Non-arteritic anterior ischemic optic neuropathy secondary to acute primary-angle closure

Ajay E Kuriyan
Byron L Lam
Bascom Palmer Eye Institute, University of Miami, Miami, FL, USA

Purpose: To describe a case of non-arteritic ischemic optic neuropathy (NAION) secondary to acute primary-angle closure (APAC).

Methods: Case report.

Results: A 50-year-old woman with painful visual loss in the right eye was found to be in APAC with a right afferent pupillary defect. Laser peripheral iridotomy relieved pain but did not improve vision. Diffuse optic disc edema in the right eye and a small cup-to-disc ratio in the left eye were evident. Magnetic resonance imaging was normal. The patient was diagnosed with non-arteritic ischemic optic neuropathy (NAION) secondary to APAC, a rare clinical entity which can result in markedly decreased visual acuity.

Conclusion: NAION secondary to APAC is a rare clinical entity that can result in severe vision loss.

Keywords: acute angle closure glaucoma, perfusion pressure, NAION, optic nerve

Introduction
The prevalence of acute primary-angle closure (APAC) is 0.04 to 0.6 per thousand in the United States in patients older than 40.1,2 The incidence of non-arteritic ischemic optic neuropathy (NAION) is two to ten per 100,000 in the population older than 50.3,4 NAION secondary to APAC was described by Sonty and Schwartz in 1981.5 There have been four subsequent case reports (five eyes) describing NAION secondary to APAC.6–9 Patients present with typical APAC symptoms and signs, which include pain, headache, nausea and vomiting, conjunctival injection, corneal edema, elevated intraocular pressure (IOP), closed angles, and a shallow anterior chamber. NAION was present at the time of presentation for APAC in four of six eyes and developed one week after APAC in two eyes.5–9

Case report
A 50-year-old woman with a history of hypertension and dyslipidemia presented with redness, pain, decreased vision, headache, and nausea, for approximately 10 hours. She reported no similar prior episodes. The patient was taking metoprolol, 50 mg, at bedtime; hydrochlorothiazide, 25 mg, daily; amlodipine, 25 mg, daily; lisinopril, 20 mg, daily for hypertension; simvastatin, 40 mg, at bedtime for dyslipidemia; and aspirin, 81 mg, daily for cardiovascular risk factors.

Best-corrected visual acuity (VA) was hand motion at 4 feet right eye and 20/20 left eye (manifest refraction: plano + 0.75 × 135). Her pupil was 5 mm in the right eye and 4 mm in the left, and she was noted to have a moderate relative afferent
pupillary defect (RAPD) in the right eye. Her IOP were 41 mmHg right eye and 15 mmHg left eye. The right eye had moderately severe ciliary injection, corneal edema, a shallow anterior chamber (closed peripherally, ¼ corneal thickness centrally), and a moderate nuclear sclerotic cataract. There was no posterior synechiae, and her pupil was minimally reactive. The left eye was quiet, had a shallow anterior segment (¼ corneal thickness peripherally, ½ corneal thickness centrally), and a mild nuclear sclerotic cataract. The view of the posterior segment in the right eye was obscured by the corneal edema. The left optic nerve head appeared normal with a cup-to-disc ratio of 0.1. Gonioscopy demonstrated that her angle was closed 360° in the right eye and was Shaffer Grade II (anterior trabecular meshwork) 360° in the left.

The patient was treated for APAC in the right eye with timolol, dorzolamide, and brimonidine drops every 30 minutes and acetazolamide 500 mg orally. One hour later, the IOP was 34 mmHg and she was still experiencing severe pain and decreased vision. After instillation of pilocarpine 2%, a laser peripheral iridotomy (LPI) was performed in the right eye without complications and the anterior chamber deepened, the IOP decreased to 14 mmHg, and the patient felt a marked improvement in pain. However, she reported that her vision did not improve. Due to the right RAPD, the patient was asked to return the following morning for a dilated fundus exam.

