

REVIEW

# Myopic Traction Maculopathy: Diagnostic and Management Strategies

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**Keywords:** pathological myopia, posterior staphyloma, retinal detachment, full thickness macular hole, macular buckle, pars plana vitrectomy, myopic traction maculopathy

#### Introduction

Myopia is a common eye disorder that causes visual disability throughout the world. The severe form, called High Myopia (HM), is defined as a spherical equivalent (SE) of more than – 6.0 diopters (D) or as an axial lengths (AL) of more than 26.5 millimeters (mm). The cut-off of – 6.0 D is still controversial, with some authors that require a SE higher than –8 D. Pathologic myopia (PM) defines high myopia associated to degenerative changes in the sclera, choroid and retina. Lewis et al in 2014, by examining the scleral biomechanics of enucleated eyes of young chicks exposed to myopia-inducing and myopia-recovery conditions, found that there are two mechanisms that contribute to scleral deformation: elasticity and creep. The first is the reversible, instantaneous deformation of the sclera, due to an applied stress; the second is the time-dependent deformation, due to persistent mechanical stress. The dysregulation of the elongation of the eye in myopia seems to be related to the inhibition of collagen crosslinking, that normally

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influences, in the mammalian sclera, the regulation of the refractive development.<sup>6</sup> This phenomenon can explain the reduced scleral rigidity in myopic eyes compared to emmetropic and hyperopic ones.7 The deformation determines an equatorial region expansion and an elongation and enlargement of the globe. The first process is directly related to the incidence of peripheral retinal degenerations like white without pressure, pigmentary degeneration, pavingstone degeneration and lattice degeneration.8 The second one induces an ectasia of the sclera, called staphyloma, which affects mainly the posterior pole (posterior staphyloma, PS) involving the optic nerve (ON) and/ or the macular area. A staphyloma may also occur laterally.

### Posterior Staphiloma

A feature that strongly suggests the presence of PM is the staphyloma, firstly described by Scarpa in 1801. 9,10 Curtin, in 1977, sorted these lesions into 10 different types, on the base of their ophthalmoscopic appearance. 11 The first five types were defined as primary staphylomas, the types from VI to X as combined ones. Posterior pole staphyloma (type I) was the most common type, involving macula and optic nerve, peripapillary (type III) and inferior staphylomas (type V) were the least common ones. Legal blindness was found to be more prevalent in eyes with staphyloma types III, VI and VII and in eyes with diffuse chorioretinal atrophy affecting the peripapillary area. 11 The most recent definition of staphyloma was proposed by Spaide as "an outpouching of the wall of the eye that has a radius of curvature that is less than the surrounding curvature of the wall of the eye". 12 This definition does not completely cover all the possible types of staphyloma: the peripapillary and nasal ones do not show an abrupt change in terms of slope of the sclera, in comparison to the surrounding scleral profile, and rather create a nasal distortion of the globe.<sup>13</sup> The staphyloma was originally considered a characteristic lesion of MM. More recently, several authors have reported an association between MTM, chorioretinal atrophy and staphyloma, supporting the hypothesis that the staphyloma itself could be considered one of the causes of MM inducing a mechanical damage to retina and ON.8,13-20 However, both MM and MTM can be present in absence of a staphyloma. A step forward in terms of staphyloma classification was made with the advent of modern imaging techniques, like three-dimensional magnetic resonance imaging (3D MRI), optical coherence tomography (OCT) and wide-field fundus imaging. These diagnostic

overcoming the limits of traditional Ultrasonography (US), that does not allow to examine the area of a large staphyloma, permitted to visualize the shape of the entire eye. Ohno Matsui et al evaluated, by 3D MRI reconstruction of the ocular globe, the shape of highly myopic eyes, dividing them into symmetric and asymmetric globes. Symmetric globes were further subdivided in barrel and cylinder types, asymmetric ones in nasal and temporal distortion types.<sup>21</sup> The same authors first defined four OCT patterns of curvature of the inner scleral surface of highly myopic eyes: a pattern with the curvature that slopes towards the ON, a pattern with symmetrical curvatures centered on the fovea, a pattern with asymmetrical curvatures and a pattern with irregular ones.<sup>22</sup> A combination of 3D MRI, OCT and wide-field fundus imaging was finally exploited by Ohno Matsui et al, to develop a new classification, by examining a total of 105 Asiatic patients with PM.<sup>13</sup> The authors classified 5 patterns of PS: wide macular, narrow macular, peripapillary, nasal, inferior and other staphylomas. A similar study was also conducted on Caucasian patients, combining A and B-scan US, 3D MRI, OCT, fundus autofluorescence (FAF), red free (RF) and color fundus photography, confirming that there is an association between the type of PS and MRI shape pattern of the eye, OCT macular profile pattern and chorioretinal atrophy.<sup>23</sup> In 2017, Shinohara et al used widefield OCT (WF OCT) combined to 3D-MRI to examine 100 eves of 57 patients affected by PS. The authors found that WF OCT could provide high-resolution images, potentially replacing 3D-MRI in assessing PS.<sup>24</sup>

