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LETTER - RESEARCH

Dermatofibroma: clinicopathological analysis of 239 cases[☆]

Dear Editor,

Dermatofibroma is one of the most commonly encountered benign, cutaneous soft tissue tumors, usually located on the extremities. Lesions are often in the form of solitary, raised, brown, firm, cutaneous nodules. Histopathologically, dermatofibroma is characterized by the proliferation of uniform spindle-shaped fibroblasts and histiocytoid cells in the dermis, with spindle cells forming a storiform appearance.^{1,2} There are sub-variants of dermatofibroma which are less frequently observed. These are lipidized, aneurysmal, hemosiderotic, cellular, clear cell, atrophic, epithelioid, and sclerotic subtypes.^{2,3} Rarely, it can also present in only subcutaneous form without causing changes on the skin surface.⁴

Common dermatofibroma is frequently observed, and rare variants are seen in 20% of histopathologically evaluated dermatofibromas.² Subtypes other than common dermatofibroma might have different clinical manifestations and may cause confusion in the differential diagnosis. Therefore, in our study, clinicopathological evaluation of cases diagnosed with dermatofibroma was aimed. Further evaluation of infrequent variants, their differential diagnosis, and correct clinical diagnosis of dermatologists about the rare variants were analyzed.

The medical records of 239 dermatofibroma cases from 232 patients with clinical and histopathological diagnoses in our dermatology clinic between the years of 2012–2021 were evaluated retrospectively. Patients with suspicious clinical or histopathological diagnoses or who had no clinical photography were excluded from the study. The following characteristics were recorded:

Demographic characteristics: Age, sex.

Clinical characteristics: Type of dermatofibroma, location of the skin lesions, the color of the lesions.

Histopathological characteristics: subtype of the dermatofibroma.

Statistical method: Mean, standard deviation, median minimum, maximum, frequency, and ratio values were used in the descriptive statistics of the data. The distribution of variables was measured with the Kolmogorov-Smirnov test. Mann-Whitney *U* test was used in the analysis of independent quantitative data. The Chi-Square test was used to analyze independent qualitative data, and Fisher's exact test was used when the Chi-Square test conditions were not met. SPSS 28.0 program was used in the analysis.

Results of 239 dermatofibroma cases from 232 patients were included in our study. Patient demographic, clinical, and histopathological characteristics are shown in Table 1. 61.2% of the patients were female, and 38.7% were male, with a mean age of 40.7 ± 12.9 . The most common site of involvement was the lower extremity (50.8%), followed by upper extremity involvement with 35.2%. The lesions had a mean size of $7 \text{ mm} \pm 6.3$. Histopathologically, 84.9% of the cases were common subtypes, followed by 7.5% atrophic, 4.6% cellular, 1.6% lipidized, 0.8% aneurysmal and 0.4% sclerotic subtypes (Figs. 1–3).

While dermatofibroma was considered the first diagnosis in 87.4% of the cases, dermatofibroma was not among the preliminary diagnoses in 12.5%. The most commonly considered differential diagnoses were foreign body granuloma, prurigo nodularis, pilomatricoma, keloid, and dermatofibrosarcoma protuberans with decreasing order (Fig. 4).

The age, gender, and size of the lesion did not differ significantly ($p > 0.05$) between the groups with and without classical Histopathology (HP). The age of the patients in the group with differential diagnosis DF was significantly ($p < 0.05$) lower than the group without DF. Gender distribution did not differ significantly ($p > 0.05$) between groups with and without differential diagnosis of DF. The size of the lesion was significantly higher ($p < 0.05$) in the group with a pre-diagnosis of DF than in the group without a pre-diagnosis of DF. The rate of common histopathology in the group with differential diagnosis of DF was significantly higher ($p < 0.05$) than in the group without differential diagnosis of DF (Table 2).

Dermatofibroma is one of the most common benign skin tumors encountered in clinical practice. Although the diagnosis of dermatofibroma is straightforward, the diagnosis of rare histopathological subtypes may be challenging. In our

[☆] Study conducted at the Istanbul Training and Research Hospital, Department of Dermatology, Istanbul, Turkey.

