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Central retinal vein occlusion with secondary cilioretinal artery occlusion following oral minoxidil use: A case report and literature review

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Abstract

Minoxidil, a drug that is widely used for hair loss treatment, is considered a safe medication. While there are minor side-effects such as scalp irritation or systemic effects like tachycardia, it is rarely associated with severe ocular complications. We report a case where a 35-year-old man, taking oral Minoxidil for alopecia, developed Central Retinal Vein Occlusion (CRVO) with Cilioretinal Artery Occlusion (CLRAO), leading to sudden vision loss. After Minoxidil discontinuation, the patient's vision significantly improved. This incident emphasises the importance of recognizing potential ocular side effects of Minoxidil among healthcare professionals and necessitates further research to understand its comprehensive impact on health.

Keywords: Central retinal vein occlusion; Cilioretinal artery occlusion; Minoxidil.

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Introduction

Initially developed for the management of hypertension in the 1970s, Minoxidil is now widely used to treat hair loss in both men and women. Functioning as an active vasodilator, Minoxidil promotes blood flow by relaxing blood vessels [1]. It is currently accessible Over The Counter (OTC) as topical solutions of 2% and 5% and is prescribed as a daily oral medication for the management of hypertension or androgenic alopecia [2].

The most notable side effects of the topical formulation of Minoxidil are scalp irritation or allergic contact dermatitis. Systemic side effects include tachycardia, increased cardiac function and stroke volume, sodium retention, and abnormal hair growth [3]. Information leaflets and drug monographs list the main ocular side effect as conjunctivitis and blurred vision, mostly associated with the topical formulation of Minoxidil. Nonetheless, there have been a few case reports that have previously shown Minoxidil-induced ocular pathologies, all associated with the use of its topical formulation [3-6]. Herein, we present a case of Central Retinal Vein Occlusion (CRVO) with secondary Cilioretinal Artery Occlusion (CLRAO), likely caused by the chronic use of Minoxidil.

Case presentation

A 35-year-old male presented to the Royal Victoria Hospital (RVH) Emergency Department (ED) with sudden vision loss in the superior nasal quadrant of the right eye (OD). He reported no significant medical history other than alopecia and migraines. His family history was non-contributory, and his only prescription medications were nortriptyline 10 Milligrams (mg) daily and Minoxidil 1.25 mg daily.

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The initial ophthalmic evaluation showed his Best-Corrected Visual Acuity (BCVA) to be 20/40 OD and 20/20 in the left eye (OS). The maximum intraocular pressure (Tmax) was 15 OD and 19 OS. His pupils were equal, symmetric, and reactive to light, with no Relevant Afferent Pupillary Defect (RAPD), and he had full Extraocular Motion (EOM) in both eyes (OU). Slit-lamp biomicroscopy examination of the anterior segment was unremarkable. However, Dilated Fundus Examination (DFE) of the posterior segment revealed scattered mild retinal hemorrhages and tortuosity with patchy retinal whitening in the inferior macula (Figure 1A). Optical Coherence Tomography (OCT) revealed inner retinal hyper reflectivity (Figure 2). He was clinically diagnosed with CRVO with secondary CLRAO.

Fluorescein Angiography (FA) performed two days later revealed a small peripheral branch with delayed filling and mild vascular leakage, in keeping with CRVO. However, the CLRA filling was found to be within normal limits. Fundus photography at day six revealed macular edema, dot-blot hemorrhages, and venous tortuosity (Figure 3). At this time, he was advised to cease the use of Minoxidil. He was also treated with a course of prednisone, tapered from 50 mg orally for 5 days between weeks two and three. Over the next few weeks to months, the patient underwent detailed systemic evaluations and investigations. An Electrocardiogram (EKG) and Transthoracic Echocardiogram (TTE) were negative for cardioembolic events. Computed Tomography (CT) of the head without Contrast (C-) showed no evidence of acute territorial infarct, intracranial hemorrhage, mass lesion, or hydrocephalus, and CT Angiography (CTA) of the neck revealed no acute occlusion or significant stenosis along the cervical and intracranial arterial vasculature. Importantly, bilateral ophthalmic arteries were patent. Additional investigation with Magnetic Resonance Imaging (MRI) with contrast of the head and orbits was negative for any ischemic lesion or optic neuropathy. Investigations for infectious causes, such as Bartonella henselae, toxoplasmosis, and Mycobacterium tuberculosis, were also negative. Investigations for inflammatory causes, such as antinuclear antibodies, protein C, antithrombin, and anti-neutrophil cytoplasmic antibodies, were all negative. Furthermore, genetic analysis for factor V and prothrombin (20210 A) gene mutations were also negative.

His BCVA OD improved to 20/20-1 within the first week following the cessation of minoxidil. His Visual Fields (VF) on presentation improved from near-total vision loss in the superior visual fields (Figure 4A) to residual VF defects superior-centrally at week 3 (Figure 4B), and to mild central VF defects at 3 months (Figure 4C). Although both the BCVA and VFs showed improvement within the first week, fundus photography at week one revealed worsening macular edema (Figure 5A). However, there was significant improvement by week 3 (Figure 5B). Evaluations after 3 months post-cessation of minoxidil showed his BCVA to be 20/20 OU, with no other complaints.

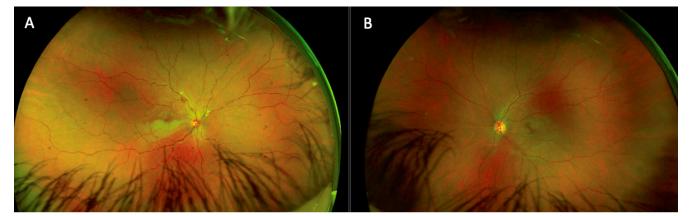


Figure 1: Fundus photography OU, on presentation. There are patchy whitening of the inferotemporal arcade with multiple dot-blot hemorrhages and dilated tortious vessels of the right eye (A). The left eye is unremarkable (B).

