Comedones como manifestación de enfermedad sistémica

López Vera E Llamas Carmona JA García-Hirschfeld JM

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CASOS PARA EL DIAGNÓSTICO

Comedones como manifestación de enfermedad sistémica

[[Translated article]]Comedones as a Sign of Systemic Disease

- Autores: López Vera, E.¹; Llamas Carmona, JA¹; García-Hirschfeld, JM²
- Centro: ¹Servicio de Dermatología y ²Servicio de Anatomía Patológica del Hospital Regional Universitario de Málaga.
- Correspondencia con autor:

López Vera, Eduardo

Email: elopez.vera@outlook.es

Medical history

A 51-year-old woman with a past medical history of hypothyroidism on 100 mg of levothyroxine daily, presented with a < 1-year history of generalized pruritic lesions.

Physical examination

Physical examination revealed the presence of multiple plaques composed of follicular plugs and open comedones with an erythematous-violaceous peripheral halo, distributed on the elbows, ears, preauricular region, knees, and palms, without involvement of the soles or mucous membranes (Fig. 1a-c). Lesions were highly pruritic and refractory to treatment with sedative antihistamines, appeared after sun exposure and were accompanied by asthenia, arthralgia, and a slight increase of hair loss. Photosensitivity, however, was not reported.

Histologic examination revealed an atrophic epidermis with vacuolar degeneration and rarefaction of the basal layer (Fig. 2a), along with superficial and deep perivascular dermatitis, a mild-to-moderate chronic lymphomonocytic inflammatory infiltrate in the superficial and middermis that spared the periannexal area, and marked follicular hyperparakeratosis with dilation of the follicular ostium and infundibular destruction with fibrinoid necrosis forming early comedones (Fig. 2b), accompanied by interstitial mucin in the reticular dermis (Fig. 2c).

Additional tests: blood tested revealed low-titer positive antinuclear antibodies (ANA 1:40), thrombocytopenia, and decreased serum complement C3 levels, and positivity for extractable nuclear antigen (ENA) antibodies, anti-Smith (anti-Sm) antibodies, anti-small nuclear ribonucleoprotein (anti-RNP) antibodies, and anti-Ro antibodies. Antiphospholipid antibodies turned out negative.

Diagnosis

Comedonic lupus cutaneous, which did not meet the classification criteria for systemic lupus erythematosus since ANA titers did not reach 1:80.

Disease progression and treatment

After an early regimen of topical tretinoin 0.5 mg/g daily and 25 mg of hydroxyzine orally every 8 hours—which proved ineffective—oral prednisone and topical fluticasone were administered, along with strict photoprotection. Afterwards, oral hydroxychloroquine at a dose of 200 mg every 12 hours was introduced, with maintenance therapy using tacrolimus 0.1% ointment. Lesions improved significantly after 2 months of treatment (Fig. 3a-c), along with the resolution of thrombocytopenia, negative autoimmune parameters, and normalization of C3 levels.

Commentary

Comedonic lupus is a rare and poorly understood form of chronic cutaneous lupus erythematosus (CCLE)¹. It is primarily described in young women in their 30s and 40s, with a slightly increased risk among smokers¹,². Lesions are typically pruritic and appear in photoexposed areas, predominantly on the malar, preauricular, and chin regions. They consist of comedones or cysts with varying degrees of surrounding inflammation and may heal with scarring¹.

The presence of comedones can be associated with elevated levels of interleukin-1 (IL-1), as this has been found to be biologically and pathologically significant in early-stage comedones and is known to play a crucial role in the pathogenesis of lupus erythematosus³,⁴. Therefore, controlling inflammatory activity is essential to prevent the growth of new lesions, whereas keratolytics or sebum regulators such as topical retinoids have proven less effective⁵.

