

STUDY OF VESICULOBULLOUS DISORDERS IN NORTH KARNATAKA PATIENTS

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ABSTRACT

Background: Blistering disorders are alarming skin conditions. Autoimmune blistering diseases are a group of bullous disorders characterized by pathogenic antibodies directed at the antigens in the epidermis or dermoepidermal junction. These diseases can be viewed histopathologically and direct immunofluorescence. **Materials and Methods:** 50 (fifty) adult patients having vesiculobullous skin lesions were studied. Skin biopsies were done and processed with paraffin. Histological examination of H and E-stained sections was viewed. The level split lesions, mechanism of blistering, and nature of infiltration was noted. **Result:** The highest plane of separation was 22 (44%) suprabasal, followed by 14 (28%) sub-epidermal, and the least was sub-corneal + sub-epidermal, dermal + sub-epidermal, 1 (2%). The highest number of inflammatory infiltrates was 21 (42%), and the least was spongiosis + BMZ destruction, 2 (4%). The observation of the mechanism of inflammation had 18 (36%). Pemphigus vulgaris had the highest number, followed by 10 (20%) bullous pemphigoid, and the least number of patients were 1 (2%) with Darier's disease, Hailey Hailey, and drug reaction. In immunofluorescence diagnosis, the highest number was 14 (28%) for pemphigus, followed by 8 (16%) for bullous pemphigoid. **Conclusion:** Histo-pathological and immune fluorescence approach is an ideal technique to rule out types of vesiculobullous disease and treat efficiently.

INTRODUCTION

Blistering disorders have been known to man since ancient times.^[1] The first recorded episode of pemphigus disease was Hippocrates (460-370 BC), who described pemphigoid fever as pemphigoides pyretol, and Galen (AD 131 to 201) named a pustular disease of the mouth as febris pemphigoides. Vesiculobullous disorders represent a heterogeneous group of dermatoses with protean manifestations. They have a remarkable impact on the patients and their families with severe economic consequences. Five principal mechanisms that can result in blister formation are genetic derangement,^[2] physical and immunological and inflammatory damage, and the drug reactions of these immunological reactions account for most of the vesiculobullous dermatological diseases.

The main pathogenic mechanism involved is the binding of antibody to the antigen, which may directly interfere with desmosomal function.^[3] This

results in spongiosis, acantholysis, reticular degeneration, cytolysis, and basement membrane zone destruction with blister formation. The plane of cleavage depends upon the antigen involved.^[4] Hence, an attempt is made to evaluate histopathological and immunofluorescence staining patterns and establish clinico-pathological correlations in patients presenting with vesiculobullous lesions.

MATERIALS AND METHODS

50 (fifty) adult patients aged between 40 to 60 years regularly visited the dermatology department of Khaja Banda Nawaz Hospital, Kalaburgi, Karnataka-585104 were studied.

Inclusion Criteria: Patients aged above 40 years presenting clinically with vesiculobullous lesions suggestive of autoimmune. The patients who gave their consent in writing for studies that were selected.

Exclusion Criteria: Patients aged below 40 years, viral herpes simplex, varicella, hand-foot, mouth disease, herpes zoster. Fungi: candidiasis, Bacterial: Congenital syphilis, bullous impetigo, and staphylococcal scalded skin syndrome, pregnant and lactation mothers were excluded from the study.

Method: The skin biopsies were taken for histopathology and immunofluorescence study. A biopsy was taken from the site of the lesion, including an intact vesicle.

An elliptical piece of skin was taken from the site of lesions. The specimen was sent in 10% buffered formalin for routine histopathological analysis to the histopathology laboratory of the pathology department. It was processed, and paraffin embedding was done. Serial sections were taken using a rotatory microtome, which were then deparaffinized and stained with hematoxylin and eosin. Histologic examination was then done with the aid of a light microscope. The level of split lesions, the presence of acantholytic cells, and the nature of the infiltrate were noted.

For direct immunofluorescence study, a part of the biopsy material from the peri-lesional area was cut and put in Michel's medium (transport medium containing ammonium sulfate N-ethyl maleimide citrate buffer). The specimen was then sent for an immunofluorescence study.

Biopsy specimen was rinsed in Phosphate Buffered Saline (PBS) at PH 7.4 for 10 minutes. Snap freezing with embedding medium was carried out in the cryostat at 20°C, and a 3-4 micron section was obtained. A drop of specific reagent, fluorescein isothiocyanate (FITC)-labeled antihuman IgG, IgM, IgA, complement C3, and fibrinogen was used. Reading was done under the fluorescence microscope (Nikon model HB 1010AF). The pattern of

distribution and type of immune reactants deposited in the intercellular space was noted.

The duration of the study was from June 2022 to January 2025.

