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Idiopathic Epiretinal Membrane and Vitreomacular Traction Preferred Practice Pattern®

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We would like to acknowledge the role of Andre Ambrus, MLIS, in the initial revisions of the Retina/Vitreous PPPs and the first meeting of the Retina/Vitreous PPP Committee.

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Preferred Practice Pattern[®] guidelines are developed by the Academy's H. Dunbar Hoskins Jr., MD Center for Quality Eye Care without any external financial support. Authors and reviewers of the guidelines are volunteers and do not receive any financial compensation for their contributions to the documents. The guidelines are externally reviewed by experts and stakeholders before publication.

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RETINA/VITREOUS PREFERRED PRACTICE PATTERN® DEVELOPMENT PROCESS AND PARTICIPANTS

The **Retina/Vitreous Preferred Practice Pattern Committee** members wrote the Idiopathic Epiretinal Membrane and Vitreomacular Traction Preferred Practice Pattern (PPP) guidelines. The Retina/Vitreous PPP Committee members discussed and reviewed successive drafts of the document, meeting in person twice and conducting other review by e-mail discussion, to develop a consensus over the final version of the document.

Retina/Vitreous Preferred Practice Pattern Committee 2023–2024

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We thank our partners, the Cochrane Eyes and Vision US Satellite (CEV@US), for identifying reliable systematic reviews that we cite and discuss in support of the PPP recommendations.

The **Preferred Practice Patterns Committee** members reviewed and discussed the document during a meeting in June 2024. The document was edited in response to the discussion and comments.

Preferred Practice Patterns Committee 2024

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The Idiopathic Epiretinal Membrane and Vitreomacular Traction PPP was sent for review in July 2024 to improve the quality of the guideline, to gather feedback on the draft recommendations and to assess feasibility for and applicability to the target audience, including assessing the facilitators and barriers to implementing recommendations (e.g., U.S. ophthalmologists and other important groups, including patients, other physicians, international ophthalmologists, research organizations, ophthalmological organizations, and experts in the field). The PPP was sent for review to the following patient organizations to solicit the views and preferences of patients and the public: Consumers United for Evidence-Based Healthcare, American Foundation for the Blind, Foundation Fighting Blindness, Lighthouse Guild, National Federation of the Blind, and Prevent Blindness. All those returning comments were required to provide disclosure of relevant relationships with industry to have their comments considered (indicated with an asterisk below). Members of the Retina/Vitreous Preferred Practice Pattern Committee reviewed these comments and determined revisions to the document.

<u>Academy Reviewers</u> Board of Trustees and Committee of Secretaries* Council* General Counsel* Ophthalmic Technology Assessment Committee Retina/Vitreous Panel Basic and Clinical Science Course Section 12 Subcommittee* Practicing Ophthalmologists Advisory Committee for Education*

Invited Reviewers

American College of Surgeons, Advisory Council for Ophthalmic Surgery American Foundation for the Blind American Ophthalmological Society* American Society of Retina Specialists* American Uveitis Society* Association for Research in Vision and Ophthalmology Association of University Professors in Ophthalmology Canadian Ophthalmological Society Consumers United for Evidence-Based Health Care Foundation Fighting Blindness International Council of Ophthalmology Lighthouse Guild Macula Society* National Eye Institute National Federation of the Blind National Medical Association, Ophthalmology Section North American Neuro-Ophthalmology Society* Prevent Blindness* Retina Society Women in Ophthalmology* Brittni Ashton Scruggs, MD Alexis Warren, MD* Steven Yeh, MD

This guideline will be formally re-evaluated and updated on a 5-year cycle in 2029. A Summary Benchmark is a resource to facilitate application of the guideline and to provide criteria that could be used to measure the application of recommendations, which will be available to all at www.aao.org/ppp.

FINANCIAL DISCLOSURES

There is no external funding, including industry/commercial support, for the development of this PPP or for the distribution of the guidelines. The Academy has fully funded the development of this PPP, and the views or interests of the Academy have not influenced the final recommendations, which are based on evidence from systematic reviews. All those individuals significantly involved in the guideline development process, including Retina/Vitreous PPP Committee members, PPP Committee members, Secretary for Quality of Care, and Academy staff, have declared competing/financial interests through a financial interest disclosure process as well as an assessment of the Open Payments website (available at https://openpaymentsdata.cms.gov/). The interests of the Retina/Vitreous PPP Committee members are provided at the beginning of each meeting and those with competing interests in a guideline topic do not participate in voting on areas of disagreement. In compliance with the Council of Medical Specialty Societies' Code for Interactions with Companies (available at https://cmss.org/code-for-interactions-with-companies/), relevant relationships with industry are listed. As per CMSS code, direct financial relationships with companies do not include food and beverage, research funds paid to the institution and relationships outside of the topic of the PPP. The Academy has Relationship with Industry Procedures to comply with the Code (available at www.aao.org/aboutpreferred-practice-patterns). A majority (86%) of the members of the Retina/Vitreous PPP Committee 2023–2024 had no financial relationship to disclose.

Retina/Vitreous Preferred Practice Pattern Committee 2023–2024

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The disclosures of relevant relationships to industry of other reviewers of the document from January to October 2024 are available online at <u>www.aao.org/ppp</u>.

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OBJECTIVES OF PREFERRED PRACTICE PATTERN® GUIDELINES

As a service to its members and the public, the American Academy of Ophthalmology has developed a series of Preferred Practice Pattern guidelines that **identify characteristics and components of quality eye care.** Appendix 1 describes the core criteria of quality eye care.

The Preferred Practice Pattern guidelines are based on the best available scientific data as interpreted by committees of knowledgeable health professionals. In some instances, such as when results of carefully conducted clinical trials are available, the data are particularly persuasive and provide clear guidance. In other instances, the committees have to rely on their collective judgment and evaluation of available evidence.

These documents provide guidance for the pattern of practice, not for the care of a particular individual. While they should generally meet the needs of most patients, they cannot possibly best meet the needs of all patients. Adherence to these PPPs will not ensure a successful outcome in every situation. These practice patterns should not be deemed inclusive of all proper methods of care or exclusive of other methods of care reasonably directed at obtaining the best results. It may be necessary to approach different patients' needs in different ways. The physician must make the ultimate judgment about the propriety of the care of a particular patient in light of all of the circumstances presented by that patient. The American Academy of Ophthalmology is available to assist members in resolving ethical dilemmas that arise in the course of ophthalmic practice.

Preferred Practice Pattern guidelines are not medical standards to be adhered to in all individual situations. The Academy specifically disclaims any and all liability for injury or other damages of any kind, from negligence or otherwise, for any and all claims that may arise out of the use of any recommendations or other information contained herein.

References to certain drugs, instruments, and other products are made for illustrative purposes only and are not intended to constitute an endorsement of such. Such material may include information on applications that are not considered community standard, that reflect indications not included in approved US Food and Drug Administration (FDA) labeling, or that are approved for use only in restricted research settings. The FDA has stated that it is the responsibility of the physician to determine the FDA status of each drug or device he or she wishes to use, and to use them with appropriate patient consent in compliance with applicable law.

Innovation in medicine is essential to ensure the future health of the American public, and the Academy encourages the development of new diagnostic and therapeutic methods that will improve eye care. It is essential to recognize that true medical excellence is achieved only when the patients' needs are the foremost consideration.

All Preferred Practice Pattern guidelines are reviewed by their parent committee annually or earlier if developments warrant and updated accordingly. To ensure that all PPPs are current, each is valid for 5 years from the approved by date unless superseded by a revision. Preferred Practice Pattern guidelines are funded by the Academy without commercial support. Authors and reviewers of PPPs are volunteers and do not receive any financial compensation for their contributions to the documents. The PPPs are externally reviewed by experts and stakeholders, including consumer representatives, before publication. The PPPs are developed in compliance with the Council of Medical Specialty Societies' Code for Interactions with Companies. The Academy has Relationship with Industry Procedures (available at <u>www.aao.org/about-preferred-practice-patterns</u>) to comply with the Code.