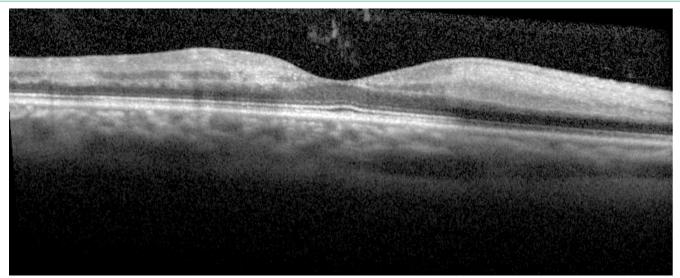


Figure 2: Optical Coherence Tomography (OCT) image taken on presentation showing inner retinal hyperreflectivity in territorial distribution of cilioretinal artery.

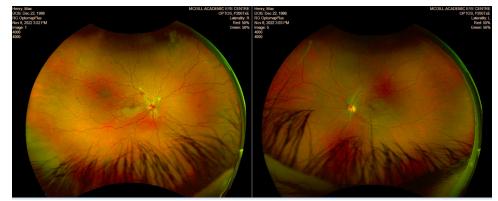
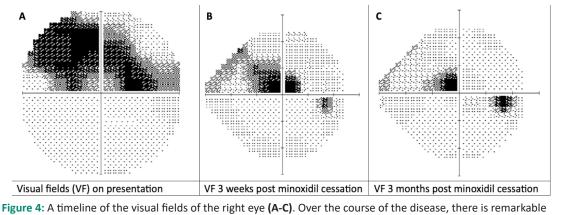


Figure 3: Fundus photography, day 6. There is macular edema, dot-blot hemorrhages, and venous tortuosity of the right eye. The left eye is unremarkable.



and gradual improvement in the visual fields (A-C).

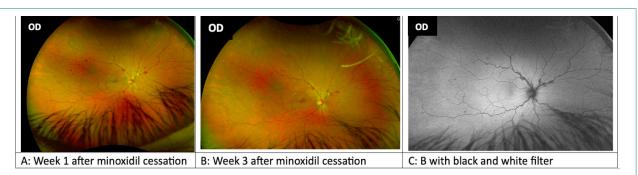


Figure 5: Timeline with fundus photography. There is improved appearance of macular edema, reduction in the vascular tortuosity, hemorrhages, and cotton wool spots **(B,C)** in comparison to 1st week **(A)**.

Discussion

We present a unique case of Central Retinal Vein Occlusion (CRVO) with secondary Cilioretinal Artery Occlusion (CLRAO) in a patient using oral Minoxidil for androgenic alopecia. While Minoxidil oral tablets are typically reserved for the treatment of refractory hypertension as it functions as direct vasodilator, the topical formulation is a popular treatment for androgenic alopecia, approved by both FDA and Health Canada [7]. Although minoxidil is generally considered to be a safe medication, few articles have shown systemic adverse events and ocular pathologies associated with the use of topical formulation of minoxidil [3-6]. However, reports of CRVO with secondary CLRAO, in the context of either topical or oral Minoxidil usage, have not been previously documented in the literature.

The cilioretinal artery arises from the posterior ciliary artery and is susceptible to sudden occlusion due to blockage of blood flow. CLRAO is a rare condition but most associated with systemic vascular diseases like atherosclerosis, giant cell arteritis, and often secondary to CRVO [8-10]. Our patient was diagnosed with CRVO and secondary CLRAO, supported by fundal examinations, Optical Coherence Tomography (OCT) findings, and Fluorescein Angiography (FA), all indicative of territorial involvement of the central retinal vein and the cilioretinal artery (Figures 1 & 2). Recent literature has suggested that a combined CRVO and CLRAO could be triggered by a temporary hemodynamic blockade in the cilioretinal artery, which occurs when a sudden increase in intraluminal pressure within the retinal capillary bed surpasses that within the cilioretinal artery [11]. This may also explain the FA findings which showed normal arterial filling with the cilioretinal artery.

Although arterial and venous occlusions are usually associated with several other etiological factors as mentioned above, our patient had no other known risk factors that could predispose to such an ocular event. Nortriptyline, another medication the patient was taking, does not have any known association with ocular pathologies according to literature review and product information, and the patient continued its usage throughout without ocular issues. Additionally, the Naranjo adverse drug reaction probability score was calculated to be 5, suggesting a probable idiosyncratic drug reaction to minoxidil usage [12]. Based on clinical history and ocular findings, the use of minoxidil, the subsequent improvement in the BCVA and VFs (Figures 4 & 5) following the cessation of minoxidil, in an otherwise young and healthy patient, this rare occurrence of combined CRVO and CLRAO is likely due to use of minoxidil.

Conclusion

It is worth noting that while there have been reports linking topical Minoxidil to ocular pathologies as mentioned above, to our knowledge, there have been no previous reports regarding ophthalmic side effect of oral minoxidil use, specifically CRVO with secondary CLRAO. Given the rarity of such reports, further investigation is warranted to explore the relationship between Minoxidil use, whether topical or oral, and adverse ophthalmic outcomes. This case underscores the necessity for increased awareness among healthcare providers regarding the potential ophthalmic side effects of Minoxidil, to ensure prompt recognition and management of such complications.

Declarations

Contributions: Motaz Bamakrid and Sidratul Rahman participated in preparing the manuscript equally.

Dr. Christian El-Hadad is the principal investigator, reviewed the manuscript, provided feedback, guidance, and expertise on the topic.

Consent: None required for case reports.

Competing interest: None.

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