## Multiple giant clear cell acanthomas

Sir,

Clear cell acanthoma, first described by Degos in 1962, was originally thought to be a benign epidermal tumor. However, recent studies suggest that it is a reactive inflammatory dermatosis. Immunohistochemically, it shows a staining pattern similar to inflammatory dermatoses such as psoriasis vulgaris, lichen planus and discoid lupus erythematosus. The etiology remains unknown. The lesion is typically slow-growing, asymptomatic, solitary and located on the legs, although all parts

of the body can be affected. It tends to affect people aged 50–70 years and usually consists of a sharply demarcated, dome-shaped nodule or plaque, 5–20 mm in diameter. The most characteristic clinical features are a thin wafer-like scale forming a peripheral collarette, slight moisture and numerous pin-point sized blanchable vascular punctae scattered over the surface.

An 81-year-old woman was seen in the department of dermatovenereology, Bezmialem Vakif University Hospital, Istanbul, Turkey, complaining of itchy papules and nodules on both lower extremities. She reported that these nodules first appeared on the right calf 40 years ago. Twenty years later, satellite papules started appearing around the initial, larger nodule on

her right calf. Subsequently, similar lesions were noted on the left leg and both thighs. The lesions had been itching and bleeding over the last few years. Her past medical history was significant for hypertension and hypothyroidism for which she was on levothyroxine and atenolol/chlorthalidone. None of her family members had similar lesions. On physical examination, there was a 40 mm  $\times$  30 mm, red, sharply demarcated nodule with satellite papulo-nodules on her right calf. There were scattered, well defined papules and nodules (5–15 mm) with a thin rim of scale on both posterior lower extremities. Three biopsies were performed from lesions from both legs [Figure 1].

On histological examination, there was epidermal acanthosis with the proliferation of clear cells containing intracytoplasmic glycogen. Periodic-acid—Schiff staining was positive. Clear demarcation was seen between the lesion and adjacent normal epidermis. There were neutrophils in the stratum corneum. The blood vessels in the papillary dermis were dilated and increased in number. Based on these findings, a diagnosis of multiple clear cell acanthoma was made [Figure 2].

The first case of multiple clear cell acanthoma was described in 1964 by Delacretaz *et al.*<sup>[3]</sup> Several variants of the condition including giant, polypoid, pigmented and eruptive patterns have been described. The eruptive form is characterized by more than 30 papules which arise within a month. Multiple and giant forms are rare; we were able to find 38 cases of multiple and 9 giant



Figure 1: Appearance of the posterior side of lower extremities (a). There was a 40 mm  $\times$  30 mm red, sharply demarcated nodule and 10 mm  $\times$  20 mm satellite papulonodules around it on her posterior side of right leg (c). There was scattered 5 mm, 10 mm, 15 mm papules and nodules with sharply demarcated margins and a thin collateral rim of scale on her posterior side of both lower extremities (b.d and e)

clear cell acanthoma in the literature. Two out of the nine cases of giant clear cell acanthomas had multiple lesions. Langtry et~al. reported a case with multiple giant (40 mm) lesions and a lesion 10 mm  $\times$  12 mm in size on the left buttock of an 87-year-old woman. [4] Kim et~al. reported a 14-year-old girl with a 2-month history of two erythematous nodules measuring 20 mm  $\times$  32 mm and 40 mm  $\times$  55 mm on the perianal area. [5]

We present a case of multiple clear cell acanthomas with a giant lesion and a total of 11 lesions. We found 38 previous reports of multiple clear cell acanthoma in the literature [Table 1]. [1-35] The most prominent associations were xerosis (6/39) and ichthyosis (4/39). Our patient had mild xerosis too.

