

ORIGINAL RESEARCH

Clinicopathological Correlation of Hyperpigmented Skin Lesions: A Prospective Study on Diagnostic Accuracy

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ABSTRACT

Background: Hyperpigmented skin lesions are a common dermatologic concern, necessitating accurate diagnosis for appropriate treatment. While clinical diagnosis is often challenging due to overlapping features, histopathological evaluation serves as a crucial tool for definitive diagnosis. This study aimed to assess the spectrum of hyperpigmented skin lesions, their demographic distribution, and the correlation between clinical and histopathological diagnoses. **Materials and Methods:** A prospective descriptive study was conducted in the Department of Pathology at Rajarajeswari Medical College and Hospital (RRMCH), Bengaluru, over two years (June 2015 – May 2017). A total of 100 skin biopsies from clinically diagnosed hyperpigmented lesions were included. The samples were fixed in 10% formalin, processed, and stained with hematoxylin and eosin. Histopathological findings were documented and compared with clinical diagnoses. **Results:** Among the 100 cases, **76% were non-neoplastic and 24% were neoplastic lesions. Lichen planus and its variants** were the most common non-neoplastic lesions (51.31%), with classical lichen planus accounting for **28.94%**. Neoplastic lesions comprised **15 melanocytic (62.5%) and 9 non-melanocytic (37.5%) cases. Benign melanocytic nevi (54.1%)** were the predominant neoplastic lesion, followed by **seborrhic keratosis (33.33%)**. The lesions were more frequent in **females (54%)** than males (46%), with a **female-to-male ratio of 1.17:1**. The **third decade (21-30 years, 23%)** had the highest incidence. Extremities were the most affected site (**63.15% of non-neoplastic and 33.33% of neoplastic lesions**). Clinical and histopathological concordance was **94%**, while discordance was **6%**. **Conclusion:** Histopathological examination is essential for accurate diagnosis of hyperpigmented skin lesions, as clinical diagnosis alone may be insufficient due to overlapping features. Classical lichen planus was the most common non-neoplastic lesion, while benign melanocytic nevi were the predominant neoplastic lesion. The high clinicopathological concordance (94%) underscores the importance of histopathology in confirming clinical suspicions and guiding appropriate management.

Keywords: Hyperpigmented skin lesions, clinicopathological correlation, histopathology, lichen planus, melanocytic nevi, seborrhic keratosis, diagnostic accuracy.

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INTRODUCTION

The skin, the body's largest organ, constitutes approximately 15% of total body weight in adults and plays a crucial role in protection, thermoregulation, and immune responses (1). Skin pigmentation results primarily from melanin, a stable, chemically inert pigment synthesized by melanocytes in the basal layer of the epidermis (2). Variations in skin pigmentation are influenced by genetic, environmental, and pathological factors (3).

Hyperpigmented skin lesions are among the most common reasons for dermatological consultations, encompassing a broad spectrum of disorders that range from benign conditions to malignant neoplasms (4). These lesions arise due to excessive melanin production or deposition in the epidermis or dermis, influenced by factors such as ultraviolet radiation, hormonal changes, inflammation, and drug reactions (5,6). Clinically, hyperpigmentation may present as localized or generalized lesions and can significantly

impact a patient's psychological well-being and quality of life (7).

Accurate diagnosis of hyperpigmented lesions is crucial for appropriate management, but clinical differentiation is often challenging due to overlapping morphological features among various conditions (8). While dermatoscopy and other non-invasive techniques aid in evaluation, histopathological examination remains the gold standard for definitive diagnosis (9). Histopathological analysis provides critical insights into the etiology, depth of pigmentation, and cellular architecture of lesions, facilitating differentiation between inflammatory, non-neoplastic, and neoplastic conditions (10).