Upon the patient’s return, she reported no pain in her right eye but noted no change in vision. VA remained at hand motion at 4 feet with a moderate RAPD and an IOP of 12 mmHg. Her conjunctival injection and corneal edema had resolved. Her LPI was patent, and her anterior chamber was shallow (¼ corneal thickness peripherally), but significantly deeper than the previous day. Gonioscopy demonstrated her angle was Shaffer Grade II 360° without peripheral anterior synechiae. The right optic nerve head had 360° edema and an area of subretinal fluid extending into the nasal aspect of macula from the optic disc temporally (Figure 1A). The examination of the left eye was unchanged (Figure 1B). Optical coherence tomography (OCT) demonstrated optic nerve edema and subretinal fluid extending to the macula in the right eye (Figure 2A and B).

A brain magnetic resonance imaging (MRI) with contrast to rule out infiltrative or compressive causes of the optic nerve edema was within normal limits. The patient had no giant cell arteritis symptoms and no headaches prior to this acute event, and the headaches resolved completely after the LPI. The patient was diagnosed with NAION related to angle closure.

The patient was counseled on the importance of stopping use of her antihypertensive medications in the evening and of improved blood pressure and dyslipidemia control. An LPI was performed in her left eye without complications. Two days after presentation, the patient was started on oral prednisone taper (60 mg prednisone daily for 2 weeks, followed by 50 mg for 5 days and then 40 mg for 5 days, and then cutting down by 5 mg every 3 days to 5 mg, followed by 2.5 mg for 3 days and then 1 mg for 3 days) and instructed to follow-up in one month. The decision to use steroids in this patient was based on a retrospective study by Hayreh and Zimmerman, which demonstrated improvement in VA and visual field in NAION patients who presented within two weeks of onset.10 A larger improvement was seen in patients with poorer presenting VA.10

One month later, VA was count fingers at two feet in the right eye and 20/20 in the left. The IOP was 12, right eye, and 14, left eye. A Humphrey visual field of the right eye demonstrated a cecocentral scotoma (Figure 3). The optic nerve edema of the right eye resolved and the disc had diffuse atrophy (Figure 1C). The patient was prescribed new polycarbonate lenses and counseled about monocular precautions.

**Discussion**

NAION secondary to APAC is a rare clinical entity that can result in severe vision loss. NAION is caused by ischemia to the optic nerve head, mainly from the posterior ciliary artery and its branches.11 Blood flow to the optic nerve head is determined by
Figure 2. OCT of the optic nerve and macula. (A) Retinal nerve fiber layer analysis demonstrates 360° optic nerve edema in the right eye and no optic nerve edema in the left. (B) Macula OCT demonstrates a small area of subretinal fluid extending from the temporal portion of the optic disc into the nasal macula in the right eye.

Abbreviations: ONH, optic nerve head; RNFL, retinal nerve fiber layer; OU, both eyes; SUP, superior; NAS, nasal; TEMP, temporal; OD, right eye; OS, left eye; OCT, optical coherence tomography; CD, cup to disc.
a ratio of perfusion pressure and resistance to flow. Resistance to flow is determined by efficiency of autoregulation, vascular integrity, and the rheological properties of blood. These factors can be impaired in patients with diabetes, hypertension, and atherosclerosis. Perfusion pressure is determined by the difference in mean arterial pressure (MAP) and IOP. The MAP is reduced with relative nocturnal hypotension and antihypertensive medications. High IOPs during phacoemulsification may contribute to the increased incidence of NAION after cataract surgery. Elevated IOP has been identified as a cause of decreased blood flow to the optic nerve head in five cases (six eyes) previously reported by other authors and our case of NAION after APAC (Table 1).

Taken together with our case, of all seven eyes with APAC-associated NAION, the duration of APAC symptoms ranged from 1 day to 1 month. Five of the seven eyes had NAION at presentation, while the remainder developed NAION one week after presentation for APAC. The IOP at presentation ranged from 40–56 mmHg. Five of seven eyes underwent LPI in the NAION eye and contralateral eye. Slavin and Margulis described one patient who developed an NAION one week after an APAC attack in the left eye and refused an LPI in the right eye. One month later, the patient had an APAC attack in the right eye and developed a right NAION a week later. Sequential NAION has not been reported in the other APAC-associated NAION cases.