Appendix 2 contains the International Statistical Classification of Diseases and Related Health Problems (ICD) codes for the disease entities that this PPP covers. The intended users of the Diabetic Retinopathy PPP are ophthalmologists.

METHODS AND KEY TO RATINGS

Preferred Practice Pattern[®] guidelines should be clinically relevant and specific enough to provide useful information to practitioners. Where evidence exists to support a recommendation for care, the recommendation should be given an explicit rating that shows the strength of evidence. To accomplish these aims, methods from the Scottish Intercollegiate Guideline Network¹ (SIGN) and the Grading of Recommendations Assessment, Development and Evaluation² (GRADE) group are used. GRADE is a systematic approach to grading the strength of the total body of evidence that is available to support recommendations on a specific clinical management issue. Organizations that have adopted GRADE include SIGN, the World Health Organization, the Agency for Healthcare Research and Policy, and the American College of Physicians.³

- All studies used to form a recommendation for care are graded for strength of evidence individually, and that grade is listed with the study citation.
- To rate individual studies, a scale based on SIGN¹ is used. The definitions and levels of evidence to rate individual studies are as follows:

I++	High-quality meta-analyses, systematic reviews of randomized controlled trials (RCTs), or RCTs with a very low risk of bias
I+	Well-conducted meta-analyses, systematic reviews of RCTs, or RCTs with a low risk of bias
I-	Meta-analyses, systematic reviews of RCTs, or RCTs with a high risk of bias
II++	High-quality systematic reviews of case-control or cohort studies High-quality case-control or cohort studies with a very low risk of confounding or bias and a high probability that the relationship is causal
II+	Well-conducted case-control or cohort studies with a low risk of confounding or bias and a moderate probability that the relationship is causal
II-	Case-control or cohort studies with a high risk of confounding or bias and a significant risk that the relationship is not causal
III	Nonanalytic studies (e.g., case reports, case series)

• Recommendations for care are formed based on the body of the evidence. The body of evidence quality ratings are defined by GRADE² as follows:

Good quality	Further research is very unlikely to change our confidence in the estimate of effect
Moderate quality	Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate
Insufficient quality	Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate Any estimate of effect is very uncertain

• Key recommendations for care are defined by GRADE² as follows:

Strong recommendation	Used when the desirable effects of an intervention clearly outweigh the undesirable effects or clearly do not
Discretionary recommendation	Used when the trade-offs are less certain—either because of low-quality evidence or because evidence suggests that desirable and undesirable effects are closely balanced

- The Highlighted Findings and Recommendations for Care section lists points determined by the Retina/Vitreous PPP Committee to be of particular importance to vision and quality of life outcomes.
- Recommendations for care in this PPP were rated using the system described above. Ratings are embedded throughout the PPP main text in italics.
- Literature searches to update the PPP were undertaken on March 6, 2023, January 23, 2024, and August 6, 2024 in PubMed. Complete details of the literature searches are available online at <u>www.aao.org/ppp</u>.
- Relevant systematic reviews were identified by the Cochrane Eyes and Vision US Satellite (CEV@US). These systematic reviews were screened by the committee and rated using the system described above by the committee methodologist.

• Recommendations are based on systematic reviews, as per the Institute of Medicine (Clinical Practice Guidelines We Can Trust, 2011). In formulating the recommendations, the health benefits, side effects/harms/risks, and the balance of benefits and risks are reviewed and considered. Final decisions are arrived at through informal consensus techniques. If there are areas of disagreement, a vote will be conducted among the members of the Retina/Vitreous PPP Committee. If there are individuals with direct financial relationships in the area of disagreement, these individuals will refrain from the vote.

HIGHLIGHTED FINDINGS AND RECOMMENDATIONS FOR CARE

Risk factors for epiretinal membrane (ERM) include increasing age and various retinal pathologies, such as posterior vitreous detachment (PVD), uveitis, retinal breaks, retinal vein occlusions, diabetic retinopathy, and ocular inflammatory diseases.

The majority of ERMs will remain relatively stable and do not require therapy. In patients who have areas of vitreomacular traction (VMT) of 1500 μ m or less, the incidence of spontaneous release of traction from the macula occurs in approximately 30% to 40% of eyes over a follow-up of 1 to 2 years.

Optical coherence tomography (OCT) is a highly sensitive test and is routinely used to diagnose and characterize ERM, VMT, and associated retinal changes.

Vitrectomy surgery is often indicated in affected patients who have a decrease in visual acuity, metamorphopsia, double vision, or difficulty using their eyes together. Vitrectomy for ERM or VMT usually improves metamorphopsia and visual acuity. On average, approximately 80% of patients with ERM or VMT will improve by at least 2 lines of visual acuity following vitrectomy surgery.

INTRODUCTION

DISEASE DEFINITION

Epiretinal membranes (ERMs) are sheet-like structures that develop on the inner surface of the neurosensory retina. Vitreomacular adhesion (VMA) is an attachment of the posterior cortical vitreous to the macula without resultant traction. Vitreomacular traction (VMT) occurs when the posterior cortical vitreous partially separates from the retina, yet some areas of adhesion remain that exert tractional forces on the neurosensory retina. Thickening, distortion, intraretinal cystoid changes, macular hole, and even subretinal fluid in the macula can result from the VMT.⁴ The macular changes that result from either ERM or VMT lead to similar symptoms: reduced visual acuity, metamorphopsia, difficulty using both eyes together, and diplopia.

PATIENT POPULATION

The patient population is predominately adults.

CLINICAL OBJECTIVES

- Describe the pathogenesis of ERM and VMT
- Recognize symptoms and signs of ERM and VMT
- Describe the natural history without treatment
- Propose a treatment strategy
- Educate the patient about treatment options
- Optimize visual function and/or relief of symptoms

BACKGROUND

Epiretinal membranes consist of fibrocellular proliferation on the surface of the neurosensory retina, with or without wrinkling of the retina. They comprise reactive cellular elements, vitreous structures, and fibrotic components.⁵ Idiopathic ERMs do not have a clearly identifiable cause.⁶

Secondary ERMs may occur after retinal breaks or detachments, or following intraocular surgery, trauma, or retinal laser surgery or cryotherapy treatment.⁵ An ERM is likely due to reactive wound healing and is associated with a proliferation of either reactive retinal pigment epithelial cells and/or retinal glial cells. Epiretinal membranes are also common in eyes with retinal vascular disease^{7, 8} (e.g., diabetic retinopathy and venous occlusions) and/or inflammation.⁹ A systematic review from 2016 that included over 49,000 subjects found that ERMs are relatively common among older populations, and the meta-analysis showed that only greater age and female gender significantly conferred a higher risk of ERM.¹⁰

The vitreous is most firmly attached at the vitreous base, the optic nerve head, and the macula.^{11, 12} A posterior vitreous detachment (PVD) evolves over a prolonged period of time.⁴ Initially, and usually associated with age, the posterior vitreous will partially detach yet will remain attached within the macular region. Eventually, a complete detachment occurs when the vitreous detaches from the macula and finally from the optic nerve head. When the vitreous detaches from the nerve head, the patient may see the acute onset of floaters, flashes, or photopsia. Combined, these represent the classic symptoms for the onset of an acute PVD. On fundus examination, a Weiss ring represents the glial remnant from the attachment at the optic nerve on the posterior cortical vitreous and is typically seen on the posterior vitreous face anterior to the optic nerve.

During the evolution of a PVD, the vitreous may remain adherent to the macula. Vitreomacular adhesion, the attachment of the posterior cortical vitreous to the neurosensory retina, may represent the normal evolution of a PVD. Vitreomacular traction occurs when the perimacular vitreous separates from the posterior retina yet remains adherent to a region or area near the center of the macula and causes distortion or change of the normal macular anatomy.^{4, 12} The pathologic mechanism responsible for such an abnormal adhesion within

the macula that leads to VMT is unclear. The combination of attachment at the macula with surrounding vitreous separation creates traction. It may lead to thickening, distortion, intraretinal cystoid changes and even subretinal fluid or tractional detachment at the macula.⁴ Epiretinal membranes can also lead to macular traction. Both ERM and VMT may lead to loss in visual acuity, metamorphopsia, difficulty in using both eyes together, and diplopia.