Table 1 Clinical and histopathological characteristics of Dermatofibroma (DF) patients.

		Min-Max	Median	Avg. ± Std dev
			N	%
Age		12.0–72.0	40.0	40.77 ± 12.92
Size (mm)		2.0 - 45.0	5.0	7.05 ± 6.31
Gender	Female		142	61.2
Male		90	38.7	
Location	Lower extremity		121	50.8
Upper extremity		84	35.2	
Trunk		29	12.1	
Face		4	1.6	
Clinical differential diagnosis of DF	(-)		30	12.5
(+)		209	87.4	
Classical DF Histopathology	(-)		36	15
(+)		203	84.9	
Histopathological sub-variant	Classical		203	84.9
Atrophic		18	7.5	
Cellular		11	4.6	
Aneurysmal		2	0.8	
Lipidized		4	1.6	
Sclerotic		1	0.4	

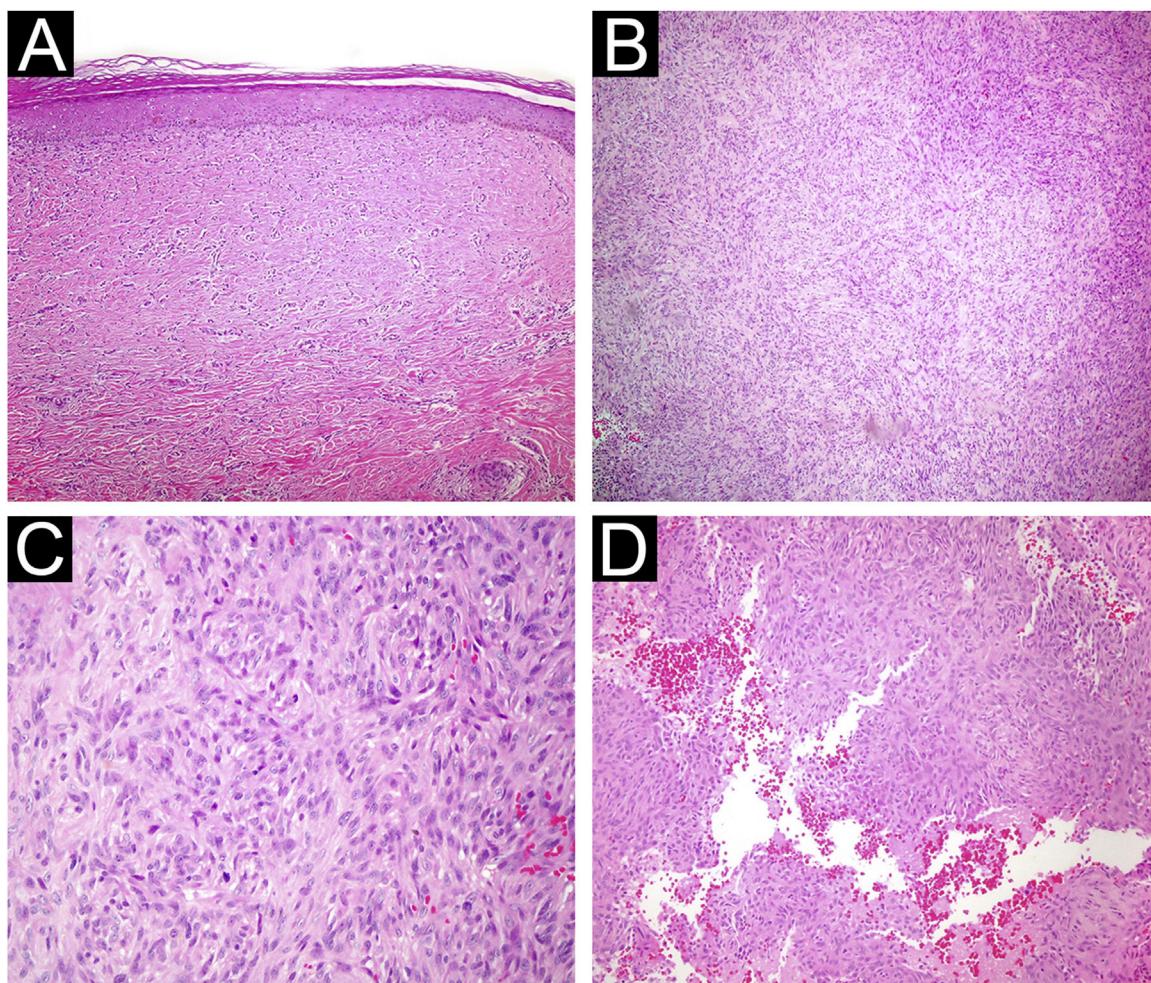


Fig. 1 (A) Atrophic dermatofibroma. The lesion is hypocellular, and exhibits marked hyalinization. (Hematoxylin & eosin, $\times 200$) (B) Cellular dermatofibroma with noticeable cellularity. (Hematoxylin & eosin, $\times 100$) (C) High magnification of cellular dermatofibroma showing prominent mitotic figures. (Hematoxylin & eosin, $\times 400$) (D) Aneurysmal fibrous histiocytoma. Cystic spaces filled with blood are observed within the lesion. (Hematoxylin & eosin, $\times 200$).

