Sympathetic ophthalmia in fellow eye after vitrectomy for massive subretinal hemorrhage secondary to polypoidal choroidal vasculopathy

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Abstract: We experienced a case of sympathetic ophthalmia in a fellow eye after vitrectomy for subretinal hemorrhage related to polypoidal choroidal vasculopathy. A 60-year-old male consulted us for polypoidal choroidal vasculopathy with subretinal hemorrhage in his left eye. The recurrence of massive subretinal hemorrhage refractory to repeated pars plana vitrectomies leads to phthisis bulbi. Two months later, multiple serous retinal detachments were observed in his right eye. Positive human leukocyte antigen-DR4 and the uveitis were helpful in distinguishing between sympathetic ophthalmia and age-related macular degeneration. High-dose pulse intravenous steroid contributed to recovery of visual acuity after resolution of serous retinal detachment.

Keywords: sympathetic ophthalmia, polypoidal choroidal vasculopathy, vitrectomy, vitreous hemorrhage, uveitis

Introduction
Sympathetic ophthalmia is a rare, bilateral granulomatous uveitis that occurs after a penetrating trauma or a surgery. 1-3 The time from trauma to onset of uveitis varies between patients, ranging from a few days to a few decades, although 90% of cases occur within a year after a triggering injury to an eye. 1 Sympathetic ophthalmia shares some similar clinical, genetic, and histopathological features with Vogt–Koyanagi–Harada (VKH) disease since both are autoimmune disorders targeting the melanocytes. 4-7 Currently, a retinal surgery is an emerging risk factor for sympathetic ophthalmia. 8-10 We describe a case where the onset of sympathetic ophthalmia is possibly associated with the vitrectomy performed to treat subretinal hemorrhage from polypoidal choroidal vasculopathy (PCV) in an exciting eye.

Case report
A 60-year-old Japanese male presented for upper visual field defect in the left eye in June 2015. His best-corrected visual acuity (BCVA) was 20/17 in the right eye and 20/22 in the left eye. There were no signs of chronic inflammation in both eyes, such as mutton-fat keratic precipitates, anterior chamber flare, and posterior synechiae, at initial presentation. In the left eye, massive submacular hemorrhage, sub-retinal pigment epithelial (RPE) hemorrhage, and an orange-red lesion characteristic to PCV were observed (Figure 1A, B). Fluorescein angiographic image obtained using Optos® California (Optos, Dunfermline, Scotland, UK) manifested hyperfluorescent area corresponding to the orange-red lesion and surrounding blocked fluorescence.
due to sub-RPE hemorrhage (Figure 1C). Also, indocyanine green angiographic image obtained using Heidelberg Retina Angiography 2 (HRA2®; Heidelberg Engineering GmbH, Heidelberg, Germany) showed focal hyperfluorescence in the area of polypoidal lesion (Figure 1D). Because of complicated vitreous hemorrhage, 25-guage pars plana vitrectomy combined with phacoemulsification and intraocular lens implantation was performed in July 2015. In November 2016, bullous hemorrhagic RPE detachment and marginal subretinal hemorrhage occurred in the right eye (Figure 1E), accompanied by vitreous hemorrhage. The patient underwent second 25-guage vitrectomy with silicone oil tamponade in December 2016. High intraocular pressure and hyphema persisted postoperatively and the left eye led to phthisis bulbi despite the third and fourth vitrectomies. No tissue plasminogen activator, gas, or cryotherapy was used during any of the surgeries.

In February 2017, the patient presented with a complaint of blurred vision in his right eye. He had tinnitus and headache with hypersensitivity of scalp from 2 weeks before the onset of blurred vision. The BCVA was 20/17 in the right eye and no light perception in the left eye. In the right eye, slit-lamp biomicroscopy showed cells in the anterior chamber. Fundus examination revealed multiple serous retinal detachments with a subretinal septum (Figure 2A, B). Fluorescein angiography revealed a broad area of hyperfluorescence in the posterior pole and optic disc during the late phase (Figure 2C). Human leukocyte antigen (HLA)-DR4 was positive, and the patient was diagnosed as sympathetic ophthalmia. Since the BCVA of the right eye dropped to 20/67 due to serous retinal detachments, a pulse steroid therapy (intravenous methylprednisolone 1,000 mg/day for 3 days) followed by oral prednisolone 40 mg/day was initiated. Two weeks later, the BCVA improved to 20/20. In May 2017, disappearance of serous retinal detachment was confirmed by optical coherence tomography (Figure 2D). Oral prednisolone was tapered gradually until December 2017 and was discontinued. No recurrence was observed thereafter. A written informed consent was obtained from the patient for use of his medical record data and images in this case report. The institutional approval was not required for publication of the case report.