The clinical differential diagnosis is broad and includes basal cell carcinoma, squamous cell carcinoma. Bowen's disease, actinic keratosis, hemangioma, dermatofibroma, inflamed seborrheic keratosis, verruca vulgaris, amelanotic melanoma, psoriasis, histiocytoma, eczematous dermatitis, nevus, parapsoriasis and pyogenic granuloma.[1-5] Our patient had multiple red nodules with a vascular appearance. Hence, our clinical differential diagnosis was Kaposi sarcoma and B-cell lymphoma. Morrison et al. reported 411 histologically confirmed cases but noted that rarely had the condition been recognised clinically.[1] They considered that this might be due to a variable clinical appearance with overlapping features of several other lesions making it difficult to distinguish, or because it was an entity that was rare and not frequently suspected.[1]

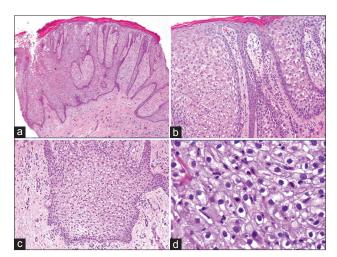


Figure 2: On histological examination epidermal acanthosis, pale appearance of the keratinocytes and intraepidermal neutrophils were seen. In the dermis, there was proliferation of blood vessels and perivascular infiltration of lymphocytes, histiocytes and neutrophils. (a) H and E,  $\times$ 40 (b) H and E,  $\times$ 100 (c) H and E,  $\times$ 100 (d) H and E.  $\times$ 200

Table 1: Features of multiple CCA\* are listed in Table 1.1-7

Case	Sex	Age	Site	Number of lesions	Associated diseases	References
1	Female	51	Legs	Multiple	-	Delacretaz et al.
2	Male	62	Legs	16	-	Duperrat et al.
3	Female	41	Thighs, legs	4	-	Delacretaz et al.
4	Male	59	Thighs, legs	7	Lamellar ichthyosis	Thorne et al.
5	Male	64	Thighs, legs	7	Lamellar ichthyosis	Landry et al.
6	Male	73	Legs	2	Ichthyosis vulgaris	Landry et al.
7	Male	68	Legs	8	Varicose veins, kserosis	Ebner et al.
8	Female	56	Legs	10	-	Kaufmann et al.
9	Female	64	Legs	10	Varicose veins	Dcsmons et al.
10	Female	45	Legs	7	-	Dosmons et al.
11	Female	14	Leg	4	-	Witkowsk et al.
12	Female	65	Lower limbs	124	-	Varotti et al.
13	Female	69	Legs, trunk, chest	12	Seborrheic keratosis, xerosis	Trau et al.
14	Male	63	Trunk, legs	6	-	Goette et al.
15	Female	58	Upper/lower limbs	30	Xerosis	Balus et al.
16	Female	37	Forehead, trunk, upper/lower limbs	16	Hodgkin's disease	Balus et al.
17	Female	64	Thighs, legs	30	-	Naeyaert et al.
18	Male	43	Lower limbs	43	-	Dupre et al.
19	Male	59	Lower limbs	4	Xerosis	Baden et al.
20	Male	57	Legs	15	Seborrheic keratosis, gynecomastia	Williams et al.
21	Male	70	Arms, lower limbs	30	-	Donati et al.
22	Male	50	Buttock, lower limbs	50	-	Donati et al.
23	Female	50	Legs	12	-	Kavanagh et al.
24	Female	32	Legs, buttocks, trunk, arms	>400	Family history: father and brother with psoriasis	Morillo et al.
25	Male	54	Legs	Multiple	Chronic venous insufficiency	Kneitz et al.
26	Male	57	Anterior thigh	Multiple	SCC**,BCC***,HT****,HC*****, condyloma acuminata	Bang et al.
27	Male	74	Lower extremities	60	-	Monari et al.
28	Female	14	Genital	2	-	Kim et al.
29	Female	65	Thighs, buttocks and lower legs	Multiple	HT****, hypercholesterolemia, and diabetes	Garcia-Gavin et al.
30	Female	52	Lateral cheek	4	-	Hatakeyama et al.
31	Female	16	Inner thighs	Multiple	-	Wilson et al.
32	Female	40	Arms, upper thigh	4	Ichthyosis bullosa (superficial epidermolytic ichthyosis) ichthyosis bullosa	Jacyk et al.
33	Female	76	Thighs	5	Multiple Bowen's disease	Shirai et al.
34	Male	52		Multiple	-	Martínez Escanamé et a
35	Male	69	Legs	20	-	Lacarrubba et al.
36	Male	59	Legs	17	Varicose veins and dry skin	Betti et al.
37	Male	52	All parts of body	>100	-	Fandrey et al.
38	Female		Legs, arms and trunk	>100	-	Burg et al.
39	Female		Legs, thighs	10	HT***, goiter, hypothyroidism	Su et al.(our case)

CCA\*: Clear cell acanthoma, SCC\*\*: Squamous cell carcinoma, BCC\*\*\*: Basal cell carcinoma, HT\*\*\*\*: Hypertension, HC\*\*\*\*\*: Hepatitis C

Treatment consists of electrofulguration, curettage, surgical excision, cryotherapy, carbon dioxide laser and topical 5-fluoruracil. The most common treatment option for multiple lesions is cryotherapy, which was also performed in our patient

## Financial support and sponsorship

Nil.

