Clinical Ophthalmology

Purpose: To evaluate the impact of intravitreal bevacizumab combined with laser vascular ablation in the management of advanced Coats’ disease presenting with exudative retinal detachment.

Methods: This was a retrospective review of 24 children that presented with exudative retinal detachments associated with advanced Coats’ disease. Mean patient age was 62 months (range 9–160 months). Presenting signs included retinal detachment in 24 children (100%), vascular telangiectasia in 24 children (100%), and retinal ischemia in 24 children (100%). Twenty of 24 children presented with elevated, vascular leakage in the fovea (83%). Two children presented with sub-retinal fibrosis associated with presumed long-standing retinal detachment without evidence of rhegmatogenous retinal detachment. Ten patients exhibited vascular alterations in the periphery of the second eye without clinical evidence of exudation. All 24 children were treated with a large-spot-size diode laser directly to areas of abnormal telangiectatic vasculature. All 24 children received intravitreal bevacizumab injection.

Results: All 24 children had resolution of exudative retinal detachment, ablation of vascular telangiectasia, and anatomic improvement of the retina. No child exhibited progressive retinal detachment and no eye required enucleation. No cases of neovascular glaucoma were seen. Fellow eyes with peripheral vascular alterations showed no progression to exudative vasculopathy during the observation period. Intravitreal bevacizumab injection was not associated with endophthalmitis or systemically-observed complications.

Conclusion: Repetitive intravitreal bevacizumab combined with laser vascular ablation may be utilized effectively for advanced Coats’ disease presenting with exudative retinal detachment.

Keywords: Coats’ disease, bevacizumab, anti-VEGF, laser ablation, retina

Introduction
Coats’ disease was first described in 1908 by Scottish ophthalmologist Georges Coats.1 This entity was characterized by unilateral retinal vascular abnormalities and retinal exudation in young males. In 1912, Leber described a disease with similar telangiectatic changes but without the massive exudation described by Coats.2 Reese merged the two diseases in a single spectrum that started as Leber disease but progressed to Coats’ disease during long-term follow-up.3

Treatment of Coats’ disease depends on the stage of the disease. Mild cases can be observed.4 However, if exudation, retinal, or subretinal fluid is present, treatment with laser ablation or cryotherapy is necessary to areas of retinal telangiectasia and retinal nonperfusion.5 Surgery may be needed in cases of nonclearing vitreous hemorrhage or severe retinal detachment.

Intravitreal anti-vascular endothelial growth factor (VEGF) has been reported to be effective as adjunctive therapy to reduce subretinal fluid and exudation both alone
and in combination with other treatment modalities.\textsuperscript{6–10} The purpose of our study is to evaluate the impact of repetitive intravitreal bevacizumab combined with laser vascular ablation in the management of advanced Coats’ disease presenting with exudative retinal detachment.

**Methods**

A retrospective review of all patients with Coats’ disease diagnosed and treated with intravitreal bevacizumab by the same senior clinician (TGM) was conducted. Patient age, sex, date of presentation, date of last follow-up, number of laser and bevacizumab application sessions, presenting stage of Coats’ disease, and final stage of Coats’ disease were recorded. Exclusion criteria for treatment included patients whose social situations prevented them from receiving ongoing care, and patients who presented with no light perception, and thus were observed without receiving laser treatment.

Patients with advanced Coats’ disease were treated with laser ablation applied at each clinical visit to all areas of retinal telangiectasis and/or exudative retinal detachment (Figures 1 and 2). Patients underwent examinations under anesthesia approximately every 2 months while treatment was continued. Laser treatments were continued until either complete anatomic resolution of exudation and telangiectasis was achieved or until neovascular glaucoma and no light perception vision developed and treatment was no longer deemed to be helpful.

Laser therapy was delivered while patients were under anesthesia using the 810 nm iris diode laser (IRIDEX Corporation, Mountain View, CA, USA). The 1.2 mm spot size was aimed through a dilated pupil using the indirect ophthalmoscope and a 20-diopter lens. The laser was generally set at a power of 350 mW. The entire surface of telangiectatic vessels and/or exudative detachment was treated. The clinical endpoint was the complete whitening of the telangiectatic vessels. Treatment was also applied to the avascular retinal areas peripheral to telangiectatic vessels. Intravitreal bevacizumab 1.25 mg was injected in the pars plana consecutively in all patients.

