Scleritis is an inflammatory process involving the outer coating of the globe which is characterized by focal or diffuse hyperemia, moderate to severe pain, and frequent impairment of vision. Most cases of scleritis are autoimmune in nature and are managed with topical and/or systemic corticosteroids. Infectious scleritis is a less common entity, occurring in 5%–10% of cases, and requiring directed antimicrobial therapy. We present a case of *Nocardia farcinica* anterior nodular scleritis diagnosed via positive culture of an excisional biopsy of a scleral nodule. The patient improved after combined surgical and medical therapy with amoxicillin-clavulanate and moxifloxacin for 12 months. Based on a literature review, a summary of reported cases of infectious scleritis is provided, and guidelines pertaining to diagnosis and management are offered.

**Keywords.** infectious scleritis; infection; scleritis; nodular scleritis; anterior scleritis.

## CASE REPORT

A previously healthy 59-year-old male presented to his ophthalmologist in October 2015 with acute onset left eye pressure and pain, followed by the development of erythema along the medial aspect of the eye. He denied any preceding injury or trauma to the eye, foreign body sensation, or prior ocular surgery. There were no associated systemic symptoms such as fevers, weight loss, rash, ulcers, or musculoskeletal complaints. He did relate that he was involved in a home renovation project for the previous several months that entailed tearing down walls. He also reported that he had been using a leaf blower in his yard quite frequently over the past several weeks, but he did not recall getting any debris in his eye. During those activities, he did not wear any eye protection. Ophthalmologic examination of the left eye revealed nasal injection from 6 o’clock to 11 o’clock with chemosis and a small nodule on the medial aspect of the globe. He was diagnosed with episcleritis of the left medial eye, and therapy with topical prednisone drops was initiated. Seven days later, he returned to his ophthalmologist with minimal resolution of his symptoms and new onset blurry vision. At that time, topical prednisone was discontinued, and ibuprofen 800 mg 3 times a day was initiated. He returned 2 days later for follow-up, at which time oral prednisone 40 mg daily for 1 week was initiated due to poor response to the ibuprofen.

The prednisone was subsequently increased to 80 mg daily due to lack of improvement, and he was referred to Wake Forest Baptist Medical Center for specialty consultation in ophthalmology. The preliminary diagnostic impression was episcleritis with possible evolving anterior nodular scleritis. His ocular examination revealed an erythematous and elevated nodule with associated scleral injection of the left eye (Figure 1). There was no associated keratitis or anterior uveitis. Serologies were obtained to rule out infectious and inflammatory etiologies and included rapid plasma reagin, interferon gamma release assay, antinuclear antibody, and rheumatoid factor, all of which were negative or nonreactive. Screening for HIV was not done due to lack of risk factors. Computed tomography of the left orbit revealed focal thickening of the left anteromedial sclera with extension of inflammation into the contiguous vitreous that measured 6 × 9 mm (Figure 2). An aspiration biopsy of the nodule for cultures was performed, and the patient was admitted to the hospital for empiric antimicrobial therapy. Infectious diseases was consulted and recommended empiric treatment with intravenous vancomycin, cefepime, and voriconazole. The aspirate from the nodule subsequently grew only diphtheroids on routine culture, the significance of which was unclear. After 5 days of inpatient parenteral antimicrobial therapy with marginal improvement in local inflammatory signs, the patient was discharged on ongoing empiric oral therapy with linezolid, ciprofloxacin, and voriconazole. He then returned to the ophthalmology clinic 2 days later, at which time he underwent excisional biopsy of the scleral nodule and injection of intrascleral moxifloxacin. Per observations at surgery, the nodule was characterized as an abscess that contained a small amount of purulent fluid. Gram stain, fungal smear, and acid-fast smear of the fluid were negative. Cytopathology revealed only acute and chronic inflammation with reactive changes; no organisms were seen. Therapy with oral linezolid, ciprofloxacin,
and voriconazole was continued while final culture data were pending. Subsequently, 2 of 3 acid-fast cultures from the biopsy grew *Nocardia farcinica*. The specimen was then sent to the University of Texas Health Sciences Center microbiology lab for susceptibility testing. Monotherapy with linezolid was initially maintained while awaiting minimum inhibitory concentration data. Treatment was then changed to oral amoxicillin–clavulanate and moxifloxacin for a total of 12 months based on the susceptibility. Trimethoprim–sulfamethoxazole was not utilized as the patient previously had a major allergy to that drug. With ongoing antimicrobial therapy, the patient’s ocular inflammation slowly improved. At his 12-month follow-up visit, all of the patient’s symptoms had resolved, and his ocular exam findings had normalized, aside from some persistent scleral thinning at the infection site.