## **Conflicts of interest**

There are no conflicts of interest.

Ozlem Su, Didem Dizman, Nahide Onsun, Dilek Biyik Ozkaya, Zeynep Tosuner<sup>1</sup>, Anil Gulsel Bahali, Cuyan Demirkesen<sup>2</sup>

Departments of Dermatology and Venereology and <sup>1</sup>Pathology, Medical Faculty, Bezmialem Vakif University, <sup>2</sup>Department of Pathology, Cerrahpasa Medical Faculty, Istanbul University, Istanbul, Turkey

Address for correspondence: Dr. Ozlem Su, Department of Dermatology and Venereology Medical Faculty, Bezmialem Vakif University, Fatih, Istanbul, Turkey. E-mail: ozlemsukucuk2@yahoo.com.tr

## **REFERENCES**

- Morrison LK, Duffey M, Janik M, Shamma HN. Clear cell acanthoma: A rare clinical diagnosis prior to biopsy. Int J Dermatol 2010;49:1008-11.
- Ohnishi T, Watanabe S. Immunohistochemical characterization of keratin expression in clear cell acanthoma. Br J Dermatol 1995;133:186-93.
- Delacretaz J, Christeler A, Loi-Zedda P, Lenggenhager R. Acanthom a cellules claires. Dermatologica 1967;135:304.
- Langtry JA, Torras H, Palou J, Lecha M, Mascaro JM. Giant clear cell acanthoma in an atypical location. J Am Acad Dermatol 1989;21 (2 Pt 1):313-5.
- Kim CY, Kim NG, Oh CW. Multiple reddish weeping nodules on the genital area of a girl. Giant clear cell acanthoma (CCA). Clin Exp Dermatol 2010;35:e67-9.
- 6. Martínez Escanamé M, Arzberger E, Hofmann-Wellenhof R, Massone C. Multiple clear-cell acanthomas. Australas J Dermatol 2015;56:e46-8.
- 7. Duperrat B, Mascaro JM, Mascaro-Galy C. 7 cases of clear cell acanthoma. Arch Belg Dermatol Syphiligr 1966;22:55-9.
- 8. Thorne EG, Fusaro RM. Multiple clear cell acanthomas in a patient with lamellar ichthyosis. Dermatologica 1971;143:105-10.
- 9. Ebner H. Multiple clear-cell acanthomas. Z Hautkr 1974;49:541-6.
- Kaufmann J, Eichmann A, Meves C, Ott F, Somazzı ST, Storck H. Klarzellakanthom (multipel): Demonstrationen. Dermatologica 1976;153:116-7.
- Dosmons F, Breuillard F, Thomas P, Leonardflli J, Hildebrand HF. Clear cell acanthoma (Degos). Histochemical and ultrastructural study of two cases. J Dermatol 1977;16:203-7.
- 12. Witkowski JA, Parish LC. Clear cell acanthoma. Int J Dermatol 1979;18:162-3.
- Varotti C, Passarini B, Baratta L. Acantoma a cellule chiare multiplo. Studio istoiogico ed ultrastrutturale di un caso. G Ital Dermatol Venereol 1979;144:765-70.
- 14. Trau H, Fisher BK, Schewach-Millet M. Multiple clear cell acanthomas. Arch Dermatol 1980;116:433-4.
- Goette DK, Diakon NC. Multiple clear cell acanthomas. Arch Dermatol 1983;119:359-61.
- Balus L, Cainelli T, Cristiani R, Donati P. Multiple familial clear cell acanthoma. Ann Dermatol Venereol 1984;111:665-6.
- Naeyaert JM, de Bersaques J, Geerts ML, Kint A. Multiple clear cell acanthomas. A clinical, histological, and ultrastructural report. Arch Dermatol 1987;123:1670-3.
- 18. Dupre A, Viraben R. Forme eruptive de tacanthome a cellules claires. Nouv Dermatol 1987;6 Suppl 4:485.
- Baden TJ, Woodley DT, Wheeler CE Jr. Multiple clear cell acanthomas. Case report and delineation of basement membrane zone antigens. J Am Acad Dermatol 1987;16 (5 Pt 2):1075-8.
- Williams RE, Lever R, Seywright M. Multiple clear cell acanthomas – treatment by cryotherapy. Clin Exp Dermatol 1989;14:300-1.
- 21. Donati P, Leonetti F, Paiamara G. Due casi di acantoma a cellule chiare multiplo. Dermatol Oggi 1991;5:29-31.
- 22. Kavanagh GM, Marshman G, Burton JL. Multiple clear cell

