



A Study of Epidemiological and Clinicopathological Correlation of Facial Hyperpigmentation in Females

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ABSTRACT: Introduction: Pigmentary disorders comprise almost one third of the dermatological consultations. The incidence, frequency and severity of pigmentary disorders vary across different geographic regions. They are particularly worse in warmer areas and in conditions where skin becomes more exposed. The pattern of reporting hyperpigmentation is age dependent, with most common complaint amongst those aged 40 to 54 years and second most in age group 15 to 30 years. Material and method: A total number of 150 female patients above 18yr of age with facial hyperpigmentation attending dermatology outpatient department of Muzaffarnagar medical college, Muzaffarnagar, U.P during 2017-2018. Woods lamp examination was conducted to determine the type of Melasma (Epidermal, Dermal or Mixed) clinically. Skin biopsy was performed to all patients at the first visit and histopathological results were compared with the clinical diagnosis. With informed consent, 3-4mm elliptical biopsy from the lesional skin of patient was taken and tissue was prepared for light microscopic examination by fixation in 10% formalin and stained with Hematoxylin & Eosin. Further, the section was preceded for special staining i.e., Masson Fontana, Congo red, KOH/PAS to categorize histopathologically and were correlated with the clinical entity. Result: On the basis of clinicodemographic profile, melasma is the most common diagnosis (52%) followed by Reihl's melanosis (21.3%), ochronosis (10%), TSDF (6.6%), ashy dermatosis (3.3%), PDL (2%), Argiria, amyloidosis and DLE (1.3% each) and Poikiloderma of civatte (0.6%). Histopathologically, a total of nine entities were identified as: melasma (80 patients; 53.3%) followed by Reihl's melanosis (40 patients; 26.6%), PIH (12 patients; 8%), ochronosis (6 patients; 4%), TSDF (5 patients; 3.3%), Ashy dermatosis and DLE (3 patients each; 2% each) and colloid milium (1 patient; 0.6%). Appropriate

statistics were applied and results indicated the concordance of the clinical assessment. Conclusion: An agreement of 65% was observed between histopathological findings and clinical diagnosis with respect to various presentations of facial pigmentation.

KEYWORDS: Pigmentary disorders, hyperpigmentation, melasma, ochronosis, TSDF

I. INTRODUCTION

Variability in pigmentation is well-established, with some skin tones, especially in Asian and Indian subjects, reported to be more susceptible to pigmentation disorders than other human groups^{[1],[2]}. Pigmentary disorders comprise almost one third of the dermatological consultations. The incidence, frequency and severity of pigmentary disorders vary across different geographic regions. They are particularly worse in warmer areas and in conditions where skin becomes more exposed. The pattern of reporting hyperpigmentation is most common complaint amongst those aged 15 to 54, regardless of skin colour and gender.³ Hyperpigmentation can be caused by a variety of diseases which may be congenital, with different patterns of inheritance, or acquired forms secondary to cutaneous (cosmetics, perfumes, sun exposure) or systemic problems (mainly hormonal) and often represent paramount causes of emotional distress. The vast majority of them are linked to alterations in the melanin pigment and may be classified as epidermal and dermal hyperpigmentation, depending on the location of the pigments. Epidermal hyperpigmentation is because of melanin pigmentation. Dermal pigmentation is may either be due to melanin or due to non-melanin pigments. Hyperpigmentation's are a group of diseases that comprise both congenital forms, with different patterns of inheritance, and acquired forms secondary



cutaneous or systemic problems. Pigmentary disorders are a frequent source of complaints, constituting the third most common reason for dermatological consultations, about 8.5% in our country. There is a different impact depending on the geographic region, being worse in places where the weather is always warm, and the skin becomes more exposed. This review will focus on the main acquired hyperpigmentation disorders associated with increased melanin, taking into account those most commonly found in clinical practice. Melasma most common in people with light brown skins, especially in people who live in areas with intense solar ultraviolet radiation (UVR)⁴. Most FM are commoner in darker races with both light and photosensitizing chemicals (occupational/ in cosmetics) playing an important

role. Based on the location of melanin (as identified by color of lesion, accentuation of color under Wood's light and histopathology, though the correlation between Wood's lamp findings and histopathology is less than satisfactory) three types of hypermelanosis are identified:

