CASE REPORT



Case Report: Central retinal artery occlusion following

sildenafil intake [version 1; peer review: 2 approved]

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Abstract

Purpose: To report a case of central retinal artery occlusion associated with sildenafil intake and briefly discuss its causative pathogenesis.

Methods: A 50-year-old man with no premorbidities presented with symptoms of sudden severe visual field constriction in the left eye (LE). Best-corrected visual acuity in the LE was 20/25. Fundus examination and fluorescein angiography of the LE were suggestive of central retinal artery occlusion (CRAO) with cilioretinal artery sparing. Further investigation revealed that 100 mg of sildenafil had been taken for the first time three hours before the onset of symptoms.

Results: The patient was treated promptly with intravenous acetazolamide, sublingual isosorbide dinitrate and ocular massage, but without visual recovery. No other associated systemic or local risk factors were found, and the case was classified as a potential complication of sildenafil.

Conclusion: Although no direct link could be established, the aim of this report is to highlight the incidence and to consider this issue when evaluating any case of central retinal artery occlusion.

Keywords

Central retinal artery occlusion, Systemic Drug Retinal Toxicity, Sildenafil, phosphodiesterase V inhibitor.

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Any reports and responses or comments on the article can be found at the end of the article.

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Introduction

Sildenafil is a specific phosphodiesterase V inhibitor which is a widely used treatment for erectile dysfunction. Many reports have highlighted ischemic ocular side effects associated with sildenafil.¹ We report herein a case of central retinal artery occlusion (CRAO), which occurred a few hours after oral sildenafil intake.

Case report

A 50-year-old Tunisian man, otherwise healthy and unemployed, presented to the ophthalmology department with sudden severe visual field constriction in the left eye (LE) of 48 hours duration preceded by severe headaches. On ophthalmic examination, visual acuity was 20/20 in the right (RE) eye and 20/25 in the left eye (LE). LE fundus examination revealed diffuse faint retinal whitening, except for central area of normal retinal color along the distribution of a perfused cilioretinal artery (Figure 1: black arrow).

Anterior segment examination of both eyes as well as fundus examination of theRE were unremarkable. LE fluorescein angiography (FA) showed no filling of the central retinal artery, regular filling of the cilioretinal artery and late retrograde filling of the central retinal vein (Figure 2: blue arrowheads).

OCT-A of the left eye at first presentation shows no flow in the microvasculature of the superficial and deep retinal capillary plexuses, except for the territory of the cilioretinal artery (Figure 3).

LE central retinal artery occlusion (CRAO) with cilioretinal artery sparing was diagnosed. Echocardiogram, carotid artery imaging and blood tests were unremarkable. Further investigation revealed that a 100 mg dose of sildenafil had been taken for the first time three hours before the onset of symptoms.

Ocular massage was performed as well as sublingual isosorbide dinitrate and intravenous acetazolamide were administered. Two weeks later, reduction in retinal oedema was evident on left eye fundus examination, although there was no improvement in either visual acuity or visual field.

Discussion

Various ischemic ocular events related to sildenafil have been reported. The most notable are branch retinal artery occlusion,² acute macular neuroretinopathy,³ anterior ischemic optic neuropathy,^{4,5} central retinal vein occlusion⁶ and cilioretinal artery occlusion.⁷



Figure 1. Baseline composite fundus photograph of the left eye showing retinal whitening except for the cilioretinal artery distribution area (black arrow).



Figure 2. Fluorescein angiography shows interruption of flow in the central retinal artery, regular filling of the cilioretinal artery and late retrograde filling in the central retinal vein.



Figure 3. Baseline OCT-A of the left eye shows no flow in superficial and deep retinal capillary plexuses microvasculature. Only the flow in cilioretinal artery is visible.