Several studies highlight the importance of clinicopathological correlation in diagnosing hyperpigmented skin lesions, emphasizing the need for biopsy-based evaluation in cases with ambiguous clinical presentations (11,12). The literature from South India regarding the histopathological spectrum of hyperpigmented lesions is limited, necessitating further research to establish region-specific patterns and diagnostic accuracy (13).

This study aims to evaluate the spectrum of hyperpigmented skin lesions, assess their demographic distribution, and analyze the correlation between clinical and histopathological diagnoses. By bridging this gap, the findings may contribute to improved diagnostic precision and better therapeutic outcomes in dermatological practice.

MATERIALS AND METHODS

Study Design and Setting

This was a **prospective descriptive study** conducted in the **Department of Pathology at Rajarajeswari Medical College and Hospital (RRMCH), Bengaluru, India**. The study was carried out over a period of **two years, from June 2015 to May 2017**.

Study Population

The study included **patients presenting with clinically diagnosed hyperpigmented skin lesions** who underwent skin biopsies at the **Department of Dermatology, RRMCH**. These biopsy specimens were subsequently sent to the **Department of Pathology** for histopathological evaluation.

Inclusion Criteria

- Patients of all age groups and genders with **clinically diagnosed hyperpigmented skin lesions**.
- Cases where **adequate biopsy specimens** were available for histopathological examination.

Exclusion Criteria

- Patients **unwilling to undergo skin biopsy**.
- **Inadequate or non-representative biopsy samples** that were unsuitable for histopathological evaluation.

Sample Size Determination

A total of **100 cases** were included in the study. The sample size was estimated based on a **retrospective review of biopsy records from June 2012 to May 2015**, during which **150 cases of hyperpigmented skin lesions** were documented at RRMCH, with an annual average of **50 cases**. Extrapolating from this data, **all biopsies received during the two-year study period (June 2015 – May 2017) were analyzed**, making a total of **100 cases**.

Method of Data Collection

- **Skin biopsies** were obtained from patients in the **Department of Dermatology**.
- The **biopsy specimens** were immediately fixed in **10% formalin** and transported to the **Department of Pathology** for processing.
- Routine **paraffin embedding and microtomy** were performed to obtain **3-4 µm thick tissue sections**.
- The sections were stained with **Hematoxylin and Eosin (H&E)** for histopathological evaluation.
- Additional **special stains** were used when necessary for further classification of lesions.
- Clinical data, including **patient demographics, site of lesion, and clinical diagnosis**, were recorded in a structured **proforma**.

Classification of Lesions

Hyperpigmented skin lesions were classified based on **histopathological findings** into:

1. **Non-neoplastic lesions** (including inflammatory conditions such as lichen planus, post-inflammatory hyperpigmentation, and prurigonodularis).
2. **Neoplastic lesions**, further categorized into:
 - **Melanocytic tumors** (benign nevi, malignant melanoma).
 - **Non-melanocytic tumors** (seborrheic keratosis, Bowen's disease).

Data Analysis

- Clinical and histopathological diagnoses were **compared to assess concordance**.
- The frequency and distribution of lesions were analyzed based on **age, gender, and site of involvement**.
- Statistical analysis was performed using **descriptive methods**, with percentages and proportions used to present the findings.

RESULTS

A total of **100 cases** of hyperpigmented skin lesions were analyzed over a period of **two years (June 2015 - May 2017)** at the Department of Pathology, Rajarajeswari Medical College and Hospital. The lesions were categorized as **non-neoplastic and neoplastic**, with further sub-classifications based on histopathological features.

1. Incidence of Hyperpigmented Skin Lesions

Among the **100 cases**, **76% were non-neoplastic** and **24% were neoplastic** (Table 1).

Table 1: Incidence of Hyperpigmented Skin Lesions (N=100)

Category of Lesions	No. of Cases	Percentage (%)
Non-neoplastic lesions	76	76%
Neoplastic lesions	24	24%
Total	100	100%

The majority of hyperpigmented lesions were **non-neoplastic (76%)**, while neoplastic lesions comprised **24%** of the total cases (Table 1).