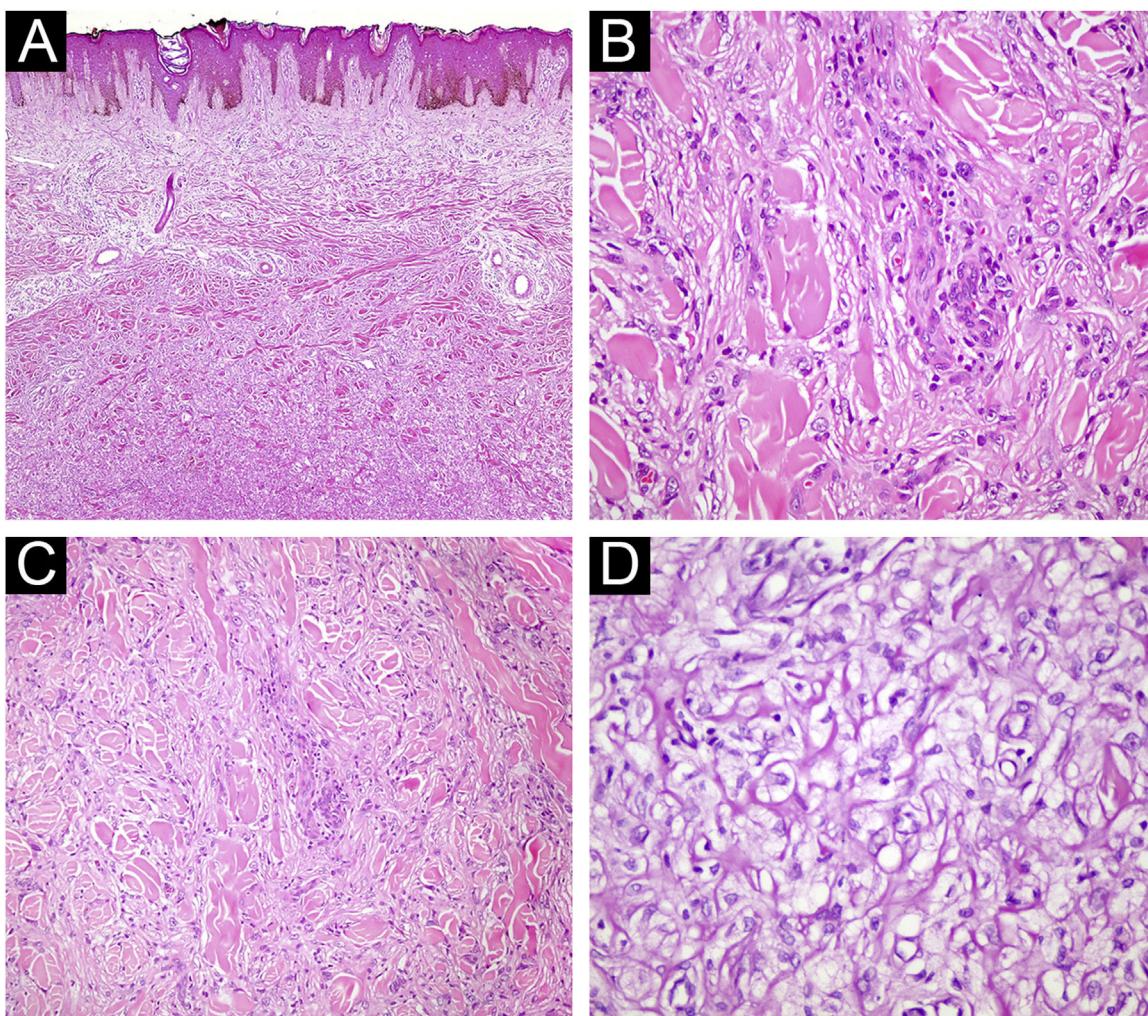


Fig. 2 (A) Fibrous histiocytoma with atypical features. Marked nuclear pleomorphism is evident (Hematoxylin & eosin, $\times 200$). (B) Fibrous histiocytoma with atypical features. Bizarre nucleated cells are observed (Hematoxylin & eosin, $\times 400$). (C) Lipidized fibrous histiocytoma. At low magnification, significant hyalinization is noticeable in the lesion (Hematoxylin & eosin, $\times 40$) (D) Foamy cytoplasmic appearance in the cytoplasm of lesional cells (Hematoxylin & eosin, $\times 400$).

Table 2 Dermatofibroma (DF) diagnosis according to clinical variables.

	Pre-diagnosis of DF(-) (n = 40)		Pre-diagnosis of DF(+) (n = 209)		Median	p
	Avg \pm SD/n (%)	Median	Avg \pm SD/n (%)	Median		
Age	47.21 \pm 13.73	50.0	39.85 \pm 12.56	39.0	0.005 ^a	
Gender	Female 15 (38.5%)		127 (62.6%)		0.263 ^b	
Male	14 (35.9%)	76 (37.4%)				
Size	5.17 \pm 4.24	4.0	7.33 \pm 6.51	5.0	0.043 ^a	
Classical DF	(-) 11 (36.7%)		25 (12.0%)		0.000 ^b	
(+)	19 (63.3%)	184 (88.0%)				

^a Mann-whitney U test.

^b Chi-Square test.

study, rare variants were observed at a rate of 15%. It has been reported in the literature that dermatofibromas can mimic many different lesions, as they can be encountered in different colors and various clinical spectrums. Foreign body granuloma, prurigo nodularis, and pilomatricoma were the most common differential diagnoses in our clinical practice.

In a study that included 30 aneurysmal fibrous histiocytomas, intradermal nevi, leiomyomas, or benign vascular tumors were often considered at the initial diagnosis. Focal thrombosis and the amount of hemosiderin were thought to be the factors affecting the color of the presentation.⁵ Similarly, in our study, a case of aneurysmal fibrous histiocytoma