The most common dermatological causes of acquired hyperpigmentation can be described as:⁵

1. Erythema dyschromicum perstans
2. Lichen planus pigmentosus
3. Amyloid melanosis
4. Melasma (cholasma)
5. Riehl's melanosis
6. Idiopathic eruptive pigmentation
7. Post-inflammatory hyperpigmentation, and
8. Hyperpigmentation due to drugs and heavy metals.

S.NO	ENTITY	HISTOPATHOLOGICAL DIAGNOSIS
1	Acanthosis nigricans	Hyperpigmentation of basal and suprabasal layers, hyperkeratosis and papillomatosis.
2	Exogenous Onychosis	Brown pigment deposits in shape of a 'banana' are found lying freely in dermis in between collagen bundles.
3	Lichen planus pigmentosus	Vacuolar interface dermatitis with a scarce lymphohistiocytic infiltrate, fibroplasia and pigmentary incontinence in papillary dermis.
4	Erythema dyschromicum perstans	Lichenoid infiltrate without fibroplasias
5	Riehl's melanosis	Basal cell degeneration and perivascular lymphocytic infiltrate in dermis
6	Poikiloderma of Civatte	Coarse collagen in the upper dermis, perivascular infiltrate of lymphocytes, histiocytes and few neutrophils.
7	Becker's melanosis	Increase melanin deposition in basal layer, acanthosis with squarish rete ridges, papillomatosis and thickening of dermis.
8	Café au lait macules	Pigmentation of basal layer
9	Macular amyloidosis	Amyloid deposits in dermal papillae close to dermoepidermal junction.
10	Frictional melanosis	Mild hyperkeratosis, acanthosis, hyperpigmentation in basal layer.
11	Pigmentary demarcation lines	Non-specific increase pigmentation in basal layer. No evidence of interface dermatitis or amyloid deposition



12	Post inflammatory hyperpigmentation	A basket-weave horn Papillary dermis thickened by fibrosis A sparse superficial perivascular lymphohistiocytic infiltrate Many melanophages in the papillary dermis
13	Nevi of Ota and Ito extending deep into the subcutis	Prominent sebaceous lobules of the facial skin Many slender dendritic pigmented melanocytes arranged horizontally between collagen bundles in the mid dermis Few or no Melanophages Melanocytic proliferation
14	Lupus erythematosus	Epidermis-hyperkeratosis, atrophy, interface dermatitis, thickening of basement membrane. Dermal-dense mononuclear cell infiltrate mainly lymphocytes and macrophages.
15	Lentigo simplex	A basket-weave cornified layer Slender elongated rete ridges. Melanophages within the papillary dermis
16	Berloque dermatitis	Increased epidermal melanin and Melanophages in the superficial dermis
17	Melasma	Increased epidermal hyperpigmentation Increased number of melanocytes Increased activity of melanogenic enzymes

II. MATERIAL AND METHODS

A total number of 150 female patients >18 years of age were taken under the study presenting with facial hypermelanosis attending outpatient department in the department of Dermatology, Venereology and Leprology at Muzaffarnagar medical college and hospital at a period of 18 months. A written consent was obtained from all patients after all details were explained to the patients. These include the duration of study, approval for skin biopsy, application and stoppage of any kind of treatment prior study. The study was preceded after taking approval from Institutional Ethical Committee. A full detailed history was obtained including the onset, patient age at beginning of pigmentation, duration, progression, associated symptoms, family history, residency, occupation and full physical examination including morphology, distribution and configuration. All skin biopsies of hyperpigmented lesion sent to the Department of Pathology, Muzaffarnagar Medical College, Muzaffarnagar, U.P

COLLECTION OF DATA

1. Onset
 2. Duration
 3. Progression
 4. Area involved
 5. Hours of sun exposure
 6. Occupation
 7. History of photosensitivity
 8. Other medication (oral/systemic)
 9. Topical medication (hair dye, fragrances, mustard oil, exposure to chemicals)
 10. Family history
 11. Pregnancy and lactation
 12. Onset during pregnancy
 13. History of use of oral contraceptives
 14. Obstetrical history
 15. History of acne, hirsutism, polycystic ovarian disease
- The following parameters would be taken into consideration.