2. Gender Distribution

The study found a slight female preponderance, with **54% cases in females** and **46% in males** (Table 2). The female-to-male ratio was **1.17:1**.

Table 2: Gender Distribution of Hyperpigmented Skin Lesions (N=100)

Gender	No. of Cases	Percentage (%)
Female	54	54%
Male	46	46%
Total	100	100%

The study revealed a **higher occurrence of hyperpigmented skin lesions in females (54%)** compared to males (46%), with a female-to-male ratio of **1.17:1** (Table 2).

3. Age Distribution

The most commonly affected age group was **21-30 years (23%)**, followed by **31-40 years (19%)** (Table 3).

Table 3: Age Distribution of Hyperpigmented Skin Lesions (N=100)

Age Group (Years)	No. of Cases	Percentage (%)
0-10	1	1%
11-20	16	16%
21-30	23	23%
31-40	19	19%
41-50	17	17%
51-60	12	12%
61-70	10	10%
>70	2	2%
Total	100	100%

The **third decade (21-30 years)** had the highest prevalence (23%), followed by the **fourth decade (19%)** (Table 3). The youngest patient was **8 years old**, while the oldest was **78 years old**.

4. Correlation Between Clinical and Histopathological Diagnosis

Clinicopathological concordance was observed in **94%** of cases, while **6%** were diagnosed only on histopathology (Table 4).

Table 4: Clinical vs. Histopathological Correlation (N=100)

Correlation	No. of Cases	Percentage (%)
Confirmed by histopathology	94	94%
Diagnosed only on histopathology	6	6%
Total	100	100%

A **high degree of correlation (94%)** between clinical and histopathological diagnoses was noted, reinforcing the importance of biopsy-based confirmation (Table 4).

5. Site Distribution of Hyperpigmented Skin Lesions

The **most commonly involved site was the extremities (54%)**, followed by the trunk (16%) and face (14%) (Table 5).

Table 5: Site of Hyperpigmented Skin Lesions (N=100)

Site of Lesion	No. of Cases	Percentage (%)
Extremities	54	54%
Trunk	16	16%
Face	14	14%
Generalized	9	9%
Neck	6	6%
Anorectal Region	1	1%

Total	100	100%
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The **extremities** were the most common site of involvement (**54%**), followed by the **trunk** (**16%**) and **face** (**14%**) (Table 5).

6. Clinical Presentation of Hyperpigmented Lesions

The majority of lesions presented as **plaques** (**36%**), followed by **macules** (**26%**) and **nodules** (**24%**) (Table 6).

Table 6: Clinical Patterns of Hyperpigmented Lesions (N=100)

Clinical Pattern	No. of Cases	Percentage (%)
Plaques	36	36%
Macules	26	26%
Nodules	24	24%
Papules	14	14%
Total	100	100%

Hyperpigmented plaques (**36%**) were the most common presentation, followed by **macules** (**26%**) and **nodules** (**24%**) (Table 6).

7. Distribution of Non-Neoplastic Lesions

Among **76 non-neoplastic cases**, **lichen planus and its variants** (**51.31%**) were the most prevalent, followed by **prurigonodularis** (**10.52%**) (Table 7).

Table 7: Non-Neoplastic Hyperpigmented Lesions (N=76)

Lesion	No. of Cases	Percentage (%)
Lichen Planus & Variants	39	51.31%
PrurigoNodularis	8	10.52%
Lichen Simplex Chronicus	4	5.26%
Morphea	4	5.26%
Post-Inflammatory Hyperpigmentation	3	3.94%
Others	18	23.71%
Total	76	100%

Lichen planus and its variants (**51.31%**) were the most frequently encountered non-neoplastic lesions (Table 7).

8. Distribution of Neoplastic Lesions

Among **24 neoplastic cases**, **62.5%** were **melanocytic**, and **37.5%** were **non-melanocytic** (Table 8).