Statistical Analysis: Types of blister separation in various diagnoses were classified with percentages. The statistical analysis was carried out using SPSS software. The ratio of male and female was 2:1.

RESULTS

Table 1: Study of Blister Separation Plane:- 6 (12%) subcorneal, 22 (44%) suprabasal, 14 (28%) subepidermal, 4 (8%) subcorneal + suprabasal, 2 (4%) suprabasal + subepidermal, 1 (2%) subcorneal + subepidermal, 1 (2%) dermal + subepidermal.

Table 2: Study of the character of inflammatory infiltrate 10 (20%) spongiosis, 21 (42%) acantholysis, 14 (28%) BMZ destruction, 3 (6%) spongiosis + acantholysis, 2 (4%) spongiosis + BMZ destruction.

Table 3: Study of Mechanism of Inflammation: 18 (36%) pemphigus vulgaris, 10 (20%) bullous pemphigoid, 4 (8%) spongiotic dermatitis, 2 (4%) pemphigus foliaceus, 2 (4%) Subcorneal pustular dermatosis, 2 (4%) lichen planus pemphigoides, 1 (2%) drug reaction, 1 (2%) Hailey Hailey, 4 (8%) inconclusive descriptions, 5 (10%) other single cases.

Table 4: Study of Immo-fluorescence Diagnosis 14 (28%) pemphigus, 8 (16%) bulous pemphigoid, 1 (2%) Bullous Pemphigoid / Epidermolysis, Bullous Acquisito, 1 (2%) chronic linear IgABullousa dermatosis, 1 (2%) Epidermolysis Bullosa Acquisita, 1 (2%) lichen Planus pemphigoides, 1 (2%) Bullous lupus erythematous, 1 (2%) lupud Bond, 2 (4%) No features of Immuno bulbous disease.

Table 1: Study of Blister Separation plane

Plane of separation	No. of cases	Percentage (%)
Sub corneal	6	12
Supra basal	22	44
Sub epidermal	14	28
Dermal	0	--
Sub corneal + Supra basal	4	8
Supra basal + subepidermal	2	4
Sub corneal + sub epidermal	1	2
Dermal + sub epidermal	1	2

Table 2: Inflammatory infiltrate

Characters of inflammatory infiltrate	No. of patients (50)	Percentage (%)
Spongiosis	10	20
Acantholysis	21	42
BMZ destruction	14	28
Spongiosis + Acantholysis	3	6
Spongiosis + BMZ destruction	2	4

Table 3: Study of Mechanism of Inflammation

Diagnosis	No. of patients (50)	Percentage (%)
Pemphigus vulgaris	18	36
Bullous pemphigoid	10	20
Spongiotic Dermatitis	4	8
Pemphigus Foliaceous	2	4
Sub Corneal pustular Dermatitis	2	4
Lichen planus pemphigoides	2	4

Drug Reaction	1	2
Darier's disease	1	2
Hailey Hailey	1	2
Inconclusive / Description	4	8
Other single cases	5	10

Table 4: Study of Immune fluorescence Diagnosis

Innume fluorescence Diagnosis	No. of Patients	Percentage
Pemphigus	14	28
Bulbous pemphigoid	8	16
Bullous pemphigoid/Epidermolysis Bullosa Acquisita	1	2
Chronic Bullous Dermatitis of childhood	1	2
Linear IgA Bullous Dermatitis	1	2
Epidermolysis Bullosa Acquisita	1	2
Lichen planus pemphigoides	1	2
Bullous Lupus Strythematosus	1	2
Lupud Band	1	2
No Features of immune bulbous Disease	2	4
Total number of patients	31	62