On examination, the most common type of ERM appears as a thin, translucent, cellophane-like membrane on the surface of the retina.^{11, 13} An ERM may not lead to tractional changes, and the underlying neurosensory retina may often appear normal. However, epiretinal membranes can contract, leading to folds in the retina, distortion of the inner and even the outer macula, traction on retinal vessels, and even displacement of the macula, or ectopia. The normal foveal depression is often absent or distorted, and the macula may develop cystoid spaces, lamellar macular hole, or even a full-thickness hole. Epiretinal membranes that have a thicker, white, fibrotic appearance that obscures the underlying retina are more likely than the thinner, more translucent ERMs to become symptomatic and displace the macula.^{6, 14}

The retinal changes associated with VMT are often similar to the changes that result from an ERM. Both ERM and VMT may be associated with adherent vitreous in a peripapillary distribution around the optic nerve head, which is referred to as vitreopapillary traction.¹⁵ Optical coherence tomography (OCT) scans through the optic nerve can help diagnose vitreopapillary traction, which may be confused with optic nerve disorders such as disc edema or, if it is present in both eyes, papilledema.¹⁵⁻¹⁷ There is some suspicion that vitreopapillary traction might be associated with decreased vision and even ischemic optic neuropathy in some cases.¹⁸

INCIDENCE AND PREVALENCE

Epiretinal membrane and VMT are relatively common retinal conditions. Higher prevalence of both conditions is associated with older age.⁶ Vitreomacular traction is less common than ERM and affects an estimated 0.4% to 2.0% in a population of U.S. adults over the age of 63.¹⁹ The prevalence of ERMs is based on several population-based studies conducted in various racial and ethnic groups worldwide over the past 20 years. It is estimated to occur in approximately 30 million adults in the United States 43 to 86 years old.²⁰ Epiretinal membranes may be bilateral in up to 20% to 35% of cases.^{8, 21-23} Prevalence rates⁶ range from a low of 2.2% and 3.4% in the Beijing Eye Study²⁴ and in the Handan Eye Study in rural China, respectively,⁸ to moderate (7% and 8.9%) in two Australian populations,^{21, 25} to a high of 18.8% and even 28.9% among Latino individuals in Los Angeles²⁶ and in the Multi-Ethnic Study of Atherosclerosis (MESA) conducted in six communities in the United States.²³ The presence or absence of ERM in most studies was based on the use of nonmydriatic retinal photography.²⁰⁻²⁸ At the 20-year follow-up examinations of the Beaver Dam Eye Study population (mean age of 74.1 years), spectral-domain optical coherence tomography (SD-OCT) was used and documented a higher prevalence of 34.1%.¹⁹ In eyes with no macular pathology on clinical exam before cataract surgery, prevalence of ERM with routine SD-OCT ranged from 2.2% to 11.0%.29,30

In most populations studied, the early asymptomatic form of ERM, also known as cellophane maculopathy, occurred more frequently than the thicker or more opaque preretinal macular fibrosis (a term used for symptomatic ERM).^{8, 22, 23, 26} The prevalence of cellophane maculopathy varied from 1.8% and 2.2% in urban and rural China^{8, 24} to as high as 16.3% among Los Angeles Latino individuals²⁶ and 25.1% in MESA.²³ Diabetes and hypercholesterolemia are associated with higher rates of cellophane maculopathy.²³ Preretinal macular fibrosis prevalence was more consistent across studies, with rates ranging from 0.7% in rural China⁸ to 3.5% among Asian Indian individuals,²⁷ 3.8% in MESA,²³ and 3.9% in Melbourne, Australia.²⁵

Several reasons might explain the variable prevalence results from different studies, including the sensitivity of the specific testing or imaging modality used, differences in classification of ERM, and differences in the populations (e.g., age, race and ethnicity, lifestyle). Optical coherence tomography is considered "the de facto standard for ERM diagnosis," but artificial intelligence is being evaluated to aid in the diagnosis of ERM using fundus photography and/or ophthalmoscopic examination, which "have advantages of price and accessibility."³¹

RISK FACTORS

Increasing age was consistently identified as a risk factor for ERM in all studies.⁶ Prevalence varies by race and ethnicity, but patterns are not consistent across studies. For example, in the United States, MESA data suggest that the prevalence of any ERM was highest in individuals of Chinese ancestry (39.0%), intermediate in Hispanic individuals (29.3%) and White individuals (27.5%), and lowest in Black individuals (26.2%),²³ whereas the data from China suggested that the ERM prevalence rates were much lower (2.2% and 3.4%).^{8, 24} Epiretinal membrane occurs more frequently in persons with retinal pathology (e.g., uveitis and other ocular inflammatory diseases,³² retinal breaks,³³ retinal vein occlusions,^{19, 20, 23} proliferative diabetic retinopathy^{6, 19}) and following cataract surgery.^{6, 19} It may be associated with impaired visual acuity or visual field loss,^{19, 24} particularly for those eyes with more severe ERMs.²⁶ A number of other more speculative risk factors have been suggested but have not been confirmed. These include female gender,⁶ myopia,³⁴ hyperopia,²⁸ smoking,^{8, 25} higher education,⁶ diabetes,⁶ hypercholesterolemia,⁶ narrow retinal arteriolar diameter,⁶ higher body mass index,²⁵ genetics,³⁵ and stroke.²⁵ There has been a case report documenting the onset of VMT with the use of 1% topical pilocarpine drops, but the VMT was resolved when the drops were discontinued.³⁶

PATHOGENESIS OF EPIRETINAL MEMBRANE AND VITREOMACULAR

TRACTION

Epiretinal Membrane

A longstanding hypothesis was that ERMs develop when a PVD results in microbreaks of the internal limiting membrane (ILM) that, in turn, allow for the migration of retinal glial or possibly retinal pigment epithelial cells onto the anterior retinal surface, where they proliferate.^{13, 37} The hypothesis was supported when retinal pigment epithelial cells, fibrous astrocytes, astrocytes, and fibrocytes were observed in ERMs of eyes that had no apparent retinal breaks, laser or cryopexy, or eye surgery.³⁸ An alternative hypothesis gaining acceptance is that ILM breaks are not necessary for ERMs to develop, and an ERM may originate from cells in the cortical vitreous remnants on the ILM that are activated into myofibroblasts resulting in membrane formation and contraction.^{6, 12, 39} Epiretinal membranes have also been observed in eyes without an obvious PVD.⁴⁰ An article evaluating posterior vitreous attachment in eyes undergoing surgery for idiopathic ERMs noted 20.1% of eyes had posterior vitreous attachment at the time of surgery.⁴¹ In eyes with a PVD, vitreous remnants have been documented on the surface of the retina.^{12, 42} The presence of a Weiss ring does not always indicate that there has been a complete separation of the posterior hyaloid membrane from the entire posterior retinal surface.⁴³

Laminocytes, vitreous cells from the posterior hyaloid membrane (hyalocytes), have been shown to represent a major cellular component of idiopathic ERMs.⁴⁴ Hyalocytes, however, are not native to the vitreous but originate from bone-marrow-derived cells and are continuously renewed.⁴⁵ Extracellular matrix material has also been consistently detected in specimens of ERMs from eye bank eyes or surgically removed membranes.⁶, ³⁸, ⁴⁴ Retinal glial cells, hyalocytes, their transdifferentiation into fibroblasts and myofibroblasts, along with the development of extracellular matrix and fibrosis, together lead to ERM formation.⁶ In summary, these and other studies show that the formation of an ERM includes some combination of vitreous collagen, several different potential cellular origins, differentiation of these cells, and the formation of new collagen and an extracellular matrix material. The constitution of ERMs varies and, therefore, ERMs are likely to have a variety of possible origins and causes.