This condition can be easily mistaken for other benign conditions such as acne vulgaris, which typically exhibits more inflammatory flare-ups, pustules, and a distribution beyond photo-exposed areas. Other differential diagnoses include follicular lichen planus tumidus, which requires histopathological and analytical findings for differentiation; milia en plaque; Favre-Racouchot disease—elastosis with cysts and comedones—which features similar lesions but less inflammation and significant actinic damage; nevus comedonicus—which is present since childhood—syringomas; or trichoepitheliomas—which require histological examination for definitive diagnosis^{1,5}. Currently, although comedonic lupus is considered an underdiagnosed form of lupus, it is gaining clinical relevance^{1,2,5}.

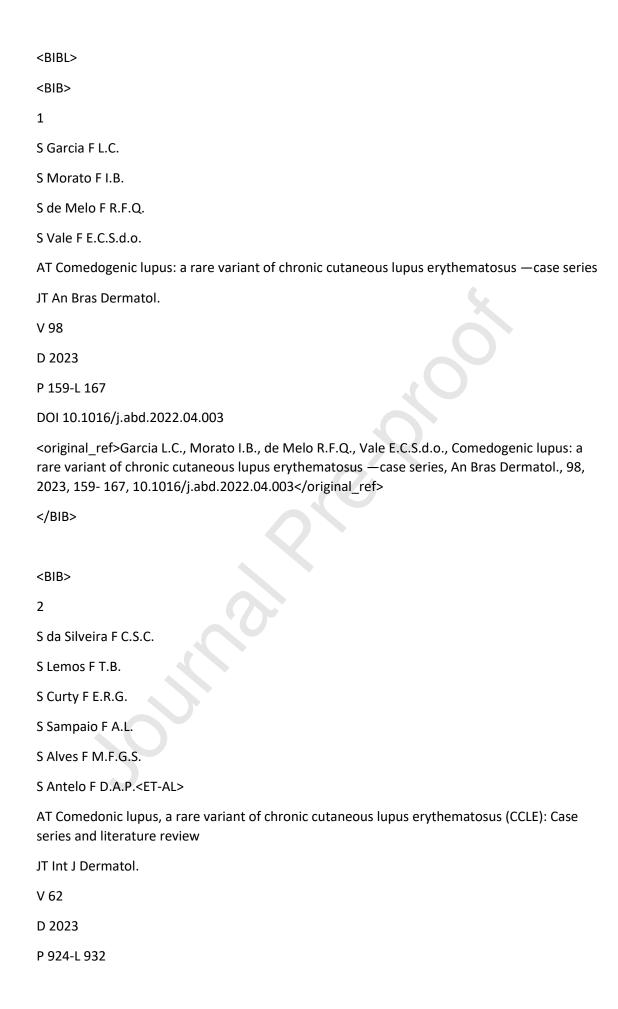
However, comedonic CCLE can be locally destructive, leading to atrophic and disfiguring scars in untreated cases⁶. Immunofluorescence assays may assist in diagnosis when histopathological findings prove inconclusive, as deposits of IgM, IgG, and C3 at the dermoepidermal junction can be observed^{1,5}.

Regarding treatment, photoprotection is essential in all cases². Oral hydroxychloroquine has demonstrated the highest efficacy, followed, to a lesser extent, by topical corticosteroids or oral tetracyclines^{1,2}. Other authors have reported satisfactory outcomes with dapsone combined with methotrexate in 2 cases¹. Topical retinoids have shown limited effectiveness⁵. Since up to one-third of cases are associated with systemic lupus erythematosus, a comprehensive evaluation and regular follow-ups are necessary to prevent future local CCLE-related complications^{1,5,6}.

Conflicts of interest

None declared.

