CLINICO-PATHOLOGICAL ANALYSIS OF MALIGNANT EYELID AND ADNEXAL TUMOURS PRESENTING TO A TERTIARY EYE HOSPITAL OF EASTERN NEPAL

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ABSTRACT

Introduction

Eye lid and adnexal tumours are one of the major problems we see in the oculoplasty department. Malignant eye lid tumours are rare in comparison to benign and diversity in its clinical presentation makes it difficult to diagnose at early stage. Eyelid malignancies can mimic a host of benign neoplasms and other less aggressive neoplastic or non-neoplastic inflammatory conditions and need differentiation before definitive therapy is planned. Out of 90% of all skin cancer occurring in head and neck region 10% occurs in the eyelid. A histopathological study confirms the diagnosis that can be correlated with patient history and other clinical data.

Objective

To study the clinico-pathological correlation and its outcome in patients with malignant eyelid and adnexal tumours presenting at Biratnagar eye hospital.

Methodology

Details of the cases of histopathologically proven malignant eyelid and adnexal tumours presenting to the department of oculoplasty from July 2016 to August 2019 were recorded. The duration of disease, presenting symptoms, risk factors, demographic data, clinico-pathological correlation with its accuracy and mode of treatment were analyzed using SPSS 17.

Result

Out of total 28 patients, male were 53.6% and female were 46.4%. The mean age of presentation was 60.5±11.8 years. 53.6% tumors were located in upper lid, 39.9% were located in lower lid and 7.1% were located in medial canthus. Basal cell carcinoma and sebaceous gland carcinoma were the most common type of malignancies (35.7% each) followed by 28.6% of squamous cell carcinoma of eyelid. The clinico-pathological correlation revealed 75% of clinical accuracy. Different types of reconstructions were done according to the size and site of the defect. Modified Hughes procedure for lower eyelid defect was the most common procedure performed (32.1%), followed by 14.3% Cutler beard, 28.6% Cutler beard with post auricular cartilage graft, 10.7% Glabellar flap rotation 7.1% had combined procedure and 4.3% direct closure.

Conclusion

Sebaceous cell carcinoma and Basal cell carcinoma were the most common malignant tumors followed by squamous cell carcinoma. The upper lid was the most common site for such malignancy. Histopathological examination was useful for the establishment of complete and accurate diagnosis, which led to further management and follow up protocol for malignant eyelid tumours.

KEYWORDS

eyelid reconstructions, histopathological correlation, location, malignant eyelid tumours,
INTRODUCTION
Eyelid and adnexal tumours are commonly encountered in ophthalmic practice and form an important part in ophthalmology practice. These tumours are significant cause of morbidity and mortality especially in developing countries like ours. Different varieties of tumor can occur at different tissue level. Out of 90% of all skin cancer occurring in head and neck region, 10% occurs in the eyelid.1 Malignant eyelid tumours are rare in comparison to benign tumours, but diversity in its clinical presentation makes it challenging to diagnose and treat early. Eyelid malignancies can mimic a host of benign neoplasms and other less aggressive neoplastic or non-neoplastic inflammatory conditions and need differentiation before definitive therapy is planned.7 Also, different types of malignant eyelid tumours may have similar gross appearance but their clinical course, outcome and prognosis may be different. Although most of the eyelid tumours are diagnosed clinically, histopathological examinations give us the final diagnosis which can be correlated with the clinical picture. This helps us to plan further management, know the prognosis of the disease and thus provides great benefit to ongoing patient care.3

The main malignant tumors affecting the eyelid are basal cell carcinoma (BCC), sebaceous gland carcinoma (SGC), squamous cell carcinoma (SCC), and malignant melanoma (MM). Basal cell carcinoma (BCC) is the most common malignant eyelid tumor and accounts for 80–95% of all eyelid malignancies in Western countries.4-5 Sebaceous gland carcinoma occur less frequently in west than in Asian countries.6-11 There are thus global variations in the type of eyelid and adnexal tumors according to the geographic location and ethnicity.

There are limited studies from Nepal related to tumors of eyelid and adnexa with their clinicopathological correlation. This study was done to analyze the demography, identify the clinical presentation of different malignant eyelid and adnexal tumors, gross and histopathological diagnosis and its clinicopathological correlation with the clinical diagnosis that makes it more helpful for further management of the patients.

METHODOLOGY
This is a hospital based, cross sectional, descriptive study which included data of all the consecutive cases of malignant eyelid tumors presenting to the Department of Oculoplasty of Biratnagar Eye Hospital between July 2016 to August 2019 and who underwent surgical excision with histopathology examination. The ethical approval was taken from institutional review committee and the study adheres to the declaration of Helsinki.

Inclusion criteria: All consecutive patients with eyelid and adnexal tumours who were surgically treated and had malignant tumour as histopathological diagnosis were included in the study.

Exclusion criteria: Patients who did not give consent for the study, those who had inflammatory, infective or benign lesion in histopathology examination and those cases which did not have histopathology reports were excluded from the study.

