Published online 2022 February 25.

Research Article

Clinicopathological Evaluation of Seborrheic Keratosis Lesions in Patients Referred to Afzalipour Hospital, Kerman, Iran

Maryam Khalili¹, Simin Shamsi Meymandi², Rezvan Amiri¹, Mahin Aflatoonian^{3,*} and Shirin Alimortazavi⁴

¹Leishmaniasis Research Center, Kerman University of Medical Sciences, Kerman, Iran

²Pathology and Stem Cell Research Center, Kerman University of Medical Sciences, Kerman, Iran

³Department of Dermatology, Kerman University of Medical Sciences, Kerman, Iran

⁴Kerman University of Medical Sciences, Kerman, Iran

corresponding author: Department of Dermatology, Kerman University of Medical Sciences, Kerman, Iran. Email: maaflatoonian@gmail.com

Received 2022 January 02; Revised 2022 February 06; Accepted 2022 February 07.

Abstract

Background: Seborrheic keratosis is a benign epidermal proliferation, which is highly common in sun-exposed areas. This study evaluated the clinicopathological characteristics of seborrheic keratosis lesions in patients referred to the Afzalipour Hospital, Kerman, Iran.

Methods: This retrospective cross-sectional study investigated ninety-nine skin biopsies of seborrheic keratosis lesions. The patients' demographic features and the clinical and pathological features of the lesions were recorded. Then the correlation between pathological subtypes and demographics and clinical features was evaluated. Independent *t*-test and chi-square tests were used to assess the correlation between quantitative and qualitative data, respectively.

Results: A majority of the patients were female (56.6%) in the sixth decade of their lives (33.3%). The lesions were more frequent in sun-exposed areas (65.6%). The most common pathological subtypes were acanthotic (47.5%), hyperkeratotic (27.3%), and adenoid (14.1%), and horn cyst (75.8%) and squamous eddies (5.1%) were the most and the least prevalent pathological features, respectively. Moreover, no significant correlation was noticed between pathological subtypes with the patients' age or sun-exposed areas (P = 0.257 & P = 0.05, respectively)

Conclusions: The most common pathological subtype in this study was the acanthotic type. There was no correlation between pathological subtypes and the patients' demographic features. The most common clinicopathological correlation was associated with the sun-protected lesions.

Keywords: Seborrheic Keratosis, Clinic, Pathology

1. Background

Seborrheic keratosis (SK) is a benign epidermal proliferation. After the fourth decade of life, SK is more common in sun-exposed areas in the Caucasians. The lesions usually develop equally in both genders. They are generally asymptomatic; however, inflamed, pruritic, or tender lesions follow trauma or infections (1-5). They usually appear as multiple brown, black, or light brown verrucous papules with about a one-centimeter diameter. The clinical subtypes of SK are dermatosis papulosa nigra, stucco keratosis, and inverted follicular keratosis. SK is mainly diagnosed clinically; however, skin biopsy is sometimes necessary to differentiate it from other skin tumors (1-5). The pathological subtypes of the disease are acanthotic, hyperkeratotic, melanoacanthoma, adenoid (reticulated), irritated, and clonal (nested) (1-5).

2. Objectives

In this study, the clinicopathological characteristics of the SK lesions in patients referred to the Afzalipour Hospital, Kerman, were evaluated.

3. Methods

This retrospective cross-sectional study examined ninety-nine skin biopsies from patients referred to the Afzalipour Hospital, Kerman, with the diagnosis of SK. The participants' demographic features (namely age and gender), the clinical features of the lesions (site and clinical differential diagnosis), and the pathological features of the lesions were recorded. Skin biopsies were revised with two dermatopathology fellowships to confirm the

Copyright © 2022, Journal of Skin and Stem Cell. This is an open-access article distributed under the terms of the Creative Commons Attribution-NonCommercial 4.0 International License (http://creativecommons.org/licenses/by-nc/4.0/) which permits copy and redistribute the material just in noncommercial usages, provided the original work is properly cited.

diagnosis and detect the pathological subtypes. Then the correlation between clinical diagnosis and pathological diagnosis was evaluated. Furthermore, the correlation between the pathological subtypes with demographic features and the site of the lesions was evaluated.