References



DOI 10.1111/ijd.16675 <original_ref>da Silveira C.S.C., Lemos T.B., Curty E.R.G., Sampaio A.L., Alves M.F.G.S., Antelo D.A.P., Comedonic lupus, a rare variant of chronic cutaneous lupus erythematosus (CCLE): Case series and literature review, Int J Dermatol., 62, 2023, 924-932, 10.1111/ijd.16675</original_ref> </BIB> <BIB> 3 S Kahlenberg F J.M. S Kaplan F M.J. AT The inflammasome and lupus: Another innate immune mechanism contributing to disease pathogenesis? JT Curr Opin Rheumatol. V 26 D 2014 P 475-L 481 DOI 10.1097/BOR.0000000000000088 <original_ref>Kahlenberg J.M., Kaplan M.J., The inflammasome and lupus: Another innate immune mechanism contributing to disease pathogenesis?, Curr Opin Rheumatol., 26, 2014, 475-481, 10.1097/BOR.000000000000088</original ref> </BIB> <BIB> S Cunliffe F W.J. S Holland F D.B. S Jeremy F A. AT Comedone formation: Etiology, clinical presentation, and treatment JT Clin Dermatol. V 22 D 2004 P 367-L 374

DOI 10.1016/j.clindermatol.2004.03.011 <original_ref>Cunliffe W.J., Holland D.B., Jeremy A., Comedone formation: Etiology, clinical presentation, and treatment, Clin Dermatol., 22, 2004, 367-374, 10.1016/j.clindermatol.2004.03.011</original_ref> </BIB> <BIB> 5 S Chessé F C. S Fernández-Tapia F M.J. S Borzotta F F. AT Comedonic lupus: An unusual presentation of cutaneous lupus JT Actas Dermosifiliogr. V 112 D 2021 P 370-L 371 DOI 10.1016/j.ad.2019.07.013 <original_ref>,, Chessé C., Fernández-Tapia M.J., Borzotta F., Comedonic lupus: An unusual presentation of cutaneous lupus, Actas Dermosifiliogr., 112, 2021, 370-371, 10.1016/j.ad.2019.07.013</original_ref> </BIB> <BIB> 6 S Cozzani F E. S Herzum F A. S Burlando F M. S Parodi F A. AT Comedonal variant of chronic cutaneous lupus erythematosus causing mutilation of the earlobe JT JAAD Case Rep. V 6

D 2020

P 843-L 844

DOI 10.1016/j.jdcr.2020.06.009

<original_ref>Cozzani E., Herzum A., Burlando M., Parodi A., Comedonal variant of chronic
cutaneous lupus erythematosus causing mutilation of the earlobe, JAAD Case Rep., 6, 2020,
843- 844, 10.1016/j.jdcr.2020.06.009/original_ref>

Figure 1.



Fig 1a



Fig 1b

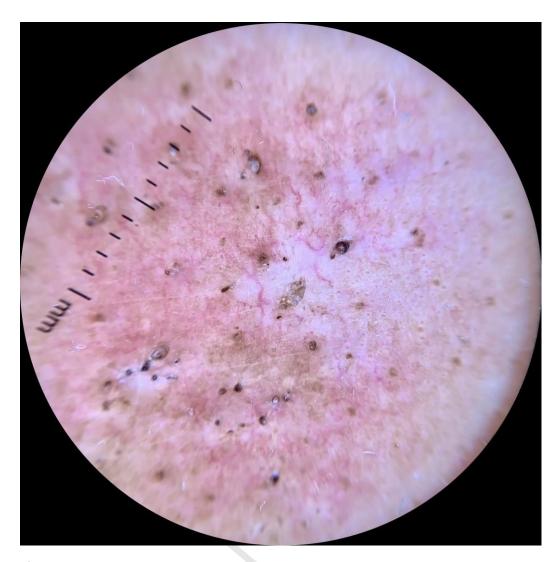


Fig 1c

Figure 2.