Vitreomacular Traction

Posterior vitreous detachment may be a prolonged process, and portions of the posterior cortical face may remain adherent to the macula and lead to tractional changes. Investigators have broadly separated VMT, based on OCT, into small and large areas of adherence. A localized vitreomacular attachment of about 500 µm causes elevation, traction, and subsequent intraretinal cystoid spaces in the foveal neurosensory retina. A broad attachment measuring

about 1500 µm (approximately 1 disc diameter) can cause more elevation of the macula, even to the point of a macular retinal detachment, yet this configuration is less likely to have intraretinal cystoid spaces.^{42, 46} Of course, there is a continuum of areas of attachment from pinpoint to over 1500 µm in diameter. The vitreous attachment may release spontaneously over time, especially in eyes with more focal areas of adherence.⁴⁷

Epiretinal membranes often contain native vitreous collagen on histopathology specimens and may evolve between the neurosensory retina and a vitreous attachment.⁴⁶ Because ERMs adhere tightly to the ILM, they may play a role in VMT by binding the remaining attachment of the vitreous to the macula.^{46, 48, 49}

CARE PROCESS

PATIENT OUTCOME CRITERIA

Patient outcome criteria include the following:

- Prevent vision loss and functional impairment
- Optimize visual function
- Minimize symptoms (e.g., metamorphopsia, diplopia)
- Maintain or improve quality of life

DIAGNOSIS

History

Many people who have an ERM have stable vision with few symptoms, whereas others are more symptomatic and have progressive loss of visual function. Patients are often especially bothered by metamorphopsia or diplopia and may experience difficulties in reading, driving, or being able to use their eyes together.⁵⁰⁻⁵³ Commonly, patients report that they close one eye while reading in order to eliminate the distortion from the affected eye.

Patients with VMT have similar symptoms of impaired visual function and metamorphopsia that may be acute or chronic depending on the severity of the traction and the resulting distortion or detachment of the macula. Frequently, the visual acuity of patients with either VMT or ERM does not change dramatically during short-term follow-up.^{14, 54, 55} Vitreomacular adhesions are often asymptomatic.

Examination

Examination includes all features of a comprehensive adult medical eye evaluation,⁵⁶ with particular emphasis on the following:

- Slit-lamp biomicroscopy of the macula and vitreoretinal interface
- Slit-lamp biomicroscopy of the optic disc to rule out an optic pit or advanced cupping
- An indirect peripheral retinal examination with scleral depression to evaluate for retinal breaks and other pathology
- Amsler grid test
- Watzke-Allen sign, which "can be used as a clinical test in cases of a suspected full thickness macular hole by shining a thin beam of light over the area of interest. The patient would perceive a 'break' in the slit beam in cases of a positive test."⁵⁷

Diagnostic Tests

Optical coherence tomography is a highly sensitive and routine method used to diagnose and characterize VMA (see Figure 1), ERM, VMT (see Figure 2), and the associated retinal changes.^{4, 30, 46, 47, 54, 58-61} Comparing the OCT images in the abnormal eye with images of a normal eye (see Figure 3) is a very helpful educational tool to help patients better understand their eye problem. An ERM on OCT appears as a hyper-reflective and sometimes irregular layer on the inner surface of the retina (see Figure 4), usually adherent across the surface of the retina. It is frequently attached by pegs emanating from the inner retinal surface with intervening hyporeflective spaces of ERM separation that gives a corrugated appearance in cross section. Optical coherence tomography commonly demonstrates that the traction from the ERM leads to elevation of the normal foveal depression. The inner retinal layers.⁶² Using OCT imaging, lamellar macular holes (see Figure 5) may have variable degrees of inner-retinal tissue loss, often with well-delineated edges that are affected by tractional elements from the ERM.⁶³⁻⁶⁶



FIGURE 1. Vitreomacular adhesion. The posterior vitreous face (blue arrows) is separated from the neurosensory retina and a foveal attachment (white arrow) or vitreomacular adhesion remains. Note that there is no secondary retinal pathology from this attachment site. (Courtesy of Timothy W. Olsen, MD)



FIGURE 2. Vitreomacular traction. (Copyright © 2015 American Academy of Ophthalmology[®])



FIGURE 3. Normal retina. The various layers of the retina are easily visualized using spectral-domain optical coherence tomography through the fovea. (Copyright © 2015 American Academy of Ophthalmology^{*})



FIGURE 4. Epiretinal membrane. Optical coherence tomography reveals a fine, moderately reflective membrane variably attached to the inner retinal surface. There is associated retinal edema. (Copyright © 2015 American Academy of Ophthalmology*)



FIGURE 5. Lamellar hole. Optical coherence tomography demonstrates an intraretinal split, with separation of the inner and outer foveal retinal layers and the absence of a full-thickness foveal defect. (Copyright © 2015 American Academy of Ophthalmology*)

The OCT findings of VMT are similar, except that the posterior hyaloid remains partially attached to the macula and is separated in the perimacular region.^{67, 68} Cystoid spaces may be present in the entire macular region in VMT. Presumably, these changes are due to anterior-posterior vitreous traction associated with VMT as opposed to a more tangential traction from an ERM. The extent of the VMT varies from a small focal adhesion to a large, broad adhesion that extends over the entire macula.^{47, 69} Both ERM and VMT often occur together; thus, the features are commonly combined.⁴⁶ In 60 eyes with ERM, the vitreous was noted to be adherent to the macula in 57%.⁷⁰ Similarly, 13/20 eyes (65%) with VMT were noted to also have an ERM.⁷¹

Ancillary Tests

A fluorescein angiogram or optical coherence tomography angiography (OCTA)^{72, 73} may be helpful to evaluate ERMs and/or VMT.⁷⁴ The fluorescein angiogram and OCTA may be useful to detect other retinal pathologies that can be associated with ERMs, such as a branch retinal vein occlusion, diabetic retinopathy, macular telangiectasia, choroidal neovascularization, and other inflammatory conditions. The fluorescein angiogram may be relatively normal in eyes with early ERM. As ERM contraction increases, the macular vessels may become tortuous near the epicenter of traction or straightened around the epicenter of traction. Some retinal vessels, especially the capillaries that are under tractional forces, may demonstrate a leakage pattern, best detected by comparing the early stages of the angiogram with the later stages. The dye may pool in cystoid spaces, especially in the recirculation phase. However, the staining and leakage in the fovea is usually not as uniformly petaloid and circular as typically seen in pseudophakic cystoid macular edema (which is often accompanied by a hyperfluorescent optic nerve in the later phase of the angiogram). Retinal vascular changes, such as capillary dropout, telangiectasia, collateral vessels, and microaneurysm formation suggest the presence of an underlying retinal vascular disease, such as diabetic retinopathy or central vein occlusion. Fluorescein crosses the placenta and it is present in breastmilk for 72 hours.^{75, 76}

MANAGEMENT

Nonsurgical

Patients should be informed that the majority of ERMs will remain relatively stable and do not require therapy.¹⁴ The visual acuity may worsen over time and rarely improves spontaneously. Patients should also be reassured that there is a very successful surgical procedure that could address worsening symptoms or decreasing visual acuity. Furthermore, patients should be encouraged to periodically test their central vision monocularly in order to detect changes that may occur over time, such as increasing metamorphopsia and/or development of a small, central scotoma. Informing patients to monitor for the signs and symptoms of progression and regular monocular Amsler grid testing are both important.