Discussion

We observed a case of sympathetic ophthalmia, which may be associated with vitrectomy performed to treat subretinal hemorrhage due to PCV. According to a population-based study conducted in the United Kingdom, the estimated risk of sympathetic ophthalmia after vitrectomy was 1 in 799 eyes (0.125%), which was higher than 1 in 1,357 eyes (0.074%) after external retinal detachment surgery.9 Another study from the United Kingdom and Republic of Ireland reported that the minimum incidence of sympathetic ophthalmia was 0.03 per 1,00,000 people. Of those reported, pars plana vitrectomy was the sole cause of sympathetic ophthalmia in 33% of the patients.10 The risk of patients developing sympathetic ophthalmia may rise due to increase in number of vitrectomies performed globally in recent years.

We carefully ruled out the possibility of the bilateral occurrence of age-related macular degeneration and other inflammatory diseases. Electrocardiogram and chest X-ray were normal. Laboratory blood test showed white blood cell count 8,560/μL, C-reactive protein 0.01 mg/dL, angiotensin I converting enzyme 13.8 IU/L, lysozyme 3.9 IU/L, Treponema pallidum hemagglutination negative, fluorescent treponemal antibody absorption negative, T-SPOT®.TB (Oxford Immunotec Ltd., Abingdon, Oxfordshire, UK) negative,
Bartonella henselae deoxyribonucleic acid negative, and HLA-DR4 positive. B. henselae was examined to rule out cat scratch disease from a pet cat that the patient had. The test results were consistent with the previous reports that Japanese patients with sympathetic ophthalmia and VKH disease were prone to express HLA-DR4, HLA-DRw53, and HLA-Bw54.1,6,7 Both diseases share similar clinical, genetic, and histopathological features, such as Dalen–Fuchs nodule (inflammatory choroidal nodules manifested as white dots in the fundus), similar HLA expression patterns, and T-cell-mediated autoimmune response.1,3–7 HLA-DR4 expression in our case was one of the supporting evidences for making a diagnosis of sympathetic ophthalmia.

A pulse steroid therapy followed by high-dose oral prednisolone was an effective treatment option for sympathetic ophthalmia. Currently, systemic steroid administration is considered to be the primary therapy. Oral prednisone 1.0–2.0 mg/kg/day tapered gradually over 3–4 months is often administered as an initial treatment. Also, a cycle of pulse steroid therapy, ie, intravenous administration of methylprednisolone 1,000 mg/day for 3 days, is often administered in severe cases.1 In our case, the patient was treated with a cycle of pulse steroid therapy followed by oral prednisolone administration. We consider that systemic steroid treatments have successfully contributed to preventing bilateral blindness from prolonged inflammation in our patient.

Currently, an enucleation is the best prevention against sympathetic ophthalmia.1 In a previous study where 55 enucleations (11 primary and 44 secondary cases) were performed among 660 open globe injuries, only two patients (0.3%) developed sympathetic ophthalmia.11 In our case, if the enucleation of the left eye was performed when ablepsia was unavoidable due to persistent high intraocular pressure after the second vitrectomy, sympathetic ophthalmia in the right eye may not have occurred. In the future, if a uveitis in the right eye recurs after the discontinuation of systemic steroid treatment, the enucleation of the left eye is an option to be considered.

Conclusion
Our case manifested sympathetic ophthalmia in the fellow eye that occurred after the vitrectomy performed to treat subretinal hemorrhage from PCV in the exciting eye. Bilateral occurrence of age-related macular degeneration and other inflammatory diseases should be carefully ruled out before making a diagnosis of sympathetic ophthalmia.

Author contributions
All authors contributed toward data analysis, drafting and critically revising the paper and agree to be accountable for all aspects of the work.

Disclosure
The authors report no conflicts of interest in this work.

References

Figure 2 Serous retinal detachments due to sympathetic ophthalmia in the right eye.

Notes: (A) Fundus photograph of the right eye showing serous retinal detachments in the posterior pole. (B) OCT of the right eye showing serous retinal detachment. An arrow indicates subretinal septum, which is the characteristic of sympathetic ophthalmia. (C) Late-phase fluorescein angiography shows a broad area of hyperfluorescence in the posterior pole and optic disc. (D) OCT of the right eye after a cycle of pulse steroid therapy and oral prednisolone administration showing disappearance of serous retinal detachment.

Abbreviation: OCT, optical coherence tomography.