INCLUSION CRITERIA

- Patient above 18 years of age
- Patient with aesthetic purpose.
- Patient with realistic expectation.



EXCLUSION CRITERIA

Patient with active herpes, chicken pox, viral exanthem infection.
Patient not willing for procedure.
Patient with keloidal tendency.
Patients with deranged clotting profile
Pregnancy
Lactation

INVESTIGATIONS: (when required)

Complete blood profile
Blood sugar
Lipid profile
Liver function test
Hormonal profile (LH, FSH values, testosterone, Serum DHEAS, serum cortisol)
Ultrasound abdomen to rule out polycystic ovarian disease
Vitamin B12
Thyroid hormones (T3, T4, TSH, Parathyroid hormone)

VISUAL ASSESSMENT

In both research and clinical settings, it is important to be able to accurately assess a subject's normal skin color or hue and their baseline degree of pigmentation so that any subsequent changes following treatment can be quantified.

PHOTOGRAPHY

All patients will be photographed and documented as per same position, angle of face exposure on day 1 of attending OPD. Minimum 3 photographs will be taken, one front view and 2 lateral view will be taken with appropriate numbering.

WOOD'S LAMP EXAMINATION

Wood's lamp was invented in 1903 by Robert W. Wood. It emits long-wave UV radiation (UVR) generated by Wood's filter which is opaque except to a band between 320 - 400 nm with peak at 365 nm.

Technique of Wood's lamp examination

The lamp was to be allowed to warm up for about 1 minute. The examination room was perfectly dark, with black occlusive shades. The light source was 4 to 5 inches from the lesion. The area was not washed before subjecting it for Wood's lamp examination since it may yield false negative results due to dilution of the pigment. Topical medicaments, lint and soap residues were wiped off from the site to be examined since these may fluoresce under Wood's light.

Hyperpigmentary dermatoses

Wood's lamp was used to determine the depth of melanin in the skin, as variations in epidermal pigmentation become more apparent under Wood's light. For dermal pigmentation, the contrast is less pronounced. Based on Wood's light findings, melasma is classified into four subtypes: epidermal, dermal, mixed and Wood's light inapparent. Wood's lamp examination was conducted to determine the type of Melasma (Epidermal, Dermal or Mixed). Examination of patients with Wood's light was useful in classifying the specific type of melasma in correlation with the localization of melanosomes in epidermis and dermis.

BIOPSY

Skin biopsy was performed to all patients at the first visit and histopathological results were compared with the clinical diagnosis. With informed consent, 3-4mm elliptical biopsy from the lesional skin of patient was taken and tissue was prepared for light microscopic examination by fixation in 10% formalin and stained with Hematoxylin & Eosin. Further, the section was preceded for special staining i.e., Masson Fontana, Congo red, KOH/PAS to categorize histopathologically and were correlated with the clinical entity. A skin biopsy is a simple procedure of great diagnostic importance and serves as permanent record of the skin pathology. In our study we considered excisional surgical biopsy method.

Steps

1. Consent and preparation
2. After taking the patient's consent, the biopsy area was cleaned with an antiseptic, prior trimming of the hair may be necessary if the region is hairy.
3. Local Anesthesia.
4. Use of 2% lignocaine with 1:1000 adrenaline is advisable for vascular area that are not supplied by terminal arteries.
5. Excisional elliptical biopsy 3-4mm skin were taken from involved area and entire lesion were removed, followed by suturing of the defect. Biopsy sent in 10% buffered formalin. Serial section were taken and stained with Hematoxylin & Eosin stain in all subjects.

