Dermatopathology

Cytokeratin expression in pilonidal sinus

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Summary

Background Pilonidal sinus (PS) is considered to belong in the category of follicular occlusion diseases (acne triad).

Objectives The aim of our study was to elucidate the pathogenesis of PS by evaluating its cytokeratin (CK) expression.

Methods CK expression in nine cases of PS was studied immunohistochemically using six antikeratin antibodies.

Results Infundibular-like epithelium contained CK1, 10 and 14 similar to normal infundibulum, but it did not contain CK17. In non-infundibular-like epithelium, CK14, 16 and 17 were detected similar to that in normal outer root sheath. CK expression in PS was similar to that in hidradenitis suppurativa, suggesting that sinus epithelium may be fragile, hyperproliferative and undifferentiated.

Conclusions PS can be classified in the same entity as follicular occlusion diseases based on CK expression.

Key words: cytokeratin, pilonidal sinus

Pilonidal sinus (PS), also called pilonidal cyst with epithelial lining as draining sinus, occurs most frequently in the middle line of the gluteal fold. It sometimes becomes inflamed, resulting in a subcutaneous abscess with recurrent infection. Histopathological findings in PS show follicular hyperkeratosis of the infundibulum with plugging and dilatation of the follicle. The early inflammatory event is perifolliculitis with neutrophils, lymphocytes and histiocytes, leading to rupture of follicular epithelium. Once the follicular epithelium had been ruptured, foreign-body materials such as corneocytes, bacteria, sebum and hairs were spilled into dermis and subcutaneous tissue, giving rise to the forming of granulomatous lesions and fibrosis. In 1956, Pillsbury et al. advocated the concept of the follicular occlusion triad (acne triad), which includes acne conglobata, hidradenitis suppurativa (HS) and dissecting cellulitis. Recently, Plewig and Steger proposed a new concept of acne tetrad that includes PS with acne triad. In addition, Plewig et al. advocated a new term: acne inversa. Although the aetiology in PS remains ambiguous, gender, onset at puberty, familial disposition, hypertrichosis and obesity have been regarded as contributory factors. The pathogenesis of PS is concerned with an HS-related disorder demonstrating follicular retention hyperkeratosis in the infundibulum.

Among the intermediate filaments, cytokeratin (CK) is the most diversified cytoskeleton tonofilament in epithelial cells, divided into 20 subclasses according to molecular weight and isometric pH. CK is a useful marker for the differentiation of epithelial cells and facilitates evaluation of the origin of epithelial tumours to elucidate the pathogenesis of PS. Recently, keratin expression of the sinus tract in HS has been investigated. However, CK expression in PS has not been studied. The aim of our study was to clarify immunohistochemically CK expression in PS, and to compare that with the previous study in HS.
Materials and methods

Samples of PS were surgically excised from nine male patients. The age of the patients ranged from 22 to 46 years. All specimens were obtained from the buttock. These specimens were cut into 5–7 μm thickness, then fixed in formalin, and embedded in paraffin. The sections were stained with haematoxylin and eosin for histopathological examination. For the immunohistochemical study, a deparaffinized sample of each respective section was incubated with six different antikeratin antibodies for 30 min. The antikeratin antibodies used in this study were 34βB4 (CK1), LHP1 (CK10), LL001 (CK14), LL025 (CK16), E3(CK17) and b190 (CK19) (all purchased from Novo Castra Laboratories Ltd, Newcastle upon Tyne, U.K.). Immunohistochemical procedures were performed according to the labelled streptavidin–biotin method (LSAB, Dako, Carpentry, CA, U.S.A.), as previously reported.

Results

Based on the histological findings, the epithelial lining in PS was composed of two distinct types of cornifying epithelium. The former, similar to infundibular-like keratinization, was classified as type A epithelium, which had a granular layer with keratohyaline granules (Fig. 1). The latter, non-infundibular-like keratinization without keratohyaline granules was classified as type B epithelium (Fig. 2). Of the nine cases, type A epithelium was found in seven, and type B in two.