Observation without Treatment

Using fundus photography, a population-based study of 3654 people showed that only 29% of ERMs progressed over 5 years; 26% regressed, and 39% remained approximately the same. Only 20% of eyes with cellophane maculopathy progressed over the same time period.¹⁴ A clinic-based study of 34 eyes with ERM and lamellar macular holes showed that the vision did not change over a mean follow-up of 18 months, although two eyes progressed to a full-thickness macular hole.⁵⁴ A study of 47 eyes with ERM found that the visual acuity and clinical appearance did not change over a mean of 38 months.⁵⁵ A study using SD-OCT images found that the ERM separated from the retina in only 16 of 1091 (1.5%) eyes with a pre-existing PVD but in 21/157 (13.6%) of eyes that did not have an apparent PVD over a mean follow-up of 33 months.⁷⁷ The separation of the ERM led to improved visual acuity in both groups. The majority of ERMs remain stable after initial presentation. Deferring surgery until symptoms develop does not have a worse outcome compared with immediate surgery.⁷⁸

In eyes with VMT of 1500 μ m or less, patients often have stable visual acuity, and the incidence of spontaneous release of traction from the macula occurs in 23% to 47% of eyes over a follow-up of 1 to 2 years.^{47, 61, 68, 69, 79-81} Usually the release of traction results in an improvement in visual acuity and less severe symptoms, assuming no full-thickness macular hole is created. An earlier study, however, found that the visual acuity in 34 of 53 eyes (64%) with VMT decreased 2 Snellen lines or more over 60 months of follow-up.⁸¹ However, 43/53 (81%) of the eyes reported in this study had cystoid macular spaces detected at baseline. Thus, eyes with cystoid spaces at baseline may represent a cohort of patients with a more guarded prognosis.⁸²

Surgery

Gas Injection for Vitreomacular Traction

The injection of intravitreal gas has been reported to also induce release of VMT within 1 month in 40% of study eyes in a relatively small cohort of 15 eyes.⁸³ A cohort of 30 eyes showed a slightly higher rate of release of 73% within 1 month.⁸⁴ Another, smaller study (9 eyes) used SF₆ gas and had similar results (56% within 1 month).⁸⁵ In another study of 56 eyes, the rate of release of VMT using 0.3 ml pure C_3F_8 was 85.7% and the rate of closure of small holes was 60%.⁸⁶ The Diabetic Retinopathy Clinical Research (DRCR) Retina

Network conducted a randomized trial evaluating pneumatic vitreolysis (PVL) with intravitreal injection of C_3F_8 gas compared with sham injection. Seventy-eight percent of eyes in the PVL group had release of VMT from the central macula compared with 9% in the control group. Higher than expected rates of retinal detachment and retinal tear in the PVL group resulted in early termination of the study due to safety concerns.⁸⁷ The small sample size and early termination of the study prevented definitive findings on safety and efficacy of PVL for VMT. Due to the lack of clear guidance from clinical trials, clinicians need to use their judgment and counsel patients closely on risks and benefits based on the available limited evidence.

Vitrectomy Surgery

A Cochrane review found no randomized controlled trial that evaluated surgery versus no intervention.⁸⁸ (I-, Insufficient quality) The decision to intervene surgically in patients with ERM/VMT usually depends on the severity of the patient's symptoms, especially the impact on their activities of daily living. Patients should be asked how much they are bothered and/or impaired by their visual dysfunction; asking about impairments of reading or driving ability is usually very important. Patients should also specifically be questioned about distortional changes. Vitrectomy surgery for ERM/VMT is elective rather than urgent. Earlier surgical intervention for ERM may result in better long-term visual acuity recovery than delayed surgery, yet the time frame of the delay is considered in months rather than in days.⁸⁹ Patients with VMT do not typically improve without vitrectomy surgery when the area of VMT is broad (>1500 µm), when there is an accompanying pathologic detachment of the macula, or when the presenting visual acuity is poor.⁴⁶ Overall, the recommendation to observe or perform surgery is mainly based on patients' discomfort with their vision, along with their understanding of the associated risks. Appropriate intervention should be made with careful informed consent and a discussion of the risk-benefit ratio of surgery.

Preoperative Discussion for Vitrectomy

The preoperative discussion should include the risks (e.g., cataract, retinal tears, retinal detachment, endophthalmitis, vision loss due to retinal damage) versus the benefits of vitrectomy surgery. Discussion should also cover the following aspects of vitrectomy surgery:

- The risk of cataract progression following pars plana vitrectomy in phakic eyes is high. Such progression occurs at variable rates and may be age dependent.
- If a cataract is present, cataract surgery may be deferred, recommended prior to vitrectomy surgery, or done at the same time as vitrectomy surgery.
- The type of anesthesia used is typically local monitored anesthesia. General anesthesia may also be used, especially for anxious or claustrophobic patients.
- Usually, the visual acuity and symptoms of distortion and diplopia will improve but not necessarily resolve completely. In some cases, visual acuity may decrease and not recover.
- There is a risk of epiretinal membrane recurrence.
- There is a risk of increase or decrease in postoperative intraocular pressure, especially in patients with glaucoma.
- The surgeon is also responsible for planning postoperative care and for communicating care instructions.^{90, 91}

Technique

Epiretinal membranes and VMT are often present in the same eye. During surgery, both the VMT and ERM must be removed from the retina surface in order to release the traction on the macula.⁴⁶ Furthermore, some suggest that removal of the ILM around the macula releases the traction even more completely and reduces the rate of recurrence.⁹² One potential explanation for the reduced rate of recurrence in eyes that undergo ERM and ILM removal could be related to residual glial and fibrotic elements seen on the

retinal surface of the ILM on histopathology after ERM removal in 80% of specimens in one study. 92

Surgical removal of ERM/VMT is usually performed using a 23-, 25-, or a 27-gauge vitrectomy system combined with local, monitored anesthesia care. The core vitreous is removed, and the surgeon induces a detachment of the posterior hyaloid from the optic nerve and macula. The off-label use of indocyanine green dye, trypan blue, or triamcinolone may be used during surgery to highlight the ILM and remaining vitreous, respectively. Food and Drug Administration approval of Brilliant Blue G Ophthalmic Solution (TISSUEBLUE®, Dutch Ophthalmic, US) in 2019 has provided surgeons with another option for visualization that has safety data.93 The posterior hyaloid is commonly separated from the retinal surface using aspiration, an illuminated pick, or forceps. The peripheral vitreous is shaved, particularly near the cannulas, to minimize the risk of iatrogenic retinal breaks during instrument exchanges. The vitreous is separated from the retinal surface, extending at least anteriorly to the equator, and removed. Next, the ERM and frequently the ILM are removed with intraocular forceps, often under specialized viewing systems to enhance visualization of the macula. Typically, forceps, microvitreoretinal blade, diamond-dusted silicone tip, loop, or a needle may be used to elevate an edge of either the ERM, ILM, or both together, which is then peeled and removed with a forceps.⁹⁰ Regardless of the technique, the surgical objectives are to gently free the macula of tractional elements.

Histopathology of the peeled membrane demonstrates variable amounts of ILM. However, often there are patches of ERM and large areas of ILM left on the retinal surface after the initial peel. These remnants can be difficult to visualize. Many surgeons choose to use agents such as indocyanine green dye, brilliant blue dye, trypan blue, or off-label triamcinolone to help visualize the ILM and facilitate the peel. The safety of these dyes remains somewhat controversial,⁹⁴ yet many surgeons agree that very low concentrations of dyes appear safe and may minimize trauma to the retina because the ILM is more easily visualized.

Minimizing excessive intraoperative exposure of the macula to light is important. The ERM is typically thicker and has a shaggy or irregular configuration, whereas the ILM is thin, homogenous, and scrolls following removal from the retinal surface.

Once the ERM, ILM, or VMT has been removed, the retina can be examined for retinal breaks or detachment. A small intraocular air bubble may be used to help seal nonsutured sclerotomies. When a surgeon suspects a full thickness or deep lamellar hole, a more complete fluid-gas exchange using a nonexpansile or minimally expansile concentration of C_3F_8 or SF_6 gas is performed.