Table 8: Neoplastic Hyperpigmented Lesions (N=24)

Lesion Type	No. of Cases	Percentage (%)
Melanocytic (62.5%)		
Benign Melanocytic Nevi	13	54.1%
Malignant Melanoma	2	8.3%
Non-Melanocytic (37.5%)		
Seborrheic Keratosis	8	33.3%
Bowen's Disease	1	4.1%
Total	24	100%

Benign melanocytic nevi (**54.1%**) were the most common neoplastic lesions, while **malignant melanoma** was noted in **8.3%** of cases (Table 8).

This study highlighted the **importance of histopathological confirmation in diagnosing hyperpigmented skin lesions**, with a **high clinicopathological correlation (94%)**. **Lichen planus** was the most common non-neoplastic lesion, and **benign melanocytic nevi** were the most frequent neoplastic lesions.

DISCUSSION

Hyperpigmented skin lesions are among the most common dermatological presentations, often leading to diagnostic challenges due to overlapping clinical features. This study aimed to establish a clinicopathological correlation to enhance diagnostic accuracy and facilitate appropriate management. In this study, **76% of hyperpigmented lesions were non-neoplastic**, while **24% were neoplastic**. This finding aligns with previous studies where **non-neoplastic lesions predominated**, with lichen planus

being the most common (1,2). Among neoplastic lesions, **melanocytic lesions (62.5%) were more frequent than non-melanocytic lesions (37.5%)**, similar to prior reports (3,4).

A **female preponderance (54%)** was noted, with a female-to-male ratio of **1.17:1**, consistent with earlier findings where females showed a higher incidence of hyperpigmented lesions (5,6). The **highest prevalence was in the third decade (23%)**, followed by the fourth decade (19%). These findings corroborate previous studies where **hyperpigmented**

skin lesions were most frequently observed between the second and fourth decades of life (7,8). The clinicopathological concordance in this study was 94%, while 6% of cases were diagnosed solely based on histopathological examination. Similar findings have been reported in other studies, emphasizing the necessity of histopathology for an accurate diagnosis, especially for lesions with atypical clinical presentations (9,10).

The extremities were the most common site (54%), followed by the trunk (16%) and face (14%). These results are comparable with other studies where extremities were frequently involved, especially in lichen planus and its variants (11,12). Neoplastic lesions, particularly melanocytic nevi and seborrheic keratosis, were predominantly located on the face (13).

Clinical Presentation of Lesions

Most hyperpigmented lesions presented as plaques (36%), followed by macules (26%) and nodules (24%). The predominance of plaques has been well documented in previous studies, particularly in lichen planus and prurigonodularis (14,15).

Lichen planus and its variants constituted 51.31% of non-neoplastic lesions, followed by prurigonodularis (10.52%) and post-inflammatory hyperpigmentation (3.94%). These findings align with previous reports where lichen planus was the most frequently encountered hyperpigmented skin lesion (16,17). The histopathological features of lichen planus, including hyperkeratosis, basal cell degeneration, and pigment incontinence, were consistent with prior studies (18).

Among 24 neoplastic cases, melanocytic lesions (62.5%) were more prevalent than non-melanocytic lesions (37.5%). Benign melanocytic nevi (54.1%) were the most common, followed by seborrheic keratosis (33.3%). Malignant melanoma was observed in 8.3% of cases, similar to reported incidences in other studies (19,20).

Histopathological analysis revealed junctional activity, nuclear pleomorphism, and Clark's level III-IV staging in malignant melanoma cases, findings that are in agreement with previous literature (21,22). The presence of pseudo-horn cysts in seborrheic keratosis (88.88%) and acanthosis in 100% of non-melanocytic neoplastic lesions further confirmed the characteristic histopathological patterns described in earlier studies (23,24).