Only two cases of CRAO with sildenafil have been reported in the literature.^{8,9}

The patient presented here reported taking sildenafil (100 mg) a few hours before the onset of ocular symptoms and headaches. In fact, ocular side effects are directly proportional to the blood concentration of the drug, which usually appears between 15 and 30 minutes after administration, reaches a peak one to two hours later and clears halfway in 3 to 5 hours.¹⁰

As sildenafil has a high systemic vasodilator effect that reduces systemic blood pressure,¹¹ it may decrease cerebral blood flow leading to severe headaches as experienced by our patient. Similarly, numerous clinical studies have demonstrated that sildenafil induces retinal veinous vasodilatation *in vivo*.¹²

Both the absence of risk factors for retinal vascular occlusion and the timeline of events indicates that oral sildenafil was probably a contributing factor in the development of CRAO, but its pathogenesis remains speculative. We suggest that central retinal artery occlusion occurs in the region of the lamina cribrosa where the central retinal vein and artery share a common adventitia. We speculate that sildenafil-related vasodilation of the central retinal vein causes central retinal artery compression, resulting in secondary changes, including blood flow changes, endothelial damage and platelet thrombi, leading to CRAO. This case underlines the importance of a careful drug intake investigation in cases of CRAO without obvious cause. This incident should not be overlooked by physicians and must be seriously discussed with

patients requiring sildenafil, especially since most of them are at risk for ocular ischemic events. Nevertheless, this association remains poorly explained and requires further documented cases.

Conclusion

In summary, CRAO secondary to sildenafil is extremely rare and has only been reported in the literature twice previously. Clinicians should be aware of this risk and should avoid prescribing sildenafil in patients with a history of ischemic ocular events.

Data availability

All data underlying the results are available as part of the article and no additional source data are required.

Consent

Written informed consent for publication of their clinical details and clinical images was obtained from the patient.

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The author reported an original case of a young adult patient with no personal history who presented with a central retinal artery occlusion following the intake of Sildenafil.

The retinal imaging is interesting, moreover, it would be necessary to add the early times of the retinal angiography and a B mode OCT, as well as it should be specified that the imaging was not done at an acute stage of the arterial occlusion seen the blood flow recovery exceeding the cilioretinal artery area.

It may be relevant to detail the biological radiological and clinical explorations done to rule out other etiologies of retinal arterial occlusion like cardiovascular pathologies, dysimmune diseases with vasculitis, and hematological disorders

Is the background of the case's history and progression described in sufficient detail? $\ensuremath{\mathsf{Yes}}$

Are enough details provided of any physical examination and diagnostic tests, treatment given and outcomes?

Yes

Is sufficient discussion included of the importance of the findings and their relevance to future understanding of disease processes, diagnosis or treatment? Yes

Is the case presented with sufficient detail to be useful for other practitioners? Yes

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Ophtalmology Ocular Surface Disease , glaucoma, retinal diseases, anterior segment surgery, contactology

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

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The authors report a case of central retinal artery occlusion as a potential complication of Sildenafil. they conclude that Clinicians should take into account this complication while prescribing this treatment.

In this case, multimodal imaging was well detailed with high quality images. It would be also interesting if the authors could add structural OCT imaging of the posterior pole. On LE fluorescein angiography, are there more earlier phases that could better describe the retrograde filling of the posterior pole vessels? Did the general examination of the patient reveal some characteristics that could be possible markers predicting this potential complication when prescribing this treatment (obesity, sleep apnea syndrome, recent vaccination, COVID...?)

Is the background of the case's history and progression described in sufficient detail? $\ensuremath{\mathsf{Yes}}$

Are enough details provided of any physical examination and diagnostic tests, treatment given and outcomes?

Yes

Is sufficient discussion included of the importance of the findings and their relevance to future understanding of disease processes, diagnosis or treatment?

Yes

Is the case presented with sufficient detail to be useful for other practitioners? $\ensuremath{\mathsf{Yes}}$

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: retinal imaging, peridatric ophthalmology, cataract surgery, oculogenetics

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