Removal of the Internal Limiting Membrane

Table 1 lists studies that compare the results of removing the ERM alone with removing both the ERM and ILM. Five of the studies found that peeling the ILM with the ERM led to a lower incidence of recurrent ERM. Two studies showed no difference between peeling or not peeling the ILM. Of note, ILM peeling can be associated with loss of inner retinal tissue, although the functional impact of this finding is unclear. A systematic review of 13 studies found no difference in visual acuity outcomes between the two groups but greater anatomical success with ILM peeling.⁹⁵ (*I*+, *Good quality, Discretionary recommendation*) Another study, a meta-analysis of randomized clinical trials, found that ILM peeling did not significantly improve the visual outcome or the ERM recurrence rate.⁹⁶ One study did report that the ILM-without-peeling-group experienced greater and faster recovery of retinal sensitivity than the ILM peeling and without ILM peeling were similar, but there was a lower rate of recurrence with ILM peeling at 6 to 12 months postoperatively.⁹⁸ (*I*-, *Moderate quality, Discretionary recommendation*)

Study	Study Design	No. of Eyes with ERM	Follow-up (mos)	Results	ERM Removal with or without ILM Peel Was Not Favored	Removal of Both ILM and ERM Was Favored	ERM without ILM Removal Favored
Park et al, 2003 ⁹⁹	Case series	44	At least 3	24 eyes no ILM peel (Group A); 20 eyes with ILM peel (Group B). Average increase in logMAR was 0.33 in Group A and 0.41 in Group B. Recurrence rate of ERM was 21% in Group A and 0% in Group B.		•	
Bovey et al, 2004 ¹⁰⁰	Case series	71	Range 6-59, mean 21.7	ERMs peeled with no attempt to peel ILM but ERM then studied by histopathology. Fifty-five of 71 eyes had long segments of ILM and 16 did not. The 55, which had ILM, had 3 lines of vision gain compared to 1 line in non-ILM group; recurrence rate of ERM was 9% in ILM group and 56% in non-ILM group.		•	
Koestinger and Bovey, 2005 ¹⁰¹	Case series	75	Mean, 20	ERM removed in only 55 eyes and ILM also peeled in 20 eyes using ICG to stain. No difference in VA between groups.	•		
Kwok et al, 2005 ¹⁰²	Case series	42	Mean, 32.8	17 ERMs removed with no ILM peel, and in 25 eyes both ERM and ILM were peeled. Postop VA was logMAR 0.65 in the non-ILM peel group and 0.46 in the peel group. ERM recurred in 3/17 of the non-ILM peel group.		•	
Shimada et al, 2009 ¹⁰³	Case series	246	12	104 eyes ERM removed only; 142 eyes ERM and ILM removed. Recurrence rate of ERM was 17/104 (16.3%) in ERM-only group and 0/142 eyes in ERM/ILM group. Postop VA did not differ between the groups.		•	
Oh et al, 2013 ¹⁰⁴	Case series	43	12	23 eyes ERM only; 20 eyes ERM and ILM peeled. ILM peel group was not favored at 3 months. No difference between two groups at 12 months for VA, central retinal thickness, and mfERG.	•		
Sandali et al, 2013 ¹⁰⁵	Case series	440	At least 12	174 eyes had no ILM peel; 266 eyes had ILM peel. VA improvement postop was the same between the two groups; VA same with dye-assisted ILM peel compared with none. Recurrence rate of ERM was in 8.6% in non-ILM peel group and 2.6% in ILM peel group.	•		
Ripandelli et al, 2015 ⁹⁷	Randomized controlled trial	60	12	ILM removed in 30 eyes, ERM only in 30 eyes. Microperimetry showed statistically significantly greater and faster recovery in ERM-only group.			٠

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Study	Study Design	No. of Eyes with ERM	Follow-up (mos)	Results	ERM Removal with or without ILM Peel Was Not Favored	Removal of Both ILM and ERM Was Favored	ERM without ILM Removal Favored
Fang et al, 2017 ⁹⁵	Systematic review	359	At least 3	Systematic review of 13 studies; no difference in BVCA at 12 months (primary outcome) between ERM/ILM group vs ERM-only group, but there was significantly increased CMT in the ILM peeling group.	•		
Tranos et al, 2017 ¹⁰⁶	Randomized controlled trial	102	12	ILM removed in 50 eyes, ERM only (no ILM) in 52 eyes. No difference in BCVA or OCT thickness.	•		
Far et al, 2021 ⁹⁸	Systematic review	387	12	Systematic review of 7 studies: 207 eyes had ERM removal with ILM peel, 180 had ERM removal only. No difference in VA outcomes between groups. Lower rate of ERM recurrence at 6–12 months with ILM peel.		•	
Sun et al, 2021 ⁹⁶	Systematic review	422	At least 6	No significant difference in BCVA, logMAR 0.03, or recurrence rate. Thicker central macular thickness in ILM peel group at final follow-up.			•
Ducloyer et al, 2024 ¹⁰⁷	Randomized controlled trial	213	12	ILM spontaneously peeled in 101 eyes, 51 eyes randomized to active ILM peeling, 49 eyes randomized to no peeling. Rate of recurrence was lower with active peeling (0%) vs no peeling (19.6%), but active peeling delayed visual recovery (BCVA and microperimetry).		•	

TABLE 1 RESULTS OF NO ILM PEEL VS. ILM PEEL IN ERM AND VMT

BCVA = best corrected visual acuity; ERM = epiretinal membrane; ICG = indocyanine green; ILM = internal limiting membrane; mfERG = multifocal electroretinography; OCT = optical coherence tomography; postop = postoperative; VA = visual acuity; VMT = vitreomacular traction.

Outcome

Vitrectomy surgery is often indicated in patients who are affected by a decrease in visual acuity, metamorphopsia, double vision, or difficulty using their eyes together. Table 2 lists results for ERM and VMT following vitrectomy. On average, the visual acuity improves by 2 lines or more after surgery. The visual results are highly variable, however; although some patients experience large visual acuity gains, it is important to note that, overall, 10% to 20% of patients will have unchanged or worse vision following surgery. Although results are variable, scores on the National Eye Institute Vision Function Questionnaire (NEI-VFQ)-25, on average, improve postoperatively both at 6 and 24 months.¹⁰⁸ Most metamorphopsia improves and may normalize. A study of 42 patients found that NEI-VFQ-25 scores improved at 6 months as well as stereopsis and distance BCVA.¹⁰⁹ Thus, even in the absence of visual acuity gain, some patients are pleased with the relief from some or all of the metamorphopsia.

A study of 43 eyes showed that preoperative OCT evidence of an intact inner photoreceptor and ellipsoid zone, also referred to as the inner segment/outer segment junction, was associated with better visual acuity after a vitrectomy for ERM.¹¹⁰ A similar study showed that the integrity of the ellipsoid zone and the cone outer segment tips line (also known as the interdigitation zone) was also associated with better visual acuity.¹¹¹ The outer retina, the ellipsoid zone, and the photoreceptors' outer segment length may improve or even normalize after vitrectomy, and each feature is associated with improved visual acuity.¹¹⁰ In another study of 101 eyes using time-domain OCT, the presence of photoreceptor disruption was found to be a predictor of poor visual outcome after surgery.¹¹³

A systematic review and meta-analysis of combined versus sequential pars plana vitrectomy and phacoemulsification for macular hole or epiretinal membrane did not find a significant difference in terms of complications and visual and refractive outcomes.¹¹⁴

Study	No. of Patients	Follow-up (mos)	Results
ERM Diagnosis			
Koerner and Garweg, 1999 ⁷⁰	60	Mean 24.7	73% improved vision; 61% 20/50 or better; 57% final VA better than preop
Wong et al, 2005 ¹¹⁵	125	10. 3	VA improved by a mean of 0.31 log units or 3 lines of vision; 16% had unchanged acuity postop
Ghazi-Nouri et al, 2006 ¹⁰⁸	20	4	No postop gain in mean VA; 40% gained 2 lines or more; metamorphopsia decreased significantly (<i>P</i> = 0.02); VFQ-25 improved significantly (<i>P</i> = 0.03)
Arndt et al, 2007 ⁵¹	85	12	56% of patients had metamorphopsia preop and 13% postop
Bouwens et al, 2008 ¹¹⁶	107	Results at 12	Mean postop VA gained 2 lines; 83% had less metamorphopsia
Okamoto et al, 2009 ¹¹⁷	28	3	LogMAR improved from 0.49 preop to 0.24 postop; 11 (39%) had no change in logMAR; VFQ-25 scores significantly improved
Matsuoka et al, 2012 ¹¹⁸	26	12	LogMAR VA 0.41 preop, 0.17 at 3 mos, 0.10 at 12 mos; metamorphopsia score (baseline, 3, and 12 mos was 202, 137 and 108 respectively); VFQ-25 scores significantly better at 3 and 12 mos