CK expression in normal pilosebaceous units, PS and HS is summarized in Table 1. Immunohistochemical expression of CK in normal pilosebaceous units was

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**Figure 1.** Type A epithelium, an infundibular-like epithelium, has granular layers with keratohyalin granules (arrow) in pilonidal sinus (haematoxylin and eosin, original magnification ×100).

**Figure 2.** Type B epithelium, non-infundibular-like epithelium, has no granular layers with keratohyalin granules (arrow) (haematoxylin and eosin, original magnification ×100).
referred to in a previous report. In brief, CK1 and 10 were present in suprabasal layers of the infundibulum above the orifice of the sebaceous duct. CK14 was found in basal layers of the infundibulum, and in all layers of the outer root sheath, and sebaceous apparatus. CK16 was detected in suprabasal layers of the outer root

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<th>SD</th>
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CK, cytokeratin; PS, pilonidal sinus; HS, hidradenitis suppurativa; IF, infundibulum; ORS, outer root sheath; SD, sebaceous duct; SC, sebaceous cells; sb, suprabasal; b, basal.
sheath below the opening of sebaceous duct. CK17 was present in suprabasal layers of the infundibulum (Fig. 3) and sebaceous duct, and in all layers of the outer root sheath. CK19 was found in the outermost layer of the outer root sheath in the hair bulge.

Concerning CK expression in PS, CK10 was found in suprabasal layers of type A epithelium (Fig. 4) as well as CK1. CK14 was observed in the basal layer of type A epithelium (Fig. 5). Compared with the presence of CK17 in normal infundibulum (Fig. 3), CK17 was absent in type A epithelium in PS (Fig. 6). CK16 and 19 were not present in type A epithelium. In type B epithelium, CK1 and 10 were not found. CK14 was seen at all layers of type B epithelium. A strong positive reaction for CK16 was found in the suprabasal layers of type B epithelium (Fig. 7) as well as CK17. CK19 was not found in type B epithelium. CK expression of overlying epidermis in PS was consistent with that in normal epidermis.

**Figure 5.** In pilonidal sinus, CK 14 was present in basal layer of type A epithelium (arrow) (original magnification ×100).

**Figure 6.** In pilonidal sinus, CK17 was absent in type A epithelium (original magnification ×100).

**Discussion**

PS belongs in the follicular occlusion tetrad. The histopathology of PS has been reported. Søndenaa and Polland showed that the epithelial lining in PS has keratohyaline granules, and is derived from the epidermis. However, in our study, type B epithelium, which had no keratohyaline granules, was clearly observed.

As a whole, CK expression in PS was comparable with that in HS as previously reported. Type A epithelium resembles infundibular epithelium because of the presence of CK1 and 10 in the suprabasal layers and CK14 in the basal layer, and the absence of CK16 and 19. However, CK17 was not found in type A epithelium. In the infundibulum, CK17 was found in the suprabasal layers of normal skin. CK17, spatial keratin, has a function related to the three-dimensional cytoskeleton structure. The absence of CK17 in type

A epithelium may reflect a fragile follicular structure, resulting in the rupture of the follicle, which subsequently forms a subcutaneous abscess. The cyst formation may be related to the mutation of CK17. The immunohistochemical expression of type B epithelium was comparable with that of the outer root sheath. Like the outer root sheath in normal skin, CK14 was found in all layers, and CK16 and 17 were present in the suprabasal layers of type B epithelium. CK1 and 10 were absent in type B epithelium as well as in the outer root sheath. The results of our study suggested that type B epithelium is more undifferentiated and hyperproliferative.

From the perspective of CK expression, CK expression in PS was comparable with that in HS, which supports the concept of Plewig et al. who have advocated that PS belongs in the follicular occlusion tetrad (acne tetrad) category.

References