

## What Is Your Diagnosis?



This patient reported recent rapid growth in a long-standing lesion.

PLEASE TURN TO PAGE 173 FOR DISCUSSION

Dirk M. Elston, MD, Departments of Dermatology and Laboratory Medicine, Geisinger Medical Center, Danville, Pennsylvania.

LTC Mary Farley, MC, USA; Maj John Albertini, USAF, MC; Skin Surgery Center, Winston-Salem, North Carolina. The authors report no conflict of interest.

The views expressed are those of the authors and are not to be construed as official or as representing those of the US Army Medical Department or the US Department of Defense. The authors were full-time federal employees at the time this work was completed. It is in the public domain.

## The Diagnosis: Periungual Squamous Cell Carcinoma



Clinically, a fungating lesion on a digit is likely to represent squamous cell carcinoma (SCC). Biopsy results of the patient's lesion demonstrated a deeply invasive SCC. Osseous involvement was present. Nodal metastases were confirmed by biopsy. The patient was treated with digital amputation, lymphadenectomy, and radiation therapy.

The diagnosis of digital SCC often is delayed. A high index of suspicion is required, as the diagnosis is complicated by its resemblance to verruca vulgaris or a variety of benign inflammatory conditions.<sup>1</sup> The lesions are commonly hyperkeratotic. Underlying induration may not be prominent. Oftentimes, the patient has had periungual warts for many years. Carpal tunnel syndrome can be an unusual presentation resulting from perineural extension of cutaneous SCC.<sup>2</sup> Because periungual lesions are distally located, nerve involvement in these lesions is less likely to produce symptoms. Skin biopsy, with adequate depth to demonstrate the deeper aspects of the lesion, is critical to establish the diagnosis. Digital SCC can be well, moderately, or poorly differentiated. Well-differentiated verrucous carcinoma accounts for a

large proportion of digital SCCs. The epithelium can appear fairly bland, and the superficial portions of the lesion can maintain a warty appearance. An adequate biopsy specimen is particularly critical when evaluating verrucous carcinoma.

Periungual Bowen disease can appear as periungual erythema with scaling and erosions, a hyperkeratotic or papillomatous plaque, fissure, or crusted ulcer. Soreness may be noted on palpation of the lesion.<sup>3</sup> Bowen disease should be considered in the differential diagnosis of any persistent scaly or crusted lesion of the finger.<sup>4</sup>

Increasing evidence supports the association of periungual SCC with human papilloma virus (HPV) infection. Periungual or nail bed SCC often retains its wartlike clinical appearance, and lesions are commonly misdiagnosed as refractory warts.<sup>5</sup> HPV type 16 DNA is commonly detected in such lesions by means of the polymerase chain reaction technique and Southern blotting.<sup>6-8</sup> HPV type 35 RNA has been detected in periungual SCC and in axillary nodal metastases.<sup>9</sup> The strongest association between skin cancer and HPV is noted in digital

and genital SCC. HPV seems to play only a minor role in the development of SCC at other cutaneous sites.<sup>10</sup> Refractory warts have been noted to evolve into SCC at multiple digital and anogenital sites in a single patient.<sup>11</sup> Multiple digits in a single patient may demonstrate lesions of Bowen disease and invasive carcinoma.<sup>12</sup>

In addition to HPV, exposure to radiation appears to predispose patients to digital SCC.<sup>13</sup> In many cases, exposure to radiation relates to one's occupation. At one time, some dentists held radiographic film in place with a digit while performing dental x-rays. Digital SCC also has occurred in the setting of radiation dermatitis from radioactive gold rings.<sup>14,15</sup> Radiation from a gold ring can amount to hundreds of cGy per week and several thousand Gy of radiation exposure over many years. Digital SCC also has been seen in the setting of hereditary ectodermal dysplasia.<sup>16</sup>