The data extracted were: age, sex, laterality, tumor location and its extent, duration of the disease, presenting symptoms, risk factors like tobacco chewing, clinical diagnosis, lymph nodes involvement at the time of presentation, preoperative and postoperative clinical photographs to ensure the clinical diagnosis. Recorded surgical procedure included excision of the mass with 4-5mm clear margin followed by different types of reconstructions like direct closure, modified Hughes, cutler beard or any of the needed combined flaps and grafts depending upon the size of defect following the tumour excision. All specimens were sent for histopathological examination with sutures placed in the specimen to label its orientation for seeing the involvement of margin and extent of tumour pathology. Histopathology reports were retrieved and clinicopathological correlation with its accuracy was noted. All the data were analyzed using SPSS 17, chi square test were used for categorical data.

RESULTS
A total of 28 patients with a histopathology proven eyelid and adnexal malignancy were included in this study. The mean age of presentation was 60.5 ± 11.8 years (range from 32 years to 81 years). Among them 53.6 % were male and 46.4 % were female. According to the types of malignancy, male had more number of squamous cell carcinoma and female had equal number of basal cell carcinoma and sebaceous gland carcinoma [Table 1]. Left eye was predominantly affected (53.6%).

| Table 1: distribution of cases according to the type of lesion |
|------------------|-----------------|------|------|
| TYPE | NUMBER | % | MALE | FEMALE |
| BCC | 10 | 36.7 | 4 | 6 |
| SGC | 10 | 35.7 | 4 | 6 |
| SCC | 8 | 28.6 | 7 | 1 |
| TOTAL | 28 | 100 | 15 | 13 |

Regarding the site of the tumor, upper lid was most commonly involved (53.6 %) followed by lower lid and medial canthus area in 39.9 % and 7.1 % respectively. Among the patient who had upper lid malignancy, 60 % had more than 2/3rd of upper lid involved, 26.7 % had involvement in lateral part and 13.3 % had involvement in medial part. Similarly, lower lid malignancy was seen in medial part in 45.5 % and in lateral part in 45.5 %.

Mean time lapse between appearance of symptoms and presentation at hospital was 25 ± 29.47 months. Due to late presentation in most of the cases, the presenting signs also varied. However, 53.6 % had only mass and 28.6 % had ulcerated mass at the time of presentation [Table 2]. Out of
total 28 cases, 25.3 % had palpable regional lymph nodes at the time of presentation. Considering tobacco as one of the commonest risk factor of malignancies, 39.3 % were active smokers and 10.7 % were tobacco chewers.

<table>
<thead>
<tr>
<th>Table 2: Clinical features at the time of presentation</th>
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<tr>
<td><strong>Presenting features</strong></td>
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<tr>
<td>Mass</td>
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<tr>
<td>Ulcerated wound</td>
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<td>Mechanical ptosis</td>
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<td>Orbital extension</td>
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<td><strong>Total</strong></td>
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Out of 28 cases, 10 cases each (35.7%) were diagnosed as Sebaceous gland carcinoma (SGC) and Basal cell carcinoma (BCC) and 8 cases (28.6 %) were diagnosed as Squamous cell carcinoma (SCC) after the histopathology examination. Among them in BCC, lower lid was the most common site of involvement (6 cases) followed by upper lid and medial canthus (2 cases in each). Likewise upper lid was mostly involved in SGC (7 cases) than lower lid (3 cases). And SCC was more commonly found in upper lid (6 cases) than in lower lid (2 cases). 75 % accuracy was seen while correlating the clinical and histopathological diagnosis in all cases [Figure1]. 67.9 % of cases had margin positive for malignancy on histopathology examinations. Among them 2 cases had palpable lymph nodes.

All the cases underwent complete mass excision with 4-5 mm surrounding clear margin and reconstruction at the same time. Different types of reconstructions were done according to the size and site of the defect [Figure 2]. Modified Hughes’s procedure for lower eyelid defect was the most common procedure performed (32.1%).

Only 3 patients (10.7 %) had more than 1 year follow-up, 5 patients (17.9 %) had 6 months and 3 months follow up in each group. At least 6 weeks of follow-up was seen in most of the cases (10/28, 35.7 %). (Picture: 1, 2 and 3). 3 patients did not come for follow up after the initial surgery. Out of the total 28 patients operated, 3 had recurrence (2 SGC and 1 BCC) and among them BCC had recurrence in 3 month, SCC and SGC and in 3months and 6 weeks respectively. All of them had margin involvement in histopathological examinations. One of the SGC case had lymph nodes involvement too. All of them were referred to oncology center for further management.

**Figure 1**: Clinicopathological correlation of different eyelid malignancies

**Figure 2**: different types of reconstructions after tumour excision

**Picture 1**: showing preoperative and post-operative (after 6 weeks) photographs of right lower lid BCC following modified Hughes procedure.

**Picture 2**: showing preoperative and post-operative (6 weeks and 1 year) photographs of right medial canthal BCC following glabellar flap rotation flap
DISCUSSION

Various studies have been done regarding eyelid and adnexal tumors in different parts of the world. These tumors may show geographical and ethnic variation. The data of our study shows the demography, various modes of presentations and ethnic variation of the malignant eyelid tumors presenting in a tertiary eye care centre in the eastern part of Nepal. This data will serve as a reference for this geographical region and guide in planning resources for screening as well as in proper management of the eyelid tumors. It will also be useful in assessing the requirements of oculoplasty set up of the hospital and will help in future eye health care provisions.