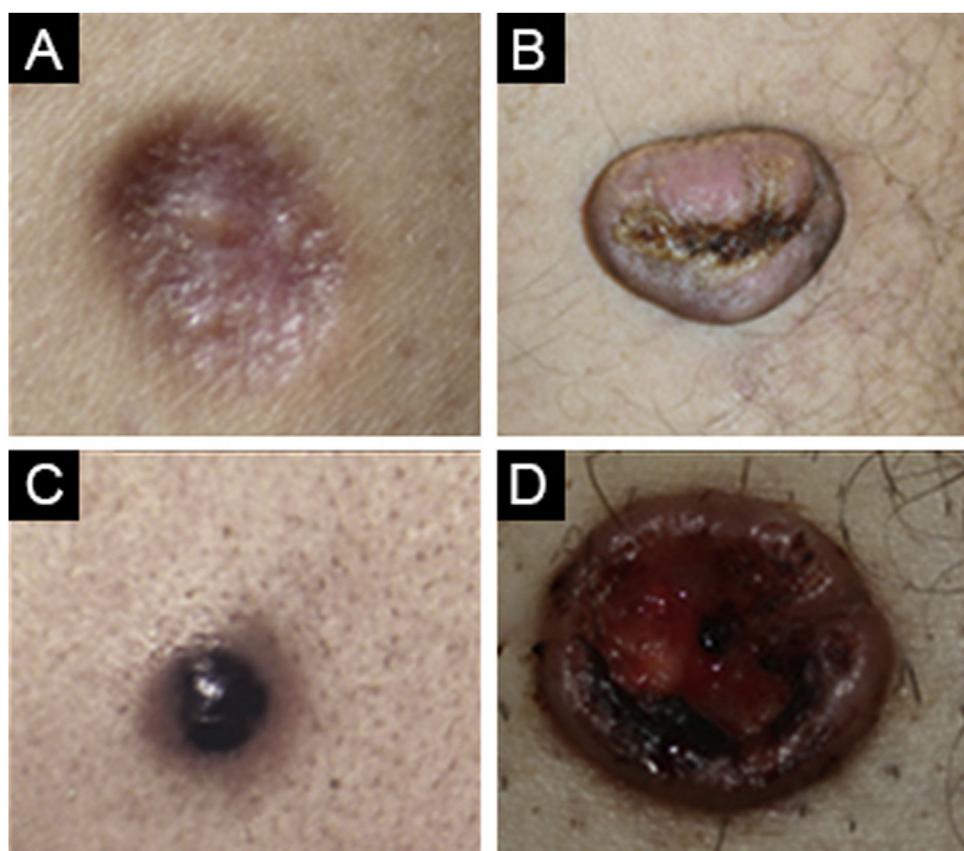


Fig. 3 (A) Atrophic dermatofibroma. (B) Cellular dermatofibroma. (C) Aneurysmal dermatofibroma. (D) Lipidized dermatofibroma.

presented with a black color and caused difficulties in the differential diagnosis.

Lipidized dermatofibroma may be in the form of a large exophytic yellow nodule, usually around the ankle, and may be clinically confused with juvenile xanthogranuloma. Lipidized dermatofibroma usually involves the lower extremities, while juvenile xanthogranuloma affects the head and neck region.^{2,6} All of our lipidized dermatofibroma cases were localized in the lower extremities.

In terms of cellular dermatofibroma, the distinction from Dermatofibrosarcoma Protuberans (DFSP) gains importance. One of its distinctive features is that it can clinically involve different localizations, such as the face, hands and feet, apart from the extremities.⁷ The human progenitor cell antigen (CD34) positivity is up to 50%–100% in DFSP, while focal positivity of CD34 can be detected at < 20% in DF.⁸ In a recent study, fat necrosis accompanied by lymphocytic infiltrate was seen in all cellular dermatofibroma cases, and its absence in DFSP cases shows that it can be helpful in histopathological diagnosis.⁹ While dermatofibroma was considered the initial diagnosis in 87.0% of the cases, dermatofibroma was not among the preliminary diagnoses in 12.5%. Among these cases, 17.0% of them were cellular dermatofibroma. Therefore differential diagnosis with DFSP or other malignancies is essential.

The main limitation of our study is its retrospective design. However, having photography of all our patients has made clinical assessment possible. The absence of other

rare histopathological subtypes of dermatofibroma might be another limitation in our study.

In conclusion, dermatofibroma is one of the most common benign soft tissue tumors with a straightforward diagnosis. On the other hand, recognizing the rare variants of dermatofibroma might help clinicians to differentiate from the mimickers.

Authors' contributions

Dilara İlhan Erdil: Approval of the final version of the manuscript; critical literature review, data collection, analysis and interpretation; effective participation in research orientation, intellectual participation in propaedeutic and/or therapeutic management of studied cases; manuscript critical review; preparation and writing of the manuscript; statistical analysis, study conception and planning.