**DISCUSSION**

Infectious scleritis (IS) is an entity not well described outside of the ophthalmologic literature, where it is noted to occur in about 5%–18% of scleritis cases [1–3]. Physicians trained in infectious diseases are generally well versed in the manifestations and causes of infection at most sites in the human host; the eye and associated ocular infections are frequently an area of less familiarity. As such, input from and collaboration with colleagues in ophthalmology are critical if management of eye infections is to be successful.

Scleral inflammatory disorders of the eye can be classified as episcleritis, in which inflammation is superficial, and scleritis, characterized by location as anterior or posterior, which is associated with deeper inflammation that is more destructive [4, 5]. In patients with scleritis, the nature of the scleral inflammatory process is often described as diffuse or nodular and may be necrotizing or non-necrotizing, with each of those descriptions providing some degree of etiologic insight [2]. Scleritis is most often associated with autoimmune or connective tissue disorders but may also be idiopathic or infectious in etiology. Clinically, autoimmune scleritis may be indistinguishable from IS, which creates challenges in the selection of appropriate therapy [2, 6–8]. If initially treated with immunosuppressive or anti-inflammatory therapies, IS can result in a number of vision-threatening complications such as cataract, glaucoma, endophthalmitis, vision loss, and loss of the globe [6, 9].

Clinical presentation of IS includes redness, pain, and tearing of the affected eye [2, 7]. Discomfort is often mild to moderate in the setting of nodular scleritis, whereas pain is frequently severe with necrotizing scleritis [2]. If pain is out of proportion to exam findings, one should consider an infectious etiology [2]. Time to presentation of infection varies from days to years and has been reported to occur up to 30 years after an injury [10, 11]. In patients with nodular IS, nodules can be solitary or multifocal and often appear to be yellow colored under the conjunctiva [2, 7]. The presence of multiple nodules has not been associated with a specific pathogen, but rather may be more related to factors such as the burden of infection [12]. Vision may be normal, but patients frequently present with diminished visual acuity [7].

Bacteria, fungi, parasites, and viruses have all been reported as etiologies of IS [2, 7, 8, 10], but bacterial infection is by far most common, with a reported range of 53%–100% [2, 7, 10]. In case series of patients with IS (Table 1), *P. aeruginosa* is etiologic in about 25% of patients, whereas all pseudomonas species account for about 40% of cases of IS (Table 1). Although fungal infection is rare, it appears to be most common in climates that are hot or humid, theorized to be due to the heavier spore load in the environment [7]. Bacterial infection tends to present more acutely, whereas infections due to fungi, *Nocardia*, and mycobacteria are more indolent in onset and presentation [2, 10]. Time to presentation varies and can be months to decades after an inciting event [2, 9, 13].

Diagnosis of IS can be difficult to achieve as the infection presents very similarly to autoimmune scleritis, which is the most common form of scleritis [6, 8, 10]. History is often a
Tissue culture, directed antimicrobial therapy can be initiated.

Treatment recommendations for infectious scleritis have not been well established. Initial management with intensive broad-spectrum antibiotics to include coverage for *P. aeruginosa* and gram-positive organisms with combined topical and systemic modalities has been recommended [1, 6]. In cases of trauma with organic material, empiric antifungal therapy with agents such as voriconazole has been suggested [11]. Early surgical debridement has also been recommended as debridement and debulking of necrotic tissue may facilitate antibiotic access [1, 6, 7, 11, 13].

Given the nature of infecting pathogens, time to diagnosis, and poor penetration of antimicrobial therapies into the avascular sclera, multiple surgical interventions may be required [7, 13]. The duration of antimicrobial therapy is poorly defined, though reference texts suggest that it should be “prolonged” [14].