The mean age of presentation was 60.5 ± 11.8 years (32 years to 81 years) in this study, which is similar with other studies done by Gautam P et al, Kaliki S. et al and I. Hussain et al where mean age of presentation was >50 years. This shows that eyelid malignancies are more commonly seen in older age group. In our study, male predominance is seen (male 53.6 % vs female 46.4 %), in contrast to other studies done by Wang CJ et al, Karan S et al, Gosai et al, where female were reported to be more affected. Pombejara et al reported that male was more commonly affected similar to our study.

The mean time lapse between appearance of symptoms and presentation at hospital was 25 ± 29.47 months in this study. The corresponding time in an Indian study was 7.2 months for SGC, 13.5 months in a Turkish study and 37.8 months in a study from Pakistan. The delayed presentation may be due to distance of hospital, lack of awareness and self-neglect in old age and socio-economic problems.

In our experience, BCC and SGC was the most common tumor (35.7 % each) followed by SCC (28.6 %). In another study from Nepal, BCC was the commonest eyelid tumor (59.4 %) followed by SCC (15.6 %) and melanoma in 15.6 %. A study from Singapore reported the frequency of different eyelid cancers over a 27-year period as follows: BCC 84 %, SGC 10.2 %, SCC 3.4 % and MM 1.2 %. Similar study from India over a 34-year period showed BCC was commonest (38.8 %) followed by SGC in 27.1 %, SCC in 22.4 % and MM in 3.5 %. In a study done in Central India by Jahagirdar et al where a series of 27 cases of eyelid malignancies were analyzed, sebaceous cell carcinoma (37 %) was almost as prevalent as basal cell carcinoma. Studies from Pakistan show BCC was commonest eyelid tumor (59 %) followed by SCC (31.5 %) and SGC (6.8 %) in north-west Pakistan while in southern Pakistan, BCC (56.32 %) was commonest, tumor followed by SCC (20.69 %) and SGC (14.94 %). Cumulative results of 26 studies from Japan (1976-2004) mentioned BCC to be 39.5 %, SCC 21.8 %, SGC 27 %. Thus, there are geographical variations in tumour frequency. In our study we could see that BCC and SCC were mostly present in lower lid (60 % and 75 % respectively) and SGC was mostly present in the upper lid (70 %). Various studies have shown the predominance of BCC in the lower eyelid. Upper lid is the most commonest site for periocular SGC, accounting for half to two thirds of cases due to a predominance of meibomian glands in the upper eyelid. No case of melanoma was seen in this study.

When we see gender predilection of these tumours, our study shows that SCC is more common in male than in female (7 cases and 1 case respectively). Whereas a study done by Kaliki S. et. al showed slight female predilection with a male: female ratio of 1: 1.22 BCC and SGC were more common in female in our study. This is similar to the study done by Kaliki S. et.al. In this study 75 % of total cases had clinical and histopathological correlation, which is statistically significant (p value <.001).

In this study 67.9 % of cases had margin positive for malignancy on histopathology examinations. This is mainly due to large size and extent of tumour which made it difficulty in total excision of tumour with clear surrounding margins.

Recurrence was seen in 3 cases (10.7 %) in our study, in one case of BCC and two cases of SGC. Kale et al reported 1.90 % of recurrence and Bagheri et al reported 19 % of BCC recurrence. None of the patients had any postoperative complications. Although most of the malignant eyelid tumors have favorable outcome in our experience, early diagnosis and intervention remains a corner stone for adequate functional and cosmetic lid reconstruction.

CONCLUSION

Basal cell carcinoma and sebaceous gland carcinoma is commonest malignant eyelid tumour in the older population. Most of the malignant lesions were confined to the eyelid only. Due to delay in presentation, the size and extent of tumour involvement varies. This leads to difficulty in total excision of tumour with clear surrounding margins. Reconstruction of bigger defects after the large tumour excision becomes more challenging. Timely diagnosis, management and proper follow-up are essential to reduce associated mortality and morbidity.

RECOMMENDATION

Every eyelid tumours should be treated with proper clinical examination, complete surgical excision and histopathological examination to reduce patient’s morbidity in subsequent follow up. In the setup where patient follow up is poor more than 5mm of clear margins should be taken to ensure free margin in histopathological examinations. Larger studies with longer follow up should be conducted to identify the accurate clinicopathological features and prognosis of malignant eyelid tumors for further management.
LIMITATION OF THE STUDY

There are few limitations of this study like the study being retrospective study, small sample size and short follow-up. Due to various reasons there are poor follow up of patients.

ACKNOWLEDGEMENT

Oculoplasty Department, Biratnagar eye hospital and Prof. Dr. Sanjay Kumar Singh for the continuous support.

CONFLICT OF INTEREST

None

FINANCIAL DISCLOSURE

None

REFERENCES