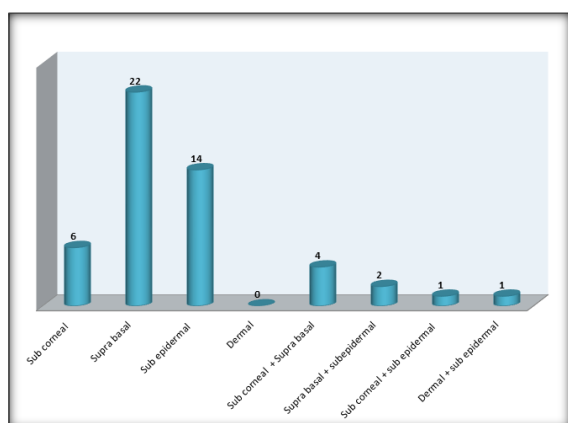


Figure 1: Study of Blister Separation plane

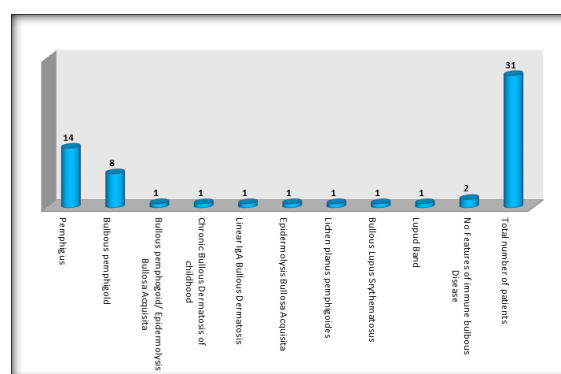


Figure 4: Study of Immune fluorescence Diagnosis

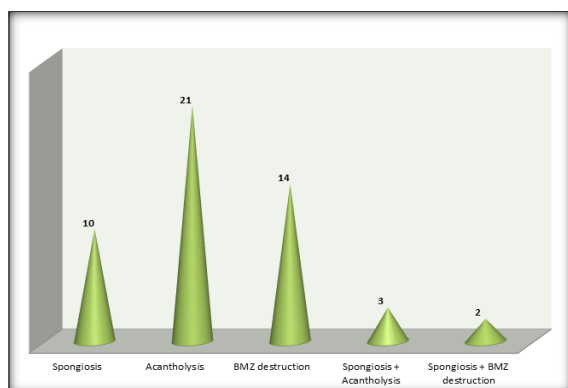


Figure 2: Inflammatory infiltrate

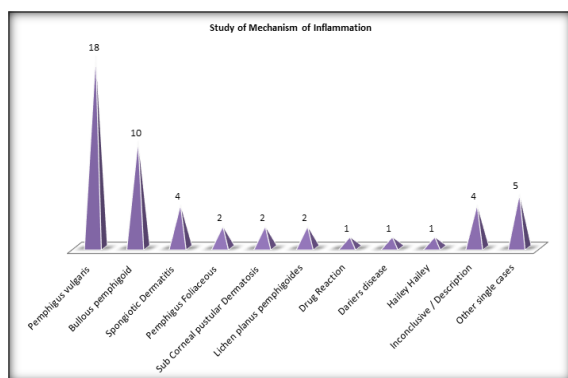


Figure3: Study of Mechanism of Inflammation

DISCUSSION

Present study of vesicobulbous disorders. The blister separation plane had the highest at 22 (44%), followed by 14 (28%) subepidermal and 1 (2%) sub-corneal + subepidermal, and dermal + subepidermal (Table 1). In the study of inflammatory infiltrate characters, the highest was 21 (42%), followed by 14 (28%) BMZ destruction, and the least was 2 (4%) spongiosis + BMZ destruction (Table 2). The diagnosis of the mechanism of inflammation had 18 (36%) pemphigus vulgaris, followed by 10 (20%) bullous pemphigoid, and the least number, 1 (2%), were Darier's disease, Hailey-Hailey, and drug reaction (Table 3). The study of immunofluorescence diagnosis had the highest number, which was 14 (28%) pemphigus, followed by 8 (16%) bulous pemphigoid (Table 4). These findings are more or less in agreement with previous studies.^[5,6,7]

Skin is the largest organ of the body. It represents a window to the internal well-being. Various diseases, along with their manifestations, can commonly involve the skin and mucous membranes, out of which vesiculobullous lesions form the predominant group.^[8] These bullous diseases in some instances are fatal if untreated. Blisters occur at different levels within the skin in the various disorders, and histological assessment is essential for accurate

diagnosis, providing insight into the pathogenic mechanism.

Apart from its protective function, skin has an immune function also, protecting from non-self antigens without reacting with self antigens. Sometimes this is distorted and leads to autoimmune disorders. Vesico-bulbous lesions are mostly immune-mediated, and the immune pathogenesis is specific for each disease, which is of diagnostic importance. The immune complex locations are intraepidermal, dermoepidermal junction, dermal blood vessels, etc.^[9] The nature of immune deposits usually used in DIF is IgG, IgA, IgM, and C3c.^[10]

Significant insights into the regulations of desmosomal adhesion of skin demonstrated the role of autoantibodies in patients suffering from the autoimmune blistering skin diseases.^[11] Autoantibodies against desmoglein cause the intraepidermal blistering seen in pemphigus, while autoantibodies against hemidesmosomes are the reason for subepidermal blistering seen in pemphigoid.^[12] The clinical examination aided by light microscopy and immunofluorescence in definitive diagnosis and proper patient management.

CONCLUSION

The present study of vesico-bulbous disorders represents a heterogeneous group of dermatoses with protean manifestations. A punch biopsy of the skin is a simple, inexpensive, safe OPD procedure causing minimal discomfort to the patient. DIF is always a gold standard and sensitive test to finalize the diagnosis. DIF aids differentiate non-immune lesions from immune-mediated lesions, which pose a diagnostic dilemma both clinically and histologically.

Limitation of study: Owing to tertiary location of research centre, small number of patients lack of latest techniques we have limited finding and results.

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