3.1. Statistical Analysis

Data were analyzed using SPSS software version 16 (software IBM, Armonk, NY, USA). Frequency, percentage, and mean \pm standard deviation were used for descriptive analysis. Independent *t*-test and chi-square tests were also used to assess the correlation between quantitative and qualitative variables, respectively.

4. Results

Ninety-nine skin biopsies with the diagnosis of SK were evaluated in this study. Most of the patients were female (56.6%). The mean age of the male patients was 49.79 \pm 15.82 years (range 12 - 85), and the mean age of the females was 52.64 \pm 16.50 years (range 16 - 80). There was no significant difference between the two genders regarding the patients' age (P = 0.4). Most of the patients were in the sixth decade of their lives (33.3%), and sun-exposed areas were the most common site of the lesions (65.6%, Table 1).

Table 1. Prevalence of Site of Lesions and Pathological Features of Seborrheic Keratosis Lesions

Variables	No. (%)			
Site of the lesions				
Head & Neck	58 (58.6)			
Trunk	10 (10.1)			
Lower limb	8 (8.1)			
Upper limb	23 (23.2)			
Pathological features				
Horn cyst	75 (75.7)			
Hyperkeratosis	71 (71.7)			
Acanthosis	60 (60.6)			
Pigmentation	50 (50.5)			
Papillomatosis	48 (48.5)			
Infiltration	39 (39.4)			
Squamous eddies	5 (5.1)			

The most common pathological subtypes were acanthotic (47.5%), hyperkeratotic (27.3%), adenoid (14.1%), clonal (7.1%), and irritated (4%), respectively. The most and the least common pathological features were horn cysts (75.8%) and squamous eddies (5.1%), respectively (Table 1). All pathological diagnoses were correlated with clinical diagnoses, and the correlation of between the first clinical diagnosis and the pathological diagnosis was 65.7%. The hyperkeratotic and acanthotic subtypes were more common in females than males (66.7% vs. 33.3% and 55.3% vs. 44.7%, respectively). Moreover, the adenoid and clonal subtypes were observed more frequently in males than females (57.2% vs. 42.9% and 57.1% vs. 42.9%, respectively); however, the difference was not statistically significant (P = 0.507, Table 2). Moreover, there was no significant correlation between pathological subtypes with either the patients' age or sun-exposed areas (P = 0.257 & P = 0.05, respectively) (Tables 2 and 3). The most common clinical diagnoses in were SK (65.7%), wart (13.1%), melanocytic nevus (6.1%), basal cell carcinoma (BCC, 5.1%), melanoma (3%), skin tag (2%), lichenoid keratosis (2%), lentigo (1%), and Bowen's disease (1%), respectively. The most common clinicopathological correlation was observed in sun-protected areas rather than sun-exposed ones (74% and 27%, respectively); however, the difference was not statistically significant (P = 0.119).

5. Discussion

In this study, the patients were mainly in the sixth decade of their lives (33.1%), and this finding is in line with the findings of other studies (6-9). Furthermore, in the present study, 7.1% of the lesions emerged in the patients aged below 20 years. In Gill et al.'s study (10), the percentage was roughly double (15.7%). This can be explained by inconsistencies in the type of the studies, geographical areas, ethnic and genetic backgrounds, cultural differences, percentage of sun-exposure, and type of covering (10).

In this study, there was no significant difference between males and females regarding the prevalence of SK. Furthermore, in line with previous studies (10-12), the male and female patients were almost homogenous in terms of age.

Although the exact pathogenesis of the SK was not wellunderstood, factors such as sun exposure, genetic predisposition, and genetic mutation in FGFR3 due to ultraviolet (especially in adenoid type) were also considered. Most of the lesions were located in sun-exposed areas (65.6%) in the present study. In other studies in Iran and South Korea, most of the lesions were similarly located in sun-exposed areas (78.3% and 56.3%, respectively) (7, 9). Furthermore, there was no significant difference between pathological subtypes and sun-exposure in the present study. In contrast, Roh et al. reported the higher prevalence of the adenoid type in sun-exposed areas (7).