- a canthomas treated by cryotherapy. Australas J Dermatol 1995; 36:33-4.
- 23. Morillo V, Manrique P, Zabalza I, Artola JL. Eruptive clear cell acanthoma. Actas Dermosifiliogr 2009;100:244-6.
- Kneitz H, Bröcker EB, Ugurel S. Slowly changing nodules on both lower limbs in a 54-year old patient. J Dtsch Dermatol Ges 2009;7:63-5.
- Bang R. Multiple clear cell acanthomas in association with hepatitis C: A novel presentation. J Am Acad Dermatol (Poster Abstracts of 68<sup>th</sup> Annual Meeting, March 5–9, 2010 Miami, Florida) 2010;623:AB39:P1155.
- Monari P, Farisoglio C, Gualdi G, Botali G, Ungari M, Calzavara-Pinton P. Multiple eruptive clear cell acanthoma. J Dermatol Case Rep 2010;4:25-7.
- García-Gavín J, González-Vilas D, Montero I, Rodríguez-Pazos L, Pereiro MM, Toribio J. Disseminated eruptive clear cellacanthoma with spontaneous regression: further evidence of an inflammatory origin? Am J Dermatopathol 2011;33:599-602.
- 28. Hatakeyama M, Oka M, Fukunaga A, Kunisada M, Funasaka Y, Nishigori C. A case of multiple facial clear cell acanthomas successfully treated by cryotherapy. Indian J Dermatol 2013;58:162.
- 29. Wilson J, Walling HW, Stone MS. Benign cutaneous Degos disease in a 16-year-old girl. Pediatr Dermatol 2007;24:18-24.
- Jacyk WK, Baran W, Essop A. Multiple pigmented clear cell acanthoma in an African patient. J Eur Acad Dermatol Venereol 2014. Doi: 10.1111/jdv.12882. [Epub ahead of print].
- 31. Shirai A, Saeki H, Matsuzaki H, Ito K, Nakagawa H. Multiple clear cell acanthoma associated with multiple Bowen's disease. Int J Dermatol 2014;53:e386-8.
- Lacarrubba F, de Pasquale R, Micali G. Videodermatoscopy improves the clinical diagnostic accuracy of multiple clear cell acanthoma. Eur J Dermatol 2003;13:596-8.
- Betti R, Bruscagin C, Inselvini E, Palvarini M, Crosti C. Successful cryotherapic treatment and overview of multiple clear cell acanthomas. Dermatol Surg 1995;21:342-4.
- Fandrey K, Rabenhorst S, Fedler R, Frosch PJ. Multiple clear cell acanthomas. Hautarzt 1995;46:865-8.
- 35. Burg G, Würsch T, Fäh J, Elsner P. Eruptive hamartomatous clear-cell acanthomas. Dermatology 1994;189:437-9.

This is an open access article distributed under the terms of the Creative Commons Attribution-NonCommercial-ShareAlike 3.0 License, which allows others to remix, tweak, and build upon the work non-commercially, as long as the author is credited and the new creations are licensed under the identical terms.



How to cite this article: Su O, Dizman D, Onsun N, Ozkaya DB, Tosuner Z, Bahali AG, et al. Multiple giant clear cell acanthomas. Indian J Dermatol Venereol Leprol 2016;82:218-21.

Received: February, 2015. Accepted: July, 2015.