Case of Rapidly Expanding Conjunctival Malignant Melanoma Initially from Primary Acquired Melanosis Diagnosed 14 Years Earlier

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Abstract: Primary acquired melanosis (PAM) of the conjunctiva is a potentially serious melanocytic lesion that can lead to the development of a melanoma. A 60-year-old woman noticed pigmentation of the conjunctiva of her left eye for more than 10 years. She underwent excisional biopsy combined with cryotherapy and was diagnosed with PAM without atypia by intraoperative consultation. She was followed for 7 years, and no changes were observed. Fourteen years after the initial biopsy, she noted a growing conjunctival tumor, and a melanoma was suspected. She underwent orbital exenteration and skin grafting procedures. Histopathological examination of the specimen led to a diagnosis of conjunctival malignant melanoma. Re-examination of the initial biopsy specimen revealed that there was a proliferation of melanocytes that partially expanded over the basal layer of the conjunctiva which had been diagnosed as PAM with moderate atypia. We conclude that this case of conjunctival PAM had progressed to a conjunctival malignant melanoma after 14 years. Pathological evaluation of intraepithelial lesions has its limitations; thus, cases of PAM, even in the absence of obvious atypia, require careful follow-up.

Keywords: primary acquired melanosis, conjunctival malignant melanoma, malignant melanoma, conjunctival melanoma, conjunctival tumor

Introduction

Primary acquired melanosis (PAM) of the conjunctiva is a potentially serious melanocytic lesion that can lead to the development of a melanoma.¹⁻⁴ A conjunctival melanoma is a rare malignant lesion of the ocular surface, and it has been estimated that approximately 75% of conjunctival melanomas arise from PAMs.⁵⁻⁷ We report a case with a conjunctival PAM that developed into a conjunctival malignant melanoma after 14 years.

Case Report

A 60-years-old woman reported that she noticed pigmentation of the conjunctiva of the left eye beginning about 10 years earlier. She was suspected of having conjunctival melanoma and visited our hospital in 2006. Our initial examination found that she had no history of ocular or systemic diseases. Her decimal best-corrected visual acuity (BCVA) was 1.2, and her intraocular pressure was normal in both eyes. Slit-lamp examinations showed that the conjunctiva of the right eye was normal but diffuse disseminated pigmentation was detected around the cornea of the left eye (Figure 1). The cornea...
was clear, the anterior chamber was normal, the lens was transparent, and no abnormalities were observed in fundus of both eyes. She underwent excisional biopsy combined with cryotherapy. Histopathological analysis by frozen section suggested PAM with subepithelial reactive lymphoid hyperplasia, as there was no melanocytic nests or obvious cellular atypia and pigmentation (Figure 2). She was followed for 7 years after the biopsy and no changes were observed (Figure 3). She was then followed at the hospital near her home.

Fourteen years after the excisional biopsy, she felt discomfort of the left eye and noted a conjunctival tumor that was growing. She then consulted our hospital in 2020. Our examination showed that her BCVA was the same; however, a black tumor that extended from the fornix and disseminated pigments at the bulbar and palpebral conjunctiva were detected in her left eye (Figure 4). A conjunctival malignant melanoma was suspected, and she underwent orbital exenteration and skin grafting procedures at the Kyoto Prefectural University of Medicine Hospital. Histopathological analysis revealed that the
tumor was a conjunctival malignant melanoma (Figure 5) but no metastasis was found by a general check-up and her postoperative course was good.

**Discussion**

PAM is usually initially managed with observation alone or with excision and cryotherapy.\(^5\)\(^-\)\(^10\) PAM with severe atypia shows progression to melanoma in 13% to 46% of the eyes whereas PAM without atypia or with mild atypia shows 0% progression to malignant melanoma.\(^1\)\(^4\) Therefore, we prepared specimen again from a permanent formalin-fixed paraffin embedded block of the frozen section diagnosed in 2006, and confirmed inconspicuous nests of melanocytes with mild to moderate atypia extending throughout epithelial thickness. It was then diagnosed by histological scoring to be a conjunctival melanocytic intraepithelial neoplasia (C-MIN) of 2–3 (Figure 6). Thus, the diagnosis was PAM with moderate atypia.

The extent of the conjunctiva involved by PAM is the most important clinical risk factor for the progression to conjunctival melanoma.\(^5\)\(^-\)\(^10\) Our case was diagnosed as conjunctival PAM without atypia but the diagnosis was made from a small piece of conjunctival lesions and the other lesions might have demonstrated worse pathological findings. Although we followed up multiple pigmented lesions carefully, these lesions were left 7 years after the initial biopsy and after then, they had developed. Thus, it might have been better to remove the remaining pigmented lesions or to treat with topical anti-tumor eyed drops such as mitomycin c.\(^10\)

In conclusion, we recommend that cautious observations and follow-up should be required for PAM.

**Figure 5** Histopathological specimen of the conjunctival tumor (H&E staining). Tumor was a conjunctival malignant melanoma.

**Figure 6** Re-examined of the initial excised sample. (A) There is reactive lymphoid hyperplasia (original magnification, X 20). (B) Melanocytic nests partially expanded throughout epithelial layer of the conjunctiva scoring for conjunctival melanocytic intraepithelial neoplasia (C-MIN) of 2–3 (original magnification, X 200). The diagnosis was PAM with moderate atypia.

**Conclusion**

Pathological evaluation of intraepithelial lesions has its limitations especially small biopsy specimen; thus, cases of PAM, even in the absence of obvious atypia, require careful follow-up.

**Abbreviations**

PAM, primary acquired melanosis; BCVA, best-corrected visual acuity; C-MIN, conjunctival melanocytic intraepithelial neoplasia.

**Data Sharing Statement**

All data supporting the conclusions of this article are included in this published article.
Consent for Publication
Written informed consent was obtained from the patient for publication of this case report and any accompanying images. A copy of the written consent is available for review by the Editor of this journal.

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Author Contributions
All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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