**Results**

Twenty-four children presented with exudative retinal detachment associated with advanced Coats’ disease. Mean patient age was 62 months (range 9–160 months). Seventeen children were boys (71%), and 7 were girls. Presenting signs included retinal detachment in 24 children (100%), vascular telangiectasia in 24 children (100%), and retinal ischemia in 24 children (100%). Twenty of 24 children presented with elevated, vascular leakage in the fovea (83%). Two children presented with sub-retinal fibrosis associated with presumed long-standing retinal detachment without evidence of rhegmatogenous retinal detachment. Ten patients exhibited vascular alterations in the periphery of the second eye without clinical evidence of exudation. All 24 children were treated with a large-spot-size diode laser in areas of abnormal

![Figure 1](https://www.dovepress.com/)

**Figure 1** The patient has a total (fovea-involving) exudative retinal detachment.  
**Notes:** Fundus photograph of patient one at presentation (A). After treatment (B).
telangiectatic vasculature. All 24 children received intravitreal bevacizumab injection. Mean number of laser treatment sessions was 7.4 (range 3–15 sessions) while mean number of intravitreal injections was 4.6 (range 2–9 injections). Mean patient follow-up was 22.4 months (range 6–36 months). All 24 children had resolution of exudative retinal detachment, ablation of vascular telangiectasia, and anatomic improvement of the retina. No child exhibited progressive retinal detachment and no eye required enucleation. No cases of neovascular glaucoma were seen. Fellow eyes with peripheral vascular alterations showed no progression to exudative vasculopathy during the observation period. Intravitreal bevacizumab injection was not associated with endophthalmitis or systemically-observed complications.

Discussion
The treatment of Coats’ disease has included the multiple modalities of observation, laser, cryotherapy, surgical management, and enucleation in cases of painful blind eyes. The mainstay of treatment remains laser ablation.

A recent study by Zhao et al has shown that the VEGF concentrations correlated with the severity of Coats’ disease. VEGF concentrations significantly correlated with extent of exudative retinal detachment. Other authors have also found elevated levels of VEGF in patients with Coats’ disease. Marked reduction in VEGF levels and improvement of exudation and retinal detachment have been seen in association with treatment with anti-VEGF agents.

In most available case reports of Coats’ disease treated with bevacizumab, VEGF appears to be well-tolerated. However, case numbers are small with limited follow-up. In our study, all patients had resolution of retinal detachment and anatomic improvement of the retina. Follow-up was unremarkable and without complications.

Anti-VEGF appears to be safe. However, a retrospective analysis of eight patients with Coats’ disease manifesting total or partial exudative retinal detachment where the retinal telangiectasia was treated with standard laser photocoagulation and/or cryotherapy plus additional intravitreal bevacizumab showed that treatment with intravitreal bevacizumab in addition to standard therapy can be associated with vitreoretinal fibrosis. Longer follow-up and larger studies will be necessary to establish safety of bevacizumab in Coats’ disease.

Weaknesses in this study include small sample size and nonrandomization. Additionally, achieving statistical superiority of laser ablation plus intravitreal bevacizumab over laser ablation alone carries intrinsic difficulties due to the rare nature of this disease. Furthermore, several authors have reported exudative recurrences after 10 years of initial laser ablation, making long-term assessment difficult. Extended serial follow-up will be important for these patients to ensure that good outcomes can be sustained over time.

Conclusion
The goal of this study was to evaluate the efficacy of repetitive intravitreal bevacizumab combined with laser vascular ablation in the management of advanced Coats’ disease presenting with exudative retinal detachment. We found that this approach resulted in resolution of retinal detachment in
all cases. Although this is a small patient series, our study results are promising and should be explored further with an enlarged cohort. We advise against intravitreal injection in all cases in which the diagnosis of retinoblastoma is uncertain.

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

References