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**Figure 3** Stimulus V 30-2 Humphrey visual field demonstrates a cecocentral scotoma in the right eye.

**Abbreviations:** POS, positive; NEG, negative; dB, decibels.
1 week delay in presentation with NAION after presentation with APAC in this patient is that the initial APAC event may have started the cycle of NAION, but it was not clinically significant for 1 week. Hayreh described the cycle of NAION as ischemia of axons resulting in axoplasmic flow stasis, which in turn cause axoplasmatic accumulation and subsequent axonal swelling, which in turn compresses the capillaries, leading to more ischemia.12

Including our patient, only two of the seven eyes (29%) with APAC-associated NAION attained acuity of 20/30 or better,5–9 which is worse than the 49% of NAION patients who attained 20/30 or better in a study by Hayreh et al.16 However, the low sample size of APAC-associated NAION patients is a barrier for meaningful comparisons with other NAION patients. It is also possible that the 2 cases of the delayed onset NAION after APAC previously reported may be spontaneous NAION.

Patients with APAC can also develop mild optic nerve edema that is not associated with NAION.17 Tsai et al found a statistically significant difference in the retinal nerve fiber layer thickness (RNFL) of the APAC eye and the contralateral eye measured by OCT 1 week after the APAC event.17 However, a statistically significant difference was not observed at the 4- and 12-week follow-ups.17 These patients can be differentiated from NAION secondary to APAC patients by the lack of subsequent optic nerve atrophy and preserved visual acuity and visual field.

NAION secondary to APAC is a rare but potentially blinding clinical entity. In addition to counseling patients about NAION risk factors (eg, evening antihypertensive medications and control of hypertension, diabetes, and dyslipidemia), patients must be counseled on the importance of performing an LPI in the contralateral eye. In APAC patients with an RAPD, a large amount of optic nerve edema, persistent decreased visual acuity, and a small cup-to-disc ratio in the contralateral eye should raise clinical suspicion for NAION.

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

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Table 1 Summary of the five previously reported patients with NAION-associated APAC (six eyes) and our additional patient

| Paper                        | Age | Eye | Symptom duration | Time to NAION | Initial VA | IOP | APD | C/D | Final VA | VF defect |
|------------------------------|-----|-----|------------------|---------------|------------|-----|-----|-----|----------|-----------|
| Sonty and Schwartz,6         | 66  | OS  | I month, intermittent | Presentation  | 20/100     | 55  | NR  | NR  | 20/50    | Nasal     |
| Slavin and Margulis,8        | 70  | OD  | I day            | I week        | 20/200     | 56  | NR  | 0.1 | 20/200   | Central scotoma and inferior |
|                             |     | OS  | I day            | I week        | HM         | 50  | NR  | 0.1 | 2/200    | Central scotoma and inferior |
| Yip et al,4                  | 64  | OS  | 5 day            | Presentation  | HM         | 54  | Y   | NR  | HM       | NR        |
| Nahum et al,7                | 59  | OS  | NR               | Presentation  | 20/30      | 40  |     |     |          | Superior   |
| Choudhari et al,9            | 59  | OS  | 1 week           | Presentation  | 20/120     | 46  | Y   | NR  |          | Constricted |
| Kuriyan and Lam,2013         | 50  | OD  | I day            | Presentation  | HM         | 41  | Y   | <0.1| HM       | Cenecentral scotoma, peripheral constriction |

Abbreviations: OS, left eye; OD, right eye; NAION, non-arteritic ischemic optic neuropathy; APAC, acute primary-angle closure; VA, visual acuity; IOP, intraocular pressure; APD, afferent pupillary defect; C/D, cup to disc ratio; VF, visual field; NR, not reported; HM, hand motion; Y, yes.
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