TABLE 2 RESULTS OF VITRECTOMY FOR EPIRETINAL MEMBRANE AND VITREOMACULAR TRACTION

Idiopathic ERM and VMT PPP

Study	No. of Patients	Follow-up (mos)	Results
Garcia-Fernandez et al, 2013 ¹¹⁹	88	12	82% had better vision but 10% worse postop
Dawson et al, 2014 ¹²⁰	237	6	Mean preop 20/120; mean postop 20/40
Elhusseiny et al, 2020 ¹²¹	49	111	The mean BCVA improved from 0.56 \pm 0.29 (20/72) preoperatively to 0.33 \pm 0.25 (20/42) at 1 year, 0.29 \pm 0.27 (20/38) at 2 years, 0.25 \pm 0.28 (20/35) at 3 years, 0.29 \pm 0.32 (20/38) at 5 years, 0.28 \pm 0.31 (20/38) at 8 years, and 0.28 \pm 0.25 (20/38) at 10 years (<i>P</i> < 0.001). The BCVA improved at each of the first 3 years postoperatively and remained stable at 5, 8, and 10 years
VMT Diagnosis			
Koerner and Garweg, 1999 ⁷⁰	50	Mean 10	73% improved vision; 66% 20/50 or better; 60% final VA better than preop
Witkin et al, 2010 ⁷¹	20	28.6	Mean VA preop was 20/122 and postop was 20/68
Jackson et al, 2013 ¹²²	Meta-analysis 259 eyes from 17 articles	Variable; range 6–35	Mean preop logMAR 0.67; mean postop 0.42; 33% gained 2 or more lines; 21% of eyes had same or decreased VA postop
Morescalchi et al 2021 ¹²³	RCT – 34 eyes	12 months	Parafoveal retinal sensitivity exhibited a significant improvement in both the foveal sparing (FS) and complete peeling (CP) groups (+2.43 \pm 0.82 dB and +1.79 \pm 0.86 dB, respectively; <i>P</i> = 0.037). Mean postoperative BCVA improved to logMAR 0.27 in the CP group and logMAR 0.14 in the FS group.

TABLE 2 RESULTS OF VITRECTOMY FOR EPIRETINAL MEMBRANE AND VITREOMACULAR TRACTION

BCVA = best corrected visual acuity; ERM = epiretinal membrane; logMAR = logarithm of the minimum angle of resolution; mos = months; postop = postoperative; preop = preoperative; RCT = randomized controlled trial; VA = visual acuity; VFQ-25 = National Eye Institute Visual Function Questionnaire; VMT = vitreomacular traction.

Complications

The majority of phakic patients develop a progressive nuclear cataract following vitrectomy for ERM.¹²⁴⁻¹²⁸

Retinal breaks and detachments are less common with current vitrectomy surgery, likely due to smaller-gauge instruments, cannulated sclerotomies, improved visualization of the retinal periphery, and management of the peripheral vitreous, including treatment of retinal breaks and or localized detachments. There may also be less vitreous incarceration leading to retinal traction with smaller-gauge sclerotomies. Retinal breaks have been reported to occur in approximately 1% of cases (8 of 548) during vitrectomies performed using a 23-gauge cannula system.¹²⁹ Another study also found that retinal detachments occur in 1% (2 of 166) of consecutive 23-gauge vitrectomies.¹³⁰ A third study reported that in a total of 349 eyes retinal detachments occurred in 1% of eyes undergoing a 23-gauge vitrectomy and in 3.5% of eyes undergoing 20-gauge vitrectomy.¹³¹ Endophthalmitis has been reported in less than 0.05% of vitrectomies.¹³²⁻¹³⁴ Macular hole formation is also a potential complication of vitrectomy surgery for ERM and VMT and has a low incidence of 2.1%.¹³⁵

Follow-up Evaluation after Surgery

Patients who have surgery should be examined on postoperative day 1 and again 1 to 2 weeks following surgery or sooner, depending on the development of new symptoms or

new findings during early postoperative examination. The primary reasons for an earlier follow-up visit or more frequent follow-up visits are high or low intraocular pressure, a wound leak, pain, worsening vision, or other concern of a retinal complication. Components of the follow-up examination should include the following:

- Interval history, including new symptoms
- Measurement of intraocular pressure
- Slit-lamp biomicroscopy of the anterior segment, including the wound sites and central retina, if possible
- Indirect binocular ophthalmoscopy of the peripheral retina
- Counseling on the use of postoperative medications
- Counseling on the signs and symptoms of retinal detachment
- Precautions about intraocular gas if it has been used
- Referral to a strabismus specialist or orthoptist (may be helpful for continued symptoms of diplopia or difficulty using the eyes together)

PROVIDER AND SETTING

Diagnosis and management of ERM, VMT, or VMA require special expertise, surgical skills, and specialized equipment to detect alterations in the retina in order to select, perform, implement, and monitor appropriate management or treatment. Referral to an ophthalmologist who has expertise or experience in managing this condition is recommended. The performance of diagnostic procedures is often delegated to appropriately trained and supervised personnel. However, the interpretation of the results of the diagnostic procedures, as well as the medical and surgical management of ERM, requires the medical training, clinical and surgical judgment, and experience of an ophthalmologist who is also trained in vitreoretinal surgery and disease.

COUNSELING AND REFERRAL

Patients should be informed to notify their ophthalmologist promptly if they have symptoms such as an increase in floaters, a loss of visual field, metamorphopsia, or a decrease in visual acuity.¹³⁶⁻¹³⁸ Patients with functionally limiting postoperative visual impairment should be referred for vision rehabilitation and social services.^{139, 140} Empathic communication and questioning by the provider is helpful for eliciting patient concerns. Referrals for counseling, vocational rehabilitation and/or peer support groups for patients with depression, anxiety, and loss of independence or employment should be considered.¹⁴¹ Such a referral is particularly important when there is a residual central or paracentral scotoma. More information on vision rehabilitation, including materials for patients, is available at www.aao.org/education/low-vision-rehab.

SOCIOECONOMIC CONSIDERATIONS

A cost-utility analysis of ERM surgery from 2008 in the better-seeing eye compared with observation resulted in a mean gain of 0.755 discounted quality-adjusted life years (QALYs) (3% annual rate) per patient treated. This model resulted in \$4,680 per QALY for this procedure. When sensitivity analysis was performed, utility values ranged from \$3,746 to \$6,245/QALY gained, and medical costs varied from \$3,510 to \$5,850/QALY gained.¹⁴² Epiretinal membrane surgery in the worse-seeing eye compared with observation resulted in a mean gain of 0.27 discounted QALYs per patient treated. The \$/QALY was \$16,146, with a range of \$12,110 to \$20,183 based on sensitivity analyses. Utility values ranged from \$12,916 to \$21,520/QALY. Overall, the results of these calculations suggest that ERM surgery is a very cost-effective procedure when compared with other interventions across medical subspecialties.

APPENDIX 1. QUALITY OF OPHTHALMIC CARE CORE CRITERIA

Providing quality care is the physician's foremost ethical obligation, and is the basis of public trust in physicians. AMA Board of Trustees, 1986

Quality ophthalmic care is provided in a manner and with the skill that is consistent with the best interests of the patient. The discussion that follows characterizes the core elements of such care.

The ophthalmologist is first and foremost a physician. As such, the ophthalmologist demonstrates compassion and concern for the individual and utilizes the science and art of medicine to help alleviate patient fear and suffering. The ophthalmologist strives to develop and maintain clinical skills at the highest feasible level, consistent with the needs of patients, through training and continuing education. The ophthalmologist evaluates those skills and medical knowledge in relation to the needs of the patient and responds accordingly. The ophthalmologist also ensures that needy patients receive necessary care directly or through referral to appropriate persons and facilities that will provide such care, and he or she supports activities that promote health and prevent disease and disability.

