

# Morphometric Analysis to Identify the Perilesional Tissue in Oral Epithelial Dysplasia: A Novel Objective Tool

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## Abstract

Histopathological changes and molecular events that are hidden in otherwise clinically normal appearing mucosa may facilitate the detection of early changes in the surrounding mucosa, which can be assessed in perilesional tissue through morphometric analysis. Aim: To determine the perilesional tissue in oral mucosal biopsies through morphometric analysis. Material Methods: 50 formalin-fixed, paraffin-embedded tissue specimens were taken, which include 40 cases of oral epithelial dysplasia with perilesional tissue marked by India ink and 10 cases of the normal buccal mucosa. Histopathological examination was done to evaluate the presence or absence of dysplastic features in lesional and perilesional tissues. Morphometric analysis was done using epithelial thickness length between the apical membrane of basal cells the basement membrane. Results: There was no significant differences in the epithelial thickness between lesional perilesional tissue. However, the length between the apical membrane of basal cells the basement membrane was found to be significantly higher on the perilesional side in comparison to lesional side.

**Index terms**— morphometry, epithelial dysplasia, perilesional tissue, oral potentially malignant disorders (OPMDs).

## 1 Introduction

Oral carcinogenesis presents a multistep model of development, as it is unlikely that there is uniformity in the way individual patients or tissues behave. Although Oral squamous cell carcinoma (OSCC) is not linear in its development, it begins as simple epithelial hyperplasia progresses through epithelial dysplasia (OED), signifying more extensive genetic aberrations. Such predecessor lesions are often referred to as 'Oral potentially malignant disorders (OPMDs)' since not all lesions and conditions described under this term transform to cancer. The transition from normal oral epithelium to oral dysplasia and cancer results from accumulated genetic alterations. Underlying the histological transition of the normal oral epithelium through a precancerous state to invasive carcinoma are multiple molecular and cellular changes. Clinically OPMDs may present as leukoplakia, erythroplakia, oral lichen planus, and Oral Submucous Fibrosis. A variety of alterations accumulate to potentiate this transition to malignancy. Oral leukoplakia, harboring histologic features of hyperplasia and/or dysplasia is frequently encountered in the oral cavity. The rate of malignant transformation in OPMDs varies from 0% to 20% in 1 to 30 years. The malignant transformation risk of leukoplakia is associated with the lesional histology. The ability to identify oral leukoplakia patients at increased risk of cancer development is critical for improving control of oral cancer. Therefore, this study was designed to establish an objective tool to assess the perilesional tissue for histopathological changes in an otherwise clinically normal-appearing mucosa to reduce subjective bias and to objectively assess the predictive value of morphometric analysis in predicting dysplasia in otherwise normal mucosa through morphometric analysis.

II. The table shows cytological and architectural alterations between Lesional and Perilesional tissues of study cases. The results show that out of 40 cases of lesional and perilesional tissues of epithelial dysplasia-Loss of polarity of basal cells, Increased N:C ratio, Increased number, and size of nucleoli and Hyperchromasia showed marked difference which was found to be statistically significant ( $p < 0.05$ ). The above table shows

the morphometric analysis of epithelial thickness and basal cell diameter in lesional and perilesional tissue of study cases. The results show that out of the total 50 cases, 40 each were of lesional and perilesional tissues of epithelial dysplasia with 10 cases of the normal buccal mucosa. The epithelial thickness was  $1310.97 \pm 58.90$  in the control group,  $1734.22 \pm 520.97$  in the lesional group and  $1836.99 \pm 634.63$  in the perilesional group, which was statistically not significant ( $p>0.05$ ). The basal cell diameter in the control group was  $50.55 \pm 3.54$ , in the lesional group  $48.82 \pm 3.77$  and in the perilesional group  $52.03 \pm 4.87$ , which was statistically significant ( $p>0.05$ ).

## 2 Materials and Method

## 3 IV.

## 4 Discussion

The transition from normal oral epithelium to oral dysplasia and cancer results from accumulated genetic alterations. ?? Underlying the histological transition of the normal oral epithelium through a precancerous state to invasive carcinoma are multiple molecular and cellular changes. Clinically OPMDs may present as leukoplakia, erythroplakia, oral lichen planus, and Oral Submucous Fibrosis. A variety of alterations accumulate to potentiate this transition to malignancy. Oral leukoplakia, harboring histologic features of hyperplasia and/or dysplasia, is frequently encountered in the oral cavity. ?? The rate of malignant transformation in OPMDs varies from 0% to 20% in 1 to 30 years. The malignant transformation risk-of leukoplakia is associated with the lesional histology. ?? The early identification of oral leukoplakia patient is critical for improving the control of oral cancer.

## 5 Numerous

histological features which differentiates neoplastic tissue from the healthy tissues have been exhaustively discussed, but the findings remain subjective among observers. The WHO classification remains the gold standard for diagnosing OED, however, subjective bias was found, As the WHO classification includes vague histopathological factors, still there is a lack of evidence on how individual features should be weighted.