Surgical excision remains the treatment of choice for most digital SCCs. Mohs micrographic surgery has been successfully used to eradicate tumors while sparing tissue on the affected digit.<sup>17</sup> Mohs micrographic surgery is appropriate for periungual SCCs that lack osseous involvement.<sup>18</sup> A combined approach using surgical excision and 5-fluorouracil has been used to treat Bowen disease involving multiple nail beds.<sup>19</sup> Digital SCC with nodal involvement may require amputation of the digit, regional lymph node dissection, postoperative radiation therapy, and systemic chemotherapy.<sup>20</sup>

## REFERENCES

1. Lumpkin LR, Rosen T, Tschien JA. Subungual squamous cell carcinoma. *J Am Acad Dermatol*. 1984;11:735-738.
2. Mackay IR, Barua JM. Perineural tumour spread: an unusual cause of carpal tunnel syndrome. *J Hand Surg [Br]*. 1990;15:104-105.
3. Baran R, Dupre A, Sayag J, et al. Bowen disease of the nail apparatus. report of 5 cases and review of the 20 cases of the literature [in French]. *Ann Dermatol Venerol*. 1979;106:227-233.
4. Kern AB, Schiff BL. Bowen's disease of the finger. *Cutis*. 1977;19:229-232.
5. Moy RL, Eliezri YD, Nuovo GJ, et al. Human papillomavirus type 16 DNA in periungual squamous cell carcinomas. *JAMA*. 1989;261:2669-2673.
6. Ashinoff R, Li JJ, Jacobson M, et al. Detection of human papillomavirus DNA in squamous cell carcinoma of the nail bed and finger determined by polymerase chain reaction. *Arch Dermatol*. 1991;127:1813-1818.
7. Eliezri YD, Silverstein SJ, Nuovo GJ. Occurrence of human papillomavirus type 16 DNA in cutaneous squamous and basal cell neoplasms. *J Am Acad Dermatol*. 1990;23:836-842.
8. Sanchez-Lanier M, Triplett C, Campion M. Possible role for human papillomavirus 16 in squamous cell carcinoma of the finger. *J Med Virol*. 1994;44:369-378.
9. McHugh RW, Hazen P, Eliezri YD, et al. Metastatic periungual squamous cell carcinoma: detection of human papilloma virus type 35 RNA in the digital tumor and axillary lymph node metastases. *J Am Acad Dermatol*. 1996;34:1080-1082.
10. Kawashima M, Favre M, Obalek S, et al. Premalignant lesions and cancers of the skin in the general population: evaluation of the role of human papillomaviruses. *J Invest Dermatol*. 1990;95:537-542.
11. Shelley WB, Wood MG. Transformation of the common wart into squamous cell carcinoma in a patient with primary lymphedema. *Cancer*. 1981;48:820-824.
12. Goodman G, Mason G, O'Brien T. Polydactylous Bowen's disease of the nail bed. *Australas J Dermatol*. 1995;36:164-165.
13. Koopelson PL, Nguyen QH, Moy RL. Verruca vulgaris and radiation exposure are associated with squamous cell carcinoma of the finger. *J Dermatol Surg Oncol*. 1994;20:38-41.
14. Stutzman CD, Schmidt GD. Squamous cell carcinoma of the skin associated with radioactive gold rings. *J Am Acad Dermatol*. 1984;10:1075-1077.
15. Miller RA, Aldrich JE. Radioactive gold ring dermatitis. *J Am Acad Dermatol*. 1990;23:360-362.
16. Mauro JA, Maslyn R, Stein AA. Squamous-cell carcinoma of nail bed in hereditary ectodermal dysplasia. *N Y State J Med*. 1972;72:1065-1066.
17. Tomsick RS, Menn H. Squamous cell carcinoma of the fingers treated with chemosurgery. *South Med J*. 1984;77:1124-1126.
18. Goldminz D, Bennett RG. Mohs micrographic surgery of the nail unit. *J Dermatol Surg Oncol*. 1992;18:721-726.
19. Strong ML. Bowen's disease in multiple nail beds—case report. *J Hand Surg [Am]*. 1983;8:329-330.
20. Lai CS, Lin SD, Tsai CW, et al. Squamous cell carcinoma of the nail bed. *Cutis*. 1996;57:341-345.