Special stains were used wherever necessary which included: Masson Fontana, Congo red and KOH/PAS (Periodic acid Schiff)

III. OBSERVATIONS AND RESULTS

The study was done to analyze the association between clinical features of facial hyperpigmented



lesions and correlate with histopathological diagnosis in adult Indian females. For this purpose a total of 150 female patients between the ages of 18-55years were taken in sampling frame and enrolled after taking proper consent in the study.

All the patients underwent a thorough demographic evaluation, exposure to risk factors, clinical history and examination followed by a histopathological evaluation.

TABLE 1: AGE OF PRESENTATION

AGE (in years)	No. OF PATIENTS	PERCENTAGE (%)
<20	5	3.3
21-30	64	42.6
31-40	70	46.6
41-50	8	5.3
>50	3	2
Mean Age±SD (Range) in years	31.82 ± 6.81	

TABLE 2: LOCATION OF PIGMENTATION

Malar, cheek, nose, mandible, forehead	85	56.6
Between malar and temporal area	37	24.6
Lateral side of face	22	14.6
Angle of mouth to lateral side of chin	6	4

TABLE 3: COLOUR OF PIGMENTATION

COLOUR OF PIGMENTATION	NO. OF PATIENTS	PERCENTAGE (%)
Bluish grey	6	4
Brown	35	23.3
Dark brown	77	51.3
Grey brown	16	10.6
Light brown	12	8
Slate grey	4	2.6

TABLE 4: DEPTH OF PIGMENTATION ON WOODS LAMP

DEPTH OF PIGMENTATION ON WOODS LAMP IN MELASMA	NO. OF PATIENTS (OUT OF 78 PATIENTS)	PERCENTAGE (%)
Dermal	17	21.7
Epidermal	17	21.7
Mixed	44	56.4

TABLE 5: EXPOSURE TO RISK FACTORS

EXPOSURE TO RISK FACTORS	NO. OF PATIENTS	PERCENTAGE (%)
Daily sun exposure		
< 2 hours	54	36
3-8 hours	66	44
>8 hours	30	20
Duration of onset		



< 1 month	17	11.3
2-3 months	25	16.6
4-6 months	30	20.0
7-12 months	35	23.3
>12 months	43	
History of Photosensitivity	85	56.6
History of Systemic drugs	36	24
History of Topical drugs	127	84.6
Onset during pregnancy	7	4.6

TABLE 6: MOST COMMON OBSERVATIONS IN OUR STUDY

Most common observations in our study	Percentage of patients out of 100% (n=150)
1) Age group	31 - 40 (46.6%)
2) Occupation	Housewives (36.6%)
3) Duration of sun exposure	3 - 8 hrs (44%)
4) H/O photosensitivity	56.6%
5) H/O systemic drugs	24%
6) H/O topical drugs	31.3%
7) Duration of onset	> 12 months (28.6%)
8) Location	Malar area, forehead, nose, mandibular (56.6%)
9) Presentation	Bilateral (96.6%)
10) Pattern	Localised (56.6%)

Table 7: DISTRIBUTION OF PATIENTS ACCORDING TO THEIR CLINICAL DIAGNOSIS

CLINICAL DIAGNOSIS	NO. OF PATIENTS
Melasma	78
Reihl's melanosis	32
Ochronosis	15
TSDF	10
Lichen planus pigmentosus	5
DLE	2
Argyria	2
PDL	3
Amyloidosis	2
Poikiloderma of civatte	1

Table 8. DISTRIBUTION OF PATIENTS ACCORDING TO THEIR HISTOPATHOLOGICAL DIAGNOSIS

HISTOPATHOLOGICAL DIAGNOSIS	NO. OF PATIENTS
Melasma	73
Reihl's melanosis	29
Ochronosis	9
TSDF	18
Lichen plans pigmentosus	4
DLE	2
PDL	5
Amyloidosis	2
Poikiloderma of civette	1

Colloid milium	1
PIH	6

Data was analyzed using Statistical Package for Social Sciences, version 23.0. Chi-square test, kappa test, Independent sample t test were used to compare the data. A p value < 0.05 indicated a statistically significant association. Sensitivity was calculated by dividing true positive with a sum of true positive and false negative

multiplied by 100. Specificity was calculated by dividing true negative with a sum of true negative and false positive multiplied by 100. A p value less than 0.05 indicated a statistically significant association.