The ophthalmologist recognizes that disease places patients in a disadvantaged, dependent state. The ophthalmologist respects the dignity and integrity of his or her patients, and does not exploit their vulnerability.

Quality ophthalmic care has the following optimal attributes, among others.

- The essence of quality care is a meaningful partnership relationship between patient and physician. The ophthalmologist strives to communicate effectively with his or her patients, listening carefully to their needs and concerns. In turn, the ophthalmologist educates his or her patients about the nature and prognosis of their condition and about proper and appropriate therapeutic modalities. This is to ensure their meaningful participation (appropriate to their unique physical, intellectual and emotional state) in decisions affecting their management and care, to improve their motivation and compliance with the agreed plan of treatment, and to help alleviate their fears and concerns.
- The ophthalmologist uses his or her best judgment in choosing and timing appropriate diagnostic and therapeutic modalities as well as the frequency of evaluation and follow-up, with due regard to the urgency and nature of the patient's condition and unique needs and desires.
- The ophthalmologist carries out only those procedures for which he or she is adequately trained, experienced and competent, or, when necessary, is assisted by someone who is, depending on the urgency of the problem and availability and accessibility of alternative providers.
- Patients are assured access to, and continuity of, needed and appropriate ophthalmic care, which can be described as follows.
 - The ophthalmologist treats patients with due regard to timeliness, appropriateness, and his or her own ability to provide such care.
 - The operating ophthalmologist makes adequate provision for appropriate pre- and postoperative patient care.
 - When the ophthalmologist is unavailable for his or her patient, he or she provides appropriate alternate ophthalmic care, with adequate mechanisms for informing patients of the existence of such care and procedures for obtaining it.
 - The ophthalmologist refers patients to other ophthalmologists and eye care providers based on the timeliness and appropriateness of such referral, the patient's needs, the competence and qualifications of the person to whom the referral is made, and access and availability.

- The ophthalmologist seeks appropriate consultation with due regard to the nature of the ocular or other medical or surgical problem. Consultants are suggested for their skill, competence, and accessibility. They receive as complete and accurate an accounting of the problem as necessary to provide efficient and effective advice or intervention, and in turn respond in an adequate and timely manner.
- The ophthalmologist maintains complete and accurate medical records.
- On appropriate request, the ophthalmologist provides a full and accurate rendering of the patient's records in his or her possession.
- The ophthalmologist reviews the results of consultations and laboratory tests in a timely and effective manner and takes appropriate actions.
- The ophthalmologist and those who assist in providing care identify themselves and their profession.
- For patients whose conditions fail to respond to treatment and for whom further treatment is unavailable, the ophthalmologist provides proper professional support, counseling, rehabilitative and social services, and referral as appropriate and accessible.
- Prior to therapeutic or invasive diagnostic procedures, the ophthalmologist becomes appropriately conversant with the patient's condition by collecting pertinent historical information and performing relevant preoperative examinations. Additionally, he or she enables the patient to reach a fully informed decision by providing an accurate and truthful explanation of the diagnosis; the nature, purpose, risks, benefits, and probability of success of the proposed treatment and of alternative treatment; and the risks and benefits of no treatment.
- The ophthalmologist adopts new technology (e.g., drugs, devices, surgical techniques) in judicious fashion, appropriate to the cost and potential benefit relative to existing alternatives and to its demonstrated safety and efficacy.
- The ophthalmologist enhances the quality of care he or she provides by periodically reviewing and assessing his or her personal performance in relation to established standards, and by revising or altering his or her practices and techniques appropriately.
- The ophthalmologist improves ophthalmic care by communicating to colleagues, through appropriate professional channels, knowledge gained through clinical research and practice. This includes alerting colleagues of instances of unusual or unexpected rates of complications and problems related to new drugs, devices or procedures.
- The ophthalmologist provides care in suitably staffed and equipped facilities adequate to deal with potential ocular and systemic complications requiring immediate attention.
- The ophthalmologist also provides ophthalmic care in a manner that is cost effective without unacceptably compromising accepted standards of quality.

Reviewed by: Council Approved by: Board of Trustees October 12, 1988

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APPENDIX 2. INTERNATIONAL STATISTICAL CLASSIFICATION OF DISEASES AND RELATED HEALTH PROBLEMS (ICD) CODES

Epiretinal membrane and vitreomacular traction, which include entities with the following ICD-9 and ICD-10 classifications:

	ICD-9 CM	ICD-10 CM	
Epiretinal membrane	362.56	H35.371	
		H35.372	
		H35.373	
Vitreomacular traction, adhesion	379.27	H43.821	
		H43.822	
		H43.823	

ICD = International Classification of Diseases; CM = Clinical Modification used in the United States

Additional information for ICD-10 codes:

- For bilateral sites, the final character of the codes in the ICD-10 CM indicates laterality. If no bilateral code is provided and the condition is bilateral, separate codes for both the left and right side should be assigned. Unspecified codes should be used only when there is no other code option available.
- When the diagnosis code specifies laterality, regardless of which digit it is found in (i.e., 4th digit, 5th digit, or 6th digit):
 - Right is always 1
 - Left is always 2
 - Bilateral is always 3

LITERATURE SEARCHES FOR THIS PPP

Literature searches of the PubMed database were conducted on March 6, 2023; the search strategies are listed below. Specific limited update searches were conducted on January 23, 2024 and August 6, 2024. The searches had added filters for human, English-language randomized controlled trials and systematic reviews and date limiters to capture literature published since June 1, 2019. The Retina/Vitreous PPP Committee analyzed 1,274 studies of which 18 were included in the PPP. The literature searches with the disease condition and the search terms patient values and patient preferences yielded 11 studies. The literature searches for economic evaluation and treatment cost yielded 0 studies.

Cost Benefit: ("Epiretinal membrane/economics"[MeSH]) OR ("Epiretinal Membrane"[MeSH] AND "Cost-Benefit Analysis"[MeSH]) NOT "Cost of Illness"[MeSH]

Cost of Illness: ("Epiretinal membrane" [MeSH] OR "epiretinal membrane" [tiab]) AND "Cost of Illness" [MeSH]

Diagnosis: "Epiretinal Membrane/diagnosis" [MeSH]

Epidemiology: "Epiretinal Membrane/epidemiology" [MeSH] OR "Epiretinal Membrane/ethnology" [MeSH]

Pathology: "Epiretinal Membrane/pathology"[MeSH] OR "Epiretinal Membrane/physiology"[MeSH] OR "Epiretinal Membrane/physiopathology"[MeSH]

Patient Values and Preferences: ("Epiretinal Membrane"[MeSH] OR "epiretinal membrane"[tiab]) AND (("patient values"[tiab] OR "patient preferences"[tiab]) OR (patient[tiab] AND (values[tiab] OR preferences[tiab])))

Quality of Life: ("Epiretinal membrane/therapy"[MeSH] OR "epiretinal membrane"[tiab]) AND "Quality of life"[MeSH]

Risk Factors: ("Epiretinal membrane" [MeSH] OR "epiretinal membrane" [tiab]) AND "Risk Factors" [MeSH]

Therapy: ("Epiretinal Membrane/surgery"[MeSH] OR "Epiretinal Membrane/therapy"[MeSH] OR "epiretinal membrane/drug therapy"[MeSH])



From: Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. BMJ 2021;372:n71. doi: 10.1136/bmj.n71

For more information, visit: <u>http://www.prisma-statement.org/</u>

RELATED ACADEMY MATERIALS

Basic and Clinical Science Course

Retina and Vitreous (Section 12, 2024-2025)

Patient Education

Face-Down Recovery After Retinal Surgery Brochure (2024) Macular Pucker Brochure (2024) Retina Patient Education Video Collection (2024) Vitrectomy Surgery Brochure (2024)

Preferred Practice Pattern Guidelines - Free download available at www.aao.org/ppp

Comprehensive Adult Medical Eye Evaluation (2020)

To order any of these products, except for the free materials, please contact the Academy's Customer Service at 866.561.8558 (U.S. only) or 415.561.8540 or <u>www.aao.org/store</u>.

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