Cem Leblebici: Approval of the final version of the manuscript; data collection, analysis and interpretation; effective participation in research orientation; intellectual participation in propaedeutic and/or therapeutic management of studied cases; manuscript critical review.

Duygu Erdil: Approval of the final version of the manuscript; data collection, analysis and interpretation; effective participation in research orientation; intellectual participation in propaedeutic and/or therapeutic management of studied cases; manuscript critical review.

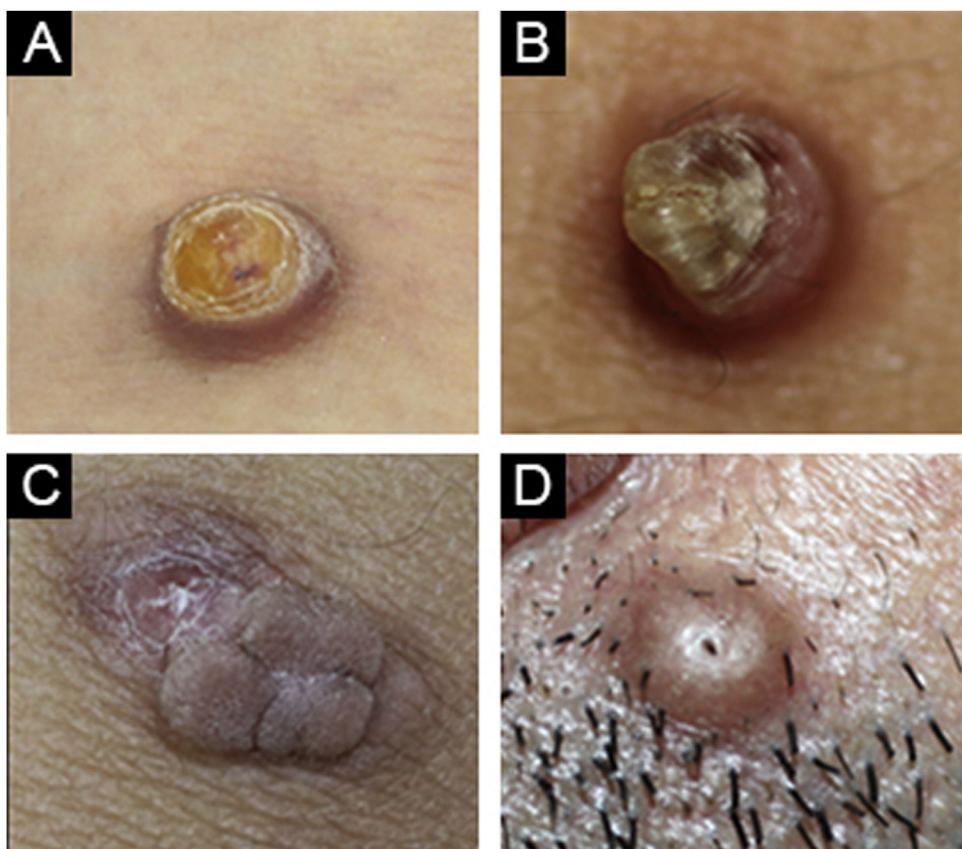


Fig. 4 (A) Cellular dermatofibroma case with pre-diagnosis of eccrine poroma. (B) Common dermatofibroma case with a pre-diagnosis of keratoacanthoma. (C) Cellular dermatofibroma case with a pre-diagnosis of irritated skin-tag. (D) Common dermatofibroma case with a pre-diagnosis of trichofolliculoma.

Vildan Manav: Approval of the final version of the manuscript; data collection, analysis and interpretation; effective participation in research orientation; intellectual participation in propaedeutic and/or therapeutic management of studied cases; manuscript critical review.

Vefa Aslı Erdemir: Approval of the final version of the manuscript; data collection, analysis and interpretation; effective participation in research orientation; intellectual participation in propaedeutic and/or therapeutic management of studied cases; manuscript critical review.

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Conflicts of interest

None declared.

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