In this study, all pathological diagnoses were correlated with clinical diagnoses, and the correlation between the first clinical diagnosis and pathological diagnosis was

Variables	Total	Hyperkeratotic	Adenoid	Acanthotic	Clonal	Irritated	P-Value
Age (y)							0.25
10 - 20	7 (7.1)	2(7.4)	1(6.7)	4 (8.5)	0(0)	0(0)	
21-30	7 (7.1)	1(3.7)	2 (13.3)	4 (8.5)	0(0)	0(0)	
30 - 40	10 (10.1)	1 (3.7)	2(20)	6 (12.8)	1 (14.3)	0(0)	
41 - 50	18 (18.2)	6 (22.6)	2 (13.4)	7 (14.9)	2 (28.6)	1(25)	
51 - 60	29 (29.3)	4 (14.8)	2 (13.3)	19 (40.4)	2 (28.6)	2 (50)	
61 - 70	13 (13.1)	4 (4.8)	2 (13.3)	4 (8.5)	2 (28.6)	1(25)	
71 - 80	14 (14.1)	9 (33.3)	2 (13.3)	3(6.4)	0(0)	0(0)	
81 - 90	1(1)	0(0)	1 (7.1)	0(0)	0(0)	0(0)	
Sex							0.5
Male	43 (43.4)	9 (33.3)	8 (57.2)	21 (44.7)	4 (57.1)	1(25)	
Female	56 (56.6)	18 (66.7)	6(42.9)	26 (55.3)	3(42.9)	3 (75)	

^a Values are expressed as No. (%).

Table 3. Distribution of Subtypes of Seborrheic Keratosis in Both Sun-exposed and Sun-protected Areas

Variables	No. (%)			
Sun-exposed				
Acanthotic	33 (50.8)			
Hyperkeratotic	18 (27.7)			
Adenoid	10 (15.4)			
Irritated	3 (4.6)			
Clonal	1(1.5)			
Sun-protected				
Acanthotic	14 (41.2)			
Hyperkeratotic	9 (26.5)			
Clonal	6 (17.6)			
Adenoid	4 (11.8)			
Irritated	1(2.9)			

65.7%. Moreover, the smallest clinicopathological correlation was associated with lesions on the sun-exposed areas. In these areas, lesions can be misdiagnosed with nonmelanocytic skin tumors [eg, BCC, squamous cell carcinoma (SCC), and melanoma]. The most common differential diagnoses in the present study were wart (13.1%), melanocytic nevus (6.1%), and skin cancers, including BCC (5.1%) and melanoma (3%). Roh et al. demonstrated a disagreement between clinical diagnosis and pathological diagnosis in 27.36% of the patients. The most prevalent clinical diagnoses were wart (42.3%), BCC (11.5%), and SCC, dysplastic nevi, or compound nevi (each 7.7%). This finding was to some extent consistent with the findings of the present study. Likewise, in Roh et al.'s study, higher disagreement between clinical and pathological diagnoses was observed in sun-exposed sites (P = 0.043)(7).

The most common pathological subtypes in the present study were acanthotic and hyperkeratotic types (47.5% & 27.3%, respectively). This finding is in line with those of the the previous studies (8, 9). In this study, the adenoid type was the third pathological subtype (14%); however, the same finding was not recorded in other studies (7, 9, 12). The clinical resemblance between the adenoid type and pigmented lesions such as solar lentigo and lentigo maligna can justify more biopsy of the lesions in the present study (8, 9).

In this study, the most common pathological features were horn cyst (75.8%), acanthosis (71.7%), and hyperkeratosis (60.6%), respectively. In India, Alapatt et al. reported pigmentation (78%), papillomatosis (68%), acanthosis (56%), and hyperkeratosis (54%) as the most common pathological features. The higher percentage of pigmentation in their study compared to the present study (50.5%) can be explained by the darker skin of Indian patients and the large number of clinical subtypes of dermatosis papulosa nigra (26%) (12).

The major limitations of this study were retrospective design and relatively low sample size. Further prospective studies with a larger sample size are recommended.

5.1. Conclusions

In this study, most of the patients were female and in the sixth decade of their lives. The most common pathological subtype was the acanthotic type, and the most prevalent pathological feature was horn cyst. There was no correlation between the pathological subtype and the patients' demographic features.