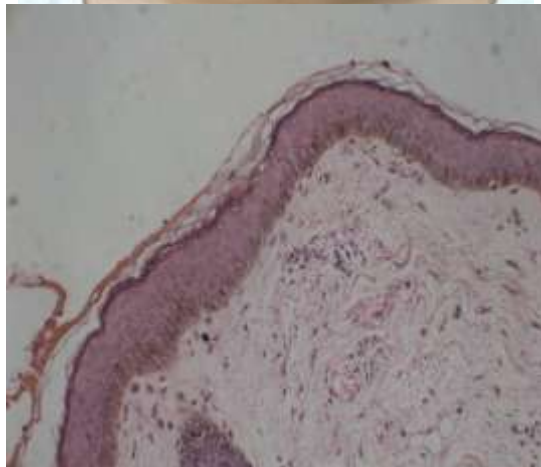


Figure 1a. Symmetric brown pigmented macules over malar, cheeks and nose in Melasma
Figure 1b. H&E shows increased atrophy, melanin deposition in basal layers and solar elastosis

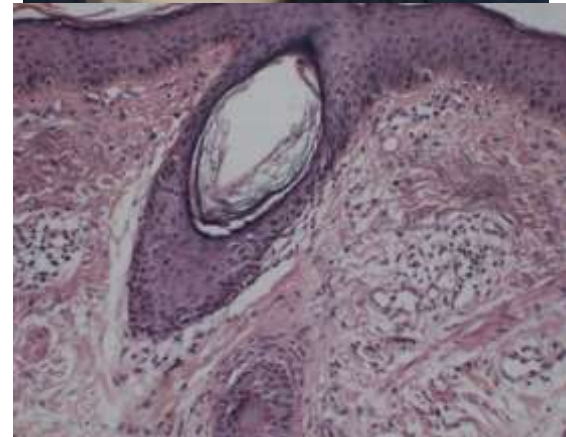


Figure 2a. Asymmetrical acquired hyperpigmentation bilaterally in Post Inflammatory Hyperpigmentation
Figure 2b. H&E showed increased amount of melanin in basal layer of epidermis and pigment incontinence



Figure 3a. black brownish hyperpigmentation with erythema over malar, cheeks, nose and around mouth in Pigmented Contact Dermatitis.

Figure 3b. H & E shows elongation of rete ridges present, pigment incontinence seen with melanin deposition at basal layers.

IV. CONCLUSION

The findings of the study showed that hyperpigmentation is an issue of concern especially among most of the females and they don't tend to report it early for treatment. Before consulting dermatologist they generally tend to use topical medication from a chemist without any prescription which effects the clinical presentation and makes it difficult to identify the disease clinically. Melasma and Reihl's Melanosis are commonly clinically overdiagnosed entities. A systematic approach for diagnosis which is the key to successful treatment and management, should be adopted and

histopathological correlation is essential before the active treatment planning is done.

REFERENCES

- [1]. Taylor SC, Cook-Bolden F, Rahman Z, Strachan D. Acne vulgaris in skin of color. *J Am Acad Dermatol* 2002;46 2 Suppl:S98-106
- [2]. Jablonski NG, Chaplin G. The evolution of human skin coloration. *J Hum Evol* 2000;39:57-106.
- [3]. Sbd.org CensoDermatologico da SBD. Brazilian Society of Dermatology Web page - Dermatological census. [Acesso 1 Dez. 2012]. [Internet] Disponível em: http://www.sbd.org.br/download/censo_dermatologico_2006.pdf2012
- [4]. Bandyopadhyay D. Topical treatment of melasma. *Indian J Dermatol* 2009;54:303-9
- [5]. Kang WH, Yoon KH, Lee ES, Kim J, Lee KB, Yim H, et al. Melasma: Histopathological characteristics in 56 Korean patients. *Br J Dermatol* 2002;146:228-37.