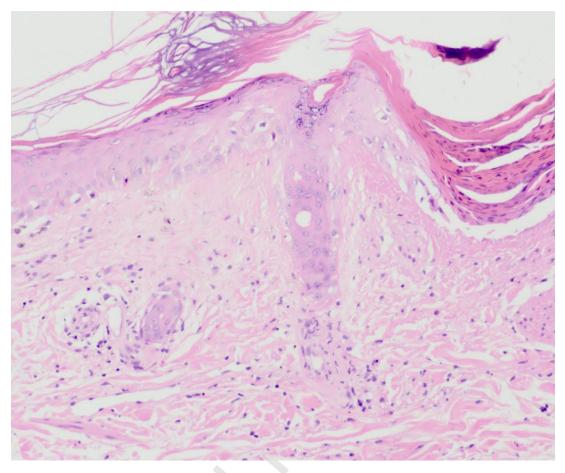


Fig 2a

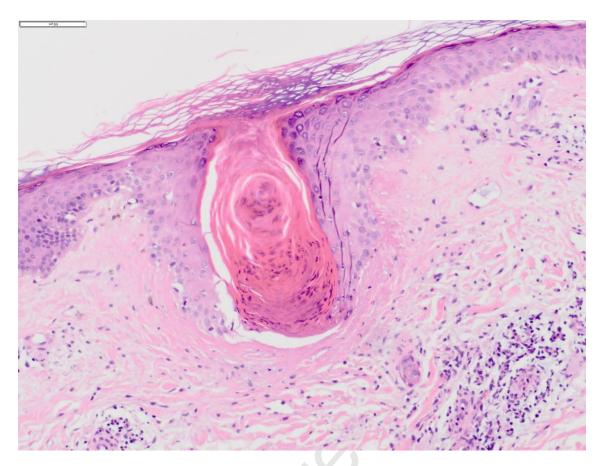


Fig 2b

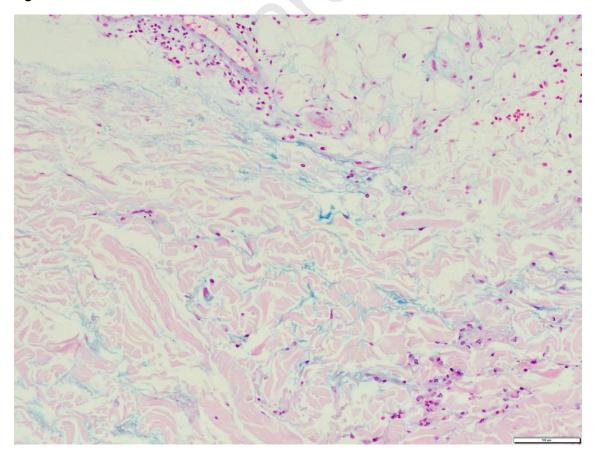


Fig 2c

Figure 3.

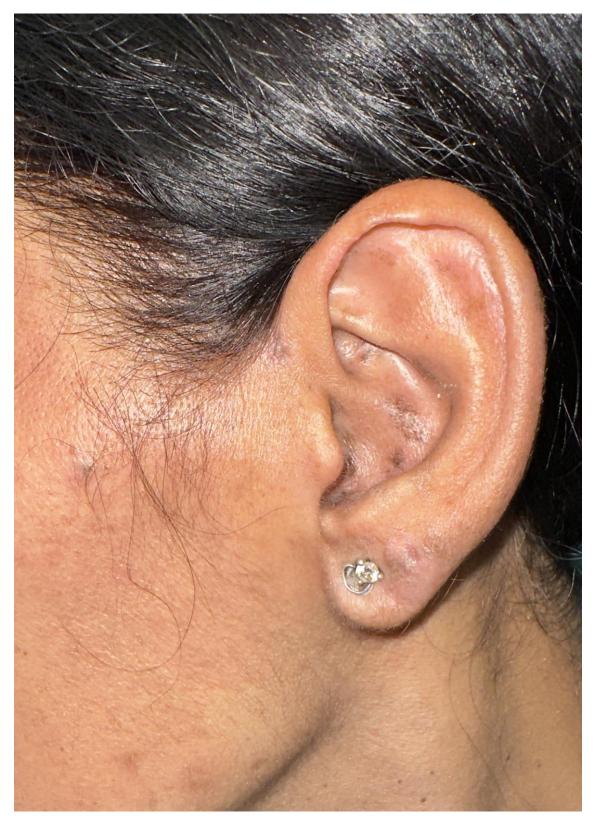


Fig 3a



Fig 3b



Fig 3c