescriptions for addition and subtraction treatments.



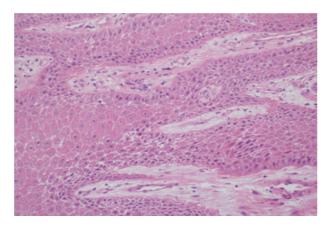


Figure 3 In July 2020, histopathological analysis confirmed well-differentiated cutaneous squamous cell carcinoma once again within the wound tissue sample.

### OUTCOME AND FOLLOW-UP

After the treatments were administered, the ulcer wound healed slowly and did not expand.

### **DISCUSSION**

Recurrent ulcers can lead to cSCC. Previous case reports have highlighted instances where toe SCCs in patients with diabetes were initially misdiagnosed as diabetic foot ulcer(DFU)[5-7]. Neglected chronic wounds and long-term scar tissue complications of cSCC have also been documented [8,9]. Additionally, there are reports in the literature of diabetic lipoid necrosis (NLD) in the legs that progressed to ulcerations and eventually degenerated into cSCC[10]. The NLD environment provides a favorable setting for the development of cSCC, emphasizing the importance of considering nonhealing ulcers as potential indicators. Early biopsy is recommended for such lesions[11]. A growing number of studies have reported clinical cases in which malignant tumors were initially misdiagnosed as DFU. A review of these cases revealed subsequent diagnoses, including melanoma (68%), Kaposi's sarcoma (14%), cSCC (11%), mantle cell lymphoma, and diffuse B-cell lymphoma (4%). Notably, older age (≥ 65 years) was associated with an increased risk of malignancy masked by DFU. The suspected risk in older patients was 145% higher than that in younger ones (< 65 years). Clinicians should remain vigilant for potentially malignant foot lesions in older patients with diabetes aged ≥ 65 years[12].

Increasing evidence has demonstrated a correlation between abnormal glucose homeostasis and the development and prognosis of certain tumors[13]. Elevated blood sugar levels can independently, directly, or indirectly increase the risk of cancer[14]. In addition, diabetes-related oxidative stress damage is believed to contribute to oncogenic transformation and cancer progression[15]. The aforementioned cases are not known to have been misdiagnosed or induced because of insufficient current clinical evidence. However, in cases of chronic ulcer wounds resembling those mentioned above, it is crucial to enhance clinical awareness of their potential progression to malignant lesions. Needle biopsies represent a simple and effective diagnostic technique for treating wounds that are resistant to healing. Histological examination can rule out cancer in the wound bed and diagnose difficult-to-cure or suspicious ulcers[16]. Studies have found that the incidence of nonvascular ulcers is 5%, to the majority of which are skin cancers. Marginal eminence and abnormal growth of granulation tissue represent the most common features of these ulcers. Therefore, a timely skin biopsy is warranted [17]. Regardless of the duration of chronic ulcer recurrence, routine skin tissue biopsies can prevent misdiagnosis. Timely surgical removal of chronic nonhealing DFU may impede the development of cSCC. Physicians should remain vigilant regarding the possibility of malignancy in patients with diabetes who present with persistent and refractory ulcerative foot lesions. Furthermore, it is anticipated that future research will uncover additional clinical evidence elucidating this "misdiagnosis", thereby preventing further instances while offering more effective and beneficial treatment options for these patients.

### CONCLUSION

In case of chronic ulcer wounds, attention should be paid to potential malignant lesions and timely treatment should prevent their malignant progression.

### **FOOTNOTES**

Author contributions: Luo Y provided medical records and asked for pathology reports; Li CY wrote the manuscript; Xiang SM and Wang YQ searched the literature; Zhao C conceived the manuscript.

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