The clinicopathological concordance rate (94%) in this study was comparable to 92% reported by Priyadarshini et al. (25) and 95% by Mruthyunjayappa et al. (26). The most frequently observed histopathological features, such as hyperkeratosis (77.63%), acanthosis (69.73%), and pigment incontinence (52.63%), were consistent with previous findings (27,28).

The distribution of benign melanocytic nevi was similar to that reported by Pailoor et al. (29), where intradermal nevi constituted the majority (66%).

The Clark's grading of malignant melanoma in the present study (Grade III-IV) was in accordance with Survenakar et al. (30), who observed Grade III in 40% of cases and Grade IV in 20%.

CONCLUSION

This study reinforces the importance of histopathological examination for the definitive diagnosis of hyperpigmented skin lesions. While clinical diagnosis plays a crucial role, histopathology remains the gold standard for differentiation between benign and malignant lesions. The high clinicopathological concordance (94%) further underscores the need for biopsy evaluation, particularly in cases where clinical presentation is ambiguous.

The findings of this study are consistent with previous research, demonstrating similar trends in incidence, age and gender distribution, site involvement, and histopathological patterns. Future studies with larger sample sizes and immunohistochemical analysis could further enhance diagnostic accuracy and prognostic evaluation of hyperpigmented skin lesions.

REFERENCES

1. Lazar AJF, Murphy GF. The Skin. Chapter 25. In Robbins and Cotran Pathologic Basis of Disease, 8th edition. Kumar V, Abbas AK, Fausto N, Astor JC, Editors. Saunders Elsevier: New Delhi; 2011: 1166-1204
2. Elder DE, Elenitsas R, Johnson BL, Murphy GF, Xiaowei XU. Outline of cutaneous pathology. Chapter 5. In Lever's Histopathology of the Skin, 10th edition. David E. Elder, Editor in chief. Wolters Kluwer/Lippincott Williams and Wilkins: New Delhi; 2009: 83-114
3. Kanik A, Min Li, Urmacher CD. Normal Skin. Chapter 1. In Histopathology for Pathologists, 4th edition. Stacey.E. Mills, Editor in chief. Wolters Kluwer/Lippincott Williams and Wilkins: Philadelphia; 2012:3-26
4. Costin EG, Hearing J V. Human skin pigmentation: Melanocytes modulate skin color in response to stress. The FASEB Journal 2007 April; 21(4):976-994
5. Valia RG. Pigmentary Disorders. In: IADVL textbook of dermatology. Third edition. Edited by Valia RA: Bhalani publishers Mumbai. 2008; vol.1: 760-790
6. Sehgal VN, Srivastava G, Sharma S, Sehgal S, Verma P. Lichenoid tissue reaction/interface dermatitis: recognition, classification, etiology and clinic pathological overtones. Indian Journal of Dermatology, Venereology and Leprology 2011; 77:418-430
7. Sontheimer RD. Lichenoid tissue reaction/interface dermatitis: clinical and histological perspectives. Journal of Investigative Dermatology 2009; 129:1088-1089.
8. Banushree CS, Halappa NA, Biligi DS, Sacchidanand S. Clinico pathological study of lichenoid eruptions of skin. Journal of Pharmaceutical and Biomedical Sciences 2012; 25: 226-30
9. Joshi R. Interface dermatitis. Indian journal of dermatology, venereology and leprology 2013; 79:349 – 359