Objective analysis based on quantitative guidelines would be extra convenient and can be considered handy than a subjective framework. ??0 In the present-study, Histopathological examination was done to evaluate the presence or absence of dysplastic features in lesional and perilesional sides of 40 cases of Oral epithelial dysplasia. The marginal areas of the tissue were excluded from the analysis because they often show hyperchromatism artifacts. All cytological and architectural alterations established by WHO in the year 2017 for OED were evaluated. The results show that out of 40 cases of the lesional and the perilesional tissues of epithelial dysplasia- Loss of polarity of basal cells, Increased N:C ratio, Increased number, and size of nucleoli and Hyperchromasia showed marked difference which was statistically significant ( $p>0.05$ ) (Table 1)

In the present study, the morphometric analysis of epithelial thickness and basal cell diameter in the lesional and the perilesional regions of study cases was performed. The results show that the epithelial thickness was  $1310.97 \pm 58.90$  in the control group,  $1734.22 \pm 520.97$  in the lesional group and  $1836.99 \pm 634.63$  in the perilesional group, which was statistically non (Figure ??) The results of our study also reveal that there is an increase in epithelial thickness and basal cell diameter of perilesional tissue in comparison with lesional tissue. (Table 2) These findings are in accordance with the study done by Okamura et al. ( ??016) ?? where he demonstrated that epithelial thickness and basal cell diameter is a reliable indicator in determining and distinguishing the lesional and perilesional tissues. They further elucidated that the disordered arrangement of the basal cells is more useful than other candidate factors for determining the extent of lesional tissues. Further Nag et al. ( ??018) 11 discussed that morphometry could be administered discriminatorily on specimens having lesional and perilesional tissue, which is problematic to assess precisely through histopathology and postulated that Gutka and Pan Masala can lead to the emergence of high levels of reactive oxygen species (ROS) near the lesion leading to the damage of the surrounding normal tissue. Carcinogens decrease the capacity of the cytoplasm to mature and boost the nuclear parameters and cause a reduction in cellular parameters. Hence, they concluded that between normal and premalignant cells, cellular mean and cellular diameter was established to be lowest in premalignant cells. 9,11

## 6 Conclusion

An early and definitive diagnosis of epithelial dysplasia followed by complete excision of the lesion with safe margins is important to prevent the progression of Oral Epithelial Dysplasia into malignancy. Morphometric analysis can be considered as an effective guide to evaluate the progression of the normal epithelium to dysplastic epithelium in otherwise clinically normal mucosa. The results of the present study indicate morphometric analysis as an emerging objective tool to assist the routine histo-pathological diagnosis. <sup>1</sup>

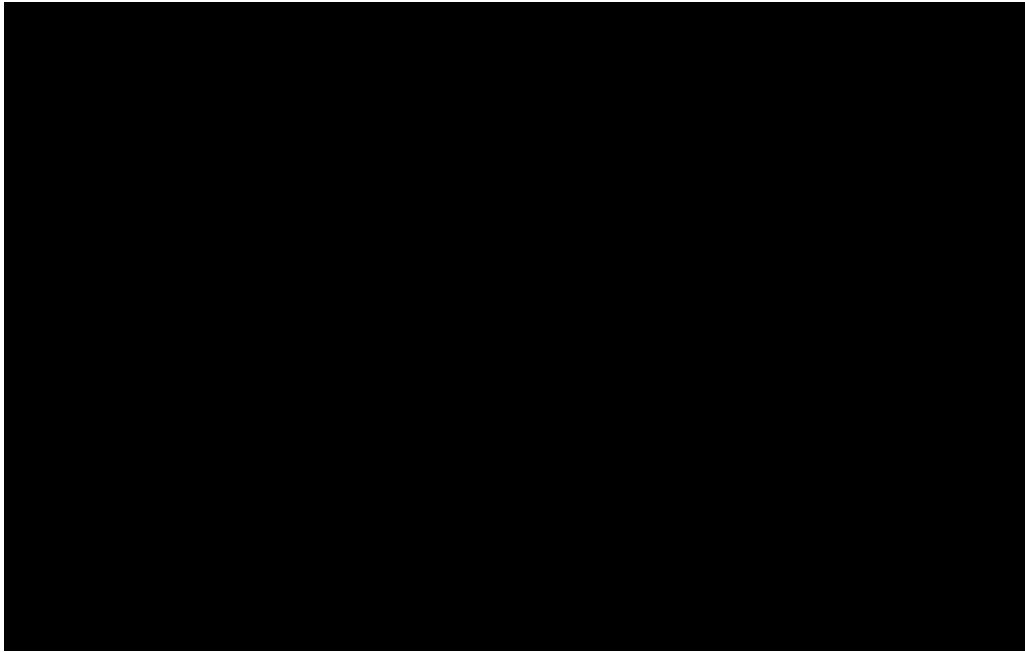


Figure 1:

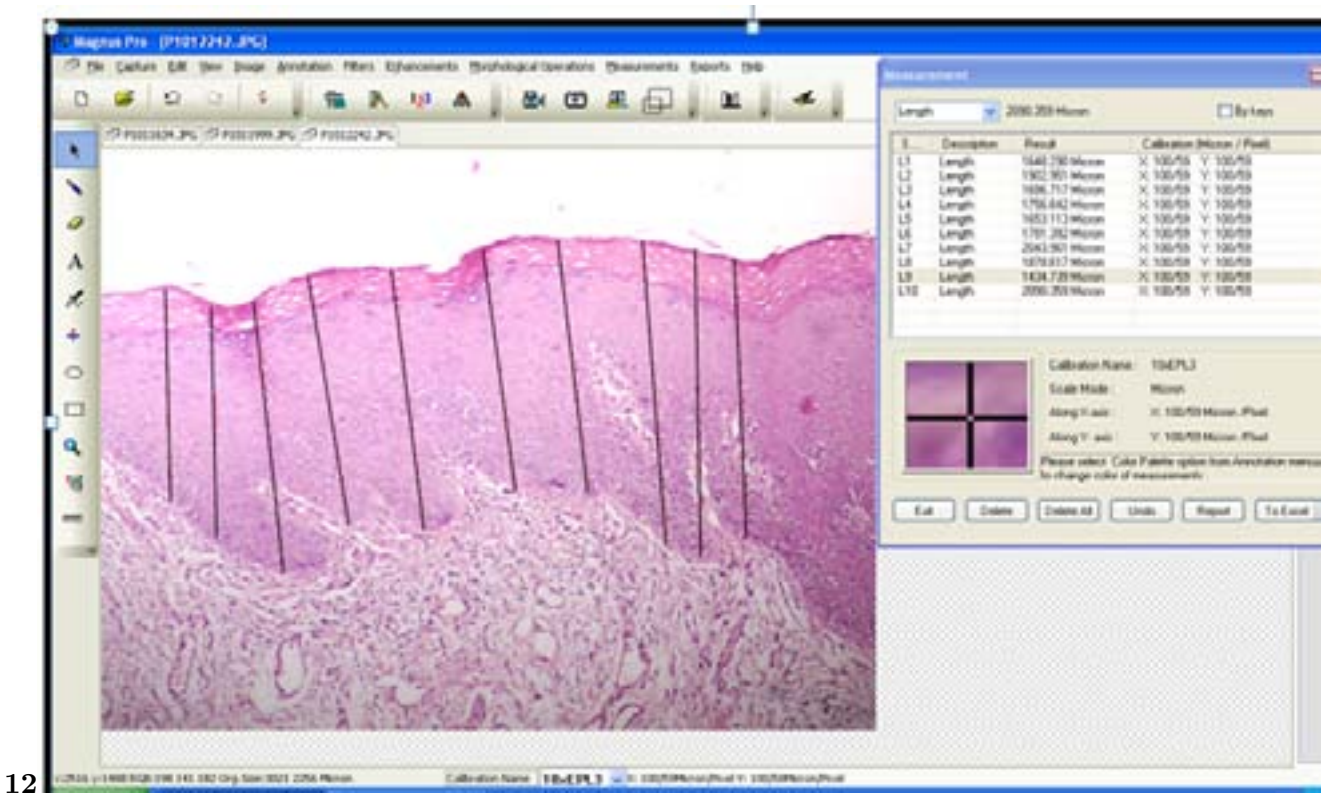


Figure 2: Fig. 1 :Fig. 2 :

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? 50 formalin- paraffin- tissue  
fixed, embedded  
specimen were taken, which include 40 cases of

[Note: 8 Following histopathological factors were evaluated in each case-in lesional and perilesional tissue Morphometric analysis was done using the image analysing software Magnus Pro. Two histopathopathological factors were quantified morphometrically in each case9 i. Epithelial thickness & ii. The length between the apical membrane of basal cells & the basement membrane. (Basal Cell Diameter) III. Results 2020 J Morphometric Analysis to Identify the Perilesional Tissue in Oral Epithelial Dysplasia: A Novel Objective Tool]

Figure 3: Table 1 :

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|                      | Groups       | N  | Mean± Std Deviation (μ) | p value |
|----------------------|--------------|----|-------------------------|---------|
| Epithelial Thickness | Control      | 10 | 1310.97±58.908          | .289    |
|                      | Lesional     | 40 | 1734.22±520.979         |         |
|                      | Perilesional | 40 | 1836.99±634.631         |         |
| Basal Cell Diameter  | Control      | 10 | 50.55±3.544             | .013    |
|                      | Lesional     | 40 | 48.82±3.777             |         |
|                      | Perilesional | 40 | 52.03±4.875             |         |

Figure 4: Table 2 :

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