Worse visual outcomes are associated with poor visual acuity at the time of presentation, medical therapy alone, and fungal infections, but they are not significantly associated with any particular organism or the presence of multifocal abscesses [1, 10, 12].

Bacterial biofilm formation is theorized to be responsible for cases of IS refractory to medical management alone [9]. Surgical intervention has been associated with better outcomes in regards to higher rates of globe preservation and improved prognosis, however, not better visual outcomes [9, 10, 12]. Hodson et al. report that only 18% of patients treated with medical therapy alone for a prolonged duration had adequate treatment, with the majority requiring surgery [2, 10]. For the best outcomes, surgical intervention in conjunction with medical therapy has been recommended [7, 9, 11, 13].

### CONCLUSION

IS should be considered in patients presenting with a painful unilateral red eye preceded by a history of ocular surgery or trauma. Often nodules or ulcers are present in the affected eye. Infectious etiologies include typical bacteria such as *P. aeruginosa* and gram-positive organisms, and fungal pathogens such as *Aspergillus*, *Fusarium*, *Pseudallescheria*, and *Scedosporium* [2, 7]. Debridement of the infected nodule or ulcer is recommended for tissue culture to establish a definitive diagnosis and may result in improved outcomes (Table 1).

It would be reasonable to initiate empiric therapy with antimicrobials such as ciprofloxacin and linezolid that would cover *P. aeruginosa*, gram-positive organisms, and *Nocardia* until tailored therapy based on culture results is feasible. Those

### Table 1. Summary of Reported Cases of Infectious Scleritis Including Demographics, Risk Factors, Treatment, Outcomes, and Microbiology [4, 5, 7, 8, 9, 10, 12, 13]

| No. (%) | Total, No. eyes | 224 |
|--------|-----------------|-----|
| Male   | 127 (56.6)      |     |
| Female | 96 (42.8)       |     |
| Age range, y | 12–89         |     |
| Risk factors |               |     |
| Trauma | 31 (13.8)       |     |
| Surgery| 170 (75.8)      |     |
| Treatment |                |     |
| Topical antibiotics | 221 (98.6)   |     |
| Systemic antibiotics | 154 (68.7)   |     |
| Intraocular antibiotics | 68 (30.4)   |     |
| Surgery | 188 (83.9)      |     |
| Outcomes |                |     |
| Resolved (VA > 20/200) | 93 (41.5)   |     |
| Recurred | 6 (0.03)       |     |
| Microbiology | 217 (96.8)   |     |
| *Pseudomonas aeruginosa* | 57 (25.4)   |     |
| *Pseudomonas* spp. | 28 (12.5)   |     |
| Other gram-negative bacilli | 16 (7.1)    |     |
| *Staphylococcus aureus* | 16 (7.1)    |     |
| *Streptococcus pneumoniae* | 7 (3.1)     |     |
| *Streptococcus* spp. | 8 (3.50)   |     |
| *Enterococcus* spp. | 4 (0.9)    |     |
| Gram-positive rods | 6 (2.7)     |     |
| *Aspergillus* spp. | 9 (4.0)    |     |
| *Fusarium* spp. | 5 (2.2)    |     |
| *Scedosporium* | 3 (1.3)   |     |
| Other fungi | 18 (8.0)    |     |
| *Nocardia* spp. | 18 (8.0)   |     |
| Nontuberculous mycobacteria | 14 (6.25)  |     |
| *M. tuberculosis* | 1 (0.4)    |     |

Abbreviation: VA, visual acuity.

Visual acuity < 20/200 confers legal blindness [15].

Klebsiella, Stenotrophomonas, Eikenella, Enterobacter, Chryseobacterium, *H. influenzae*, Citrobacter, Alcaligenes, Serratia.

Corynebacterium, Propionibacterium, Brevobacterium.

Paecilomyces, Candida, Colletotrichum, Cladosporium, Acremonium, Curvularia, Rhizoctonia.
patients with trauma, particularly involving organic material, could potentially be started on empiric antifungal therapy with an agent such as voriconazole. Duration of therapy has not been established for infectious scleritis, and further studies are needed to define the appropriate duration of medical therapy depending on the etiology of the infection and the presence or absence of surgical intervention.

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