Footnotes

Authors' Contribution: M. Kh. and M. A. contributed to the study conception and design. Material preparation and data collection were undertaken by all authors. M.A. Sh. A., S. M., and M.Kh. performed the data acquisition, analysis, and interpretation. M. A. and M. Kh. wrote the first draft of the manuscript. All authors revised the final version of the manuscript.

Conflict of Interests: There was no conflict of interest.

Data Reproducibility: The data presented in this study are openly available in one of the repositories or will be available on request from the corresponding author by this journal representative at any time during submission or after publication. Otherwise, all consequences of possible withdrawal or future retraction will be with the corresponding author.

Ethical Approval: IR.KMU.AH.REC.1397.2697.

Funding/Support: No funding was received for this study.

References

- Karadag AS, Parish LC. The status of the seborrheic keratosis. *Clin Dermatol.* 2018;**36**(2):275-7. doi: 10.1016/j.clindermatol.2017.09.011. [PubMed: 29566932].
- Braun RP, Ludwig S, Marghoob AA. Differential Diagnosis of Seborrheic Keratosis: Clinical and Dermoscopic Features. J Drugs Dermatol. 2017;16(9):835–42.

- Rajesh G, Thappa DM, Jaisankar TJ, Chandrashekar L. Spectrum of seborrheic keratoses in South Indians: a clinical and dermoscopic study. *Indian J Dermatol Venereol Leprol.* 2011;77(4):483–8. doi: 10.4103/0378-6323.82408. [PubMed: 21727696].
- Hafner C, Vogt T. Seborrheic keratosis. J Dtsch Dermatol Ges. 2008;6(8):664–77. doi: 10.1111/j.1610-0387.2008.06788.x. [PubMed: 18801147].
- Zalaudek I, Ferrara G, Argenziano G. Clonal Seborrheic Keratosis: A Dermoscopic Pitfall. Arch Dermatol. 2004;140(9). doi: 10.1001/archderm.140.9.1169-b.
- Jackson JM, Alexis A, Berman B, Berson DS, Taylor S, Weiss JS. Current understanding of seborrheic keratosis: prevalence, etiology, clinical presentation, diagnosis, and management. J Drugs Dermatol. 2015;14(10):1119–25.
- Roh NK, Hahn HJ, Lee YW, Choe YB, Ahn KJ. Clinical and Histopathological Investigation of Seborrheic Keratosis. *Ann Dermatol.* 2016;**28**(2):152-8. doi: 10.5021/ad.2016.28.2.152. [PubMed: 27081260]. [PubMed Central: PMC4828376].
- Hafner C, van Oers JM, Hartmann A, Landthaler M, Stoehr R, Blaszyk H, et al. High frequency of FGFR3 mutations in adenoid seborrheic keratoses. J Invest Dermatol. 2006;126(11):2404-7. doi: 10.1038/sj.jid.5700422. [PubMed: 16778799].
- Rajabi P, Adibi N, Nematollahi P, Heidarpour M, Eftekhari M, Siadat AH. Bowenoid transformation in seborrheic keratosis: A retrospective analysis of 429 patients. *J Res Med Sci.* 2012;17(3):217.
- Gill D, Dorevitch A, Marks R. The prevalence of seborrheic keratoses in people aged 15 to 30 years: is the term senile keratosis redundant? *Arch Dermatol.* 2000;**136**(6):759–62. doi: 10.1001/archderm.136.6.759. [PubMed: 10871940].
- Kwon OS, Hwang EJ, Bae JH, Park HE, Lee JC, Youn JI, et al. Seborrheic keratosis in the Korean males: causative role of sunlight. *Photodermatol Photoimmunol Photomed*. 2003;**19**(2):73–80. doi: 10.1034/j.1600-0781.2003.00025.x. [PubMed: 12945806].
- Alapatt GF, Sukumar D, Bhat MR. A Clinicopathological and Dermoscopic Correlation of Seborrheic Keratosis. *Indian J Dermatol.* 2016;61(6):622-7. doi: 10.4103/0019-5154.193667. [PubMed: 27904179]. [PubMed Central: PMC5122276].