Is There an Increased Prevalence of Asteroid Hyalosis in Eyes with Uveal Melanoma? A Histopathologic Study

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Keywords

Asteroid hyalosis · Uveal melanoma · Prevalence

Abstract

During the planning meeting for the Collaborative Ocular Melanoma Study (COMS) prior to the start of patient recruitment in 1986, there was an interest expressed in determining whether a relationship existed between the presence of uveal melanoma (UM) and asteroid hyalosis (AH). To answer this question, the ophthalmic examination form (unlike the pathology form for enucleated eyes) for each COMS patient asked whether AH was present or not. Though an increased prevalence was not found, this result was never published. A recent unpublished study at the University of Wisconsin School of Veterinary Medicine indicated a higher prevalence of AH in canine eyes with UM when compared to control eyes (without tumor) enucleated for goniodysgenesis. This further increased our inter-

est in revisiting the published literature, clinical records, and histopathology slides of the enucleated eyes from the COMS study, as well as the histopathology slides on file in the University of Wisconsin Eye Pathology Laboratory. While cases with both AH and UM were occasionally encountered in the literature, clinically, we could not find a previous study focusing on these two processes. This study was conducted to explore whether such an association exists.

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Background

Asteroid hyalosis (AH) is a degenerative and age-related process in the vitreous of unknown etiology with a reported clinical and histological prevalence ranging from 0.36 to 2.9% [1–3]. Histopathologically, the asteroid bodies appear as basophilic crystalline spherules that stain

Fig. 1. Images depict round spherules characteristic of asteroid hyalosis (arrows) in an eye with uveal melanoma (UM) on hematoxylin and eosin (**a**, original magnification $\times 100$), stained blue with Alcian blue staining (**b**, original magnification $\times 100$; **c**, original magnification $\times 400$).

positively for acid mucopolysaccharide, and are birefringent with a Maltese cross pattern under polarized light.

Uveal melanoma (UM) has an incidence of 7 cases per million per year in the United States, making it the most common primary intraocular malignancy in adults. The mean age at presentation is 59 years old and it affects both sexes equally [4]. Its etiology is unknown, but many studies have given insight into the associated chromosomal and genetic changes.

Materials and Methods

The clinical prevalence of AH in eyes with UM was determined by reviewing the clinical records of eyes enucleated during the Collaborative Ocular Melanoma Study (COMS). As a control, the clinical prevalence of AH was established from patients without UM seen in the retina department in the University of Wisconsin Eye Clinic by a single observer (J.L.G.), who systematically documented the presence of AH in his patients. The clinical association between UM and AH was determined by comparing the prevalence between the two groups and compared using a two-sample proportion test.

The histopathological prevalence in eyes with UM was obtained by reviewing archived histology slides and specimens from

the COMS and eyes with UM on file in the University of Wisconsin Eye Pathology Laboratory. Multiple hematoxylin and eosin slides in each case were examined by several experienced reviewers (M.R.K., A.A.A., H.D.P., V.L., R.R.D., and D.M.A.), and cases with a sufficient amount of vitreous had the vitreous screened for the presence of basophilic crystalline spherules. The unstained slides in suspicious cases were stained with Alcian blue. The spherules were confirmed to be AH with Alcian blue staining and polarization (Fig. 1a–c).

Results

Review of the clinical records of 1,548 eyes from the COMS revealed a clinical prevalence of 1.2% (19 of 1,548 eyes) amongst eyes with UM. In the control group, the clinical prevalence of AH in eyes without UM was 0.82% (43 of 5,218 eyes). This 1.5-fold increase in the clinical prevalence of AH in UM eyes, when compared to the controls was not found to be statistically significant (95% CI 0.871–2.55, $p = 0.19$).

There were a total of 1,985 UM histopathology specimens (1,759 eyes from the COMS and 226 eyes from the eye pathology laboratory at the University of Wisconsin);

out of these, 28 (1.4%) cases had anterior uveal involvement and 1,957 (98.6%) cases had only posterior uveal involvement. The histological prevalence of AH amongst the COMS and University of Wisconsin eyes was 0.7% (13 of 1,985 eyes).

Discussion

AH is a complex but poorly understood entity associated with a number of biochemical changes in the eye. It has been suggested that it is caused by toxins diffusing into the vitreous from degenerating retina [5, 6]. It is well demonstrated that the prevalence of both AH and UM increases with age [2, 7, 8]. While the prevalence of AH in our control group was comparable to previously reported rates, including the largest study from Bergren et al. [9], it was less than the rates found in the Beaver Dam Eye Study and a study performed at the University of California at Los Angeles (UCLA) [2, 3]. These differences can be explained by the different data collection and investigation methods employed by each study. Additionally, patients from the Beaver Dam and UCLA studies were older than our control group, which may have contributed to the higher prevalence in those studies.

As with all retrospective clinical studies, there were several limitations. We did not have a histological control group and due to the sampling of different groups, the clinical versus histological prevalence of AH was unable to be assessed via McNemar's test. Secondly, clinical detection of AH by the examination of a single clinician may

not have the same sensitivity as examination by several clinicians. Part of the increased clinical prevalence of AH in eyes with UM versus controls may reflect more thorough posterior segment examinations and/or more detailed record keeping in eyes with known UM. Histological examination of the vitreous has its own pitfalls, since much of the vitreous is lost during processing. This may account for our clinical detection rate being higher than our histological rate in eyes with UM. Furthermore, there is a possibility that the patients in either of our groups had other systemic and/or ocular conditions contributing to AH. In conclusion, there does not appear to be a statistically significant association between AH and UM.

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Statement of Ethics

The authors have no ethical conflicts to disclose.

Disclosure Statement

The authors have no conflicts of interest.

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