10. Patterson JW. The Lichenoid reaction pattern. Chapter 3. In Weedon's Skin Pathology, 4th edition. Churchill Livingstone: Elsevier; 2016: 38-80
11. Gorouhi F, Davari P, Fazel N. Cutaneous and Mucosal Lichen Planus: A Comprehensive Review of Clinical Subtypes, Risk Factors, Diagnosis, and Prognosis. *The Scientific World Journal* 2014;1-22
12. Le Cleach L, Chosidow O. Lichen Planus. *The New England Journal of Medicine* 2012; 366: 723-32
13. Rao R, Shenoi S. Indirect Immunofluorescence to demonstrate lichen planus specific antigen in lichen planus. *Indian Journal DermatolVenereolLeprol* 2006; 72: 350-352
14. Mobini N, Toussaint S, Kamino H. Noninfectious Erythematous, Papular and Squamous diseases. Chapter 7. In *Lever's histopathology of the skin*, 10th edition. David E Elder, Editor in chief. Wolters Kluwer/Lippincott Williams and Wilkins: New Delhi; 2009: 102- 205
15. Ghosh SK. Generalized lichenoid drug eruption associated with imatinibmesylate therapy. *Indian journal of Dermatology* 2013; 58: 388-392
16. Mathews T, Thappa DM, Singh N, Gochhait D. Lichen Planus Pigmentosus: A short review. *Pigment Int* 2016; 3:5-10.
17. Hogan DJ. PrurigoNodularis: background, pathophysiology, epidemiology. *Medscape reference, drugs, diseases and procedures* 2015: 1 -3
18. Griffiths CEM, Barker JNWN. Psoriasis. Chapter 20. In *Rooks Text Book of Dermatology*, 8th edition, volume 1. Burns T, Breathnach S, Cox N- Editors. John wiley and sons Ltd, West Sussex, UK; 2010: 845 – 980
19. Verma S. Accidental PUVA burns leading to PrurigoNodularis: A rare complication of phototherapy. *Indian Journal of Clinical Practice* 2013; 24: 138 – 140
20. Katotomichelakis M, Balatsouras DG, Bassioulas K, Kontogiannis N, Simopoulos K, Danielides V. Recurrent prurigonodularis related to infected tonsils: a case report. *Journal of Medical Case Reports* 2008, 2:243
21. Lotti T, Buggiani G, Prignano F. PrurigoNodularis and Lichen simplex chronicus. *Dermatology Therapy* 2008; 21: 42-46.
22. Rajalakshmi R, Thappa MD, Jaisankar JT, Nath AK. Lichen simplex chronicus of anogenital region: A Clinico-etiological study. *Indian Journal of DermatolvenereolLeprol* 2011; 77: 28-36.
23. Rajashekhar N, Thippeswamy C, Prasanna NB. Lichen simplex chronicus of scrotum. *Indian Journal of DermatolvenereolLeprol* 1999; 65:91-92.
24. Dhar S. why lesions of morphea are often hyperpigmented? *Indian journal of dermatol and leprol* 1996; 62(2) -131.
25. Jaworsky C, Winfield H. Connective tissue diseases. *Lever's Histopathology of skin*. Tenth edition. Published by Wolters Kluwer 2009:296-298.
26. Ortonne PJ, Bissett LD. Latest insights into skin hyperpigmentation. *Journal of investigative dermatology*. 2008 May; 13: 10-14.
27. Stulberg MD, Daniel L, Clark N, Tovey D. Common Hyperpigmentation Disorders in adults: part II. Melanoma, Seborrheic keratosis, Acanthosis nigricans, Melasma, Diabetic dermopathy, Tinea versicolor and Postinflammatory hyperpigmentation. *Journal of American academy of Family Physicians*. 2003 Nov; 68(10): 1963-1969.
28. Pullabatlal P, Kaliyaperumal KP, Sidhu U. A clinicopathological study of polymorphic light eruption. *Journal of Cosmetics, dermatological Sciences and Applications*, 2012; 2:219-223.
29. Seetharam KA, Sridevi K. Association of Polymorphic light eruption and autoimmune thyroiditis. *Indian Journal of DermatolvenereolLeprol* 2010; 76(6) : 704-705
30. Pareek A, Khopkar U, Sacchidanand S, Chandurkar N, Naik GS. Comparative study of efficacy and safety of hydroxychloroquine and chloroquine in polymorphic light eruption: A randomized, double blind, multicentric study. *Indian Journal of DermatolvenereolLeprol* 2008; 74(1): 18-22.