# Myopic Maculopathy

Curtin and Karlin, in 1970, described 5 fundus changes associated with PM: optic nerve crescent, chorioretinal atrophy, central pigment spot (Fuchs's), lacquer cracks (Lc), and PS.<sup>25</sup> These findings were further implemented by Tokoro et al, that classified the lesions of myopic maculopathy (MM) into four categories: tessellated fundus, diffuse chorioretinal atrophy, patchy chorioretinal atrophy and macular haemorrhage, the last one associated or not to myopic choroidal neovascularization (mCNV).<sup>26</sup> In 1984, Avila et al proposed a classification on the basis of the increasing severity from a normal appearing posterior pole (M0) to choroidal pallor and tessellation (M1), with further addition of PS (M2), Lc (M3), focal areas of deep choroidal atrophy (M4) and large geographic areas of deep chorioretinal atrophy and bare sclera (M5).<sup>27</sup> In 2010, Hayashi et al proposed a new classification based on the long-term

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progression pattern of MM by analysing 806 highly myopic eyes of 429 patients. Differently to Tokoro's, Hayashi et al defined Lc as a stand-alone lesion, not included in the category of diffuse atrophy. Tokoro divided macular haemorrhage as associated or not to mCNV, whereas Hayashi included haemorrhages into the category of mCNV or Lc. Finally, Hayashi divided patchy atrophy into three subtypes: associated with Lc. diffuse atrophy or PS. 28 Since none of these classification schemes became globally accepted, an international group of experts in HM developed, on the base of a meta-analysis of PM (META-PM), a systematic classification with 5 different categories of MM: no myopic retinal lesions (category 0), tessellated fundus only (category 1), diffuse chorioretinal atrophy (category 2), patchy chorioretinal atrophy (category 3) and macular atrophy (category 4). The presence of Lc, mCNV, and Fuch's spot was added as 'plus' sign.<sup>29</sup> A correlation between the presence of the atrophic lesions, the neovascular lesions and the tractional ones and the degree of PS had been previously hypothesized by several authors. Steidl and Pruett reported a correlation between PS and the presence of atrophy or Lc.<sup>30</sup> Ohno Matsui et al reported a correlation between the width of PS and chorioretinal atrophy and tractional alterations. 13 mCNV seems to be more common in eyes with less advanced PS, because of a better preserved choriocapillaris, but Ishida et al found, in cases of defect of Bruch's membrane, a possible communication between mCNV and short posterior ciliary arteries.<sup>31</sup> Recently, Medrano et al proposed a new definition of MM as macular alterations induced by PM, in which an excessive AL and/ or PS is the main common factor but not the only factor.

These alterations included not only the atrophic lesions but also the tractional and neovascular ones. The new ATN classification of MM was based on three faces of the disease: atrophic MM (A0-no myopic retinal lesions; A1-tessellated fundus only; A2-diffuse chorioretinal atrophy; A3-patchy chorioretinal atrophy; A4-complete macular atrophy), neovascular MM (N0- no mCNV; N1-macular Lc; N2a-active CNV; N2b-scar or Fuch's spot) and tractional MM (T0-no macular schisis; T1-inner or outer foveoschisis; T2-inner + outer foveoschisis; T3-foveal detachment; T4-full-thickness macular hole; T5-macular hole + retinal detachment).

# **Myopic Traction Maculopathy** Definition

For years, the key factors defining MM were PS and the fundus changes. More recently, the focus was moved on

all the tractional features of PM thanks to the development of OCT.

In 2014 Panozzo coined the term myopic traction maculopathy (MTM) to describe the spectrum of foveal tractional changes in highly myopic eyes.<sup>32</sup> MTM included the following alterations: foveoschisis/maculoschisis/retinoschisis (FS/MS/RS), retinal/foveal detachment (RD/FD), lamellar macular holes (LMH) and full-thickness macular holes (FTMH) with (MHRD) or without RD.<sup>32</sup> They analysed the OCT images of 125 eyes with HM, focusing on epiretinal tractions (epiretinal membrane, ERM, and vitreomacular traction, VMT) and retinal damage (retinal thickening, MS/RS, RD, LMH), finding a prevalence of epiretinal traction of 46.4% and retinal damage in 34.4% of eyes (Figure 1).

### **Pathogenesis**

The development of MTM was described to be caused by two different groups of forces that act on the retina: preretinal and subretinal factors.<sup>33</sup> Preretinal factors included forces that cause centrifugal and tangential tractions to the retina. such as incomplete posterior vitreous detachment (PVD), VMT and ERM. Subretinal factors included forces that cause centrifugal tractions to the retina such as deformation of the scleral eyewall. Both can potentially damage the retina if they overcome its elasticity, which is limited by small retinal vessels, thickened or stiffened internal limiting membrane (ILM) or vitreous remnants. 3,17,34,35 Additionally, the reduced retinal blood supply due to choroidal atrophy and retinal pigment epithelium (RPE) atrophy was hypothesized to contribute to reduce adhesions between the retinal nerve layers and between the photoreceptors and the RPE itself.<sup>33</sup> A factor that accelerates the creation of tractions was reported to be the partial or complete PVD, that in myopic eyes occurs earlier and more frequently than in emmetropic eyes. 36,37 In addition, in highly myopic eyes, the formation of a physiological lacuna was reported, between the vitreous cortex, in contact with the ILM, and vitreous. The lacuna was called posterior precortical vitreous pocket (PPVP) and could simulate a PVD.37

# Morphological Changes in MTM Foveoschisis

The earliest retinal alteration occurring in the progression of MTM is RS, defined as MS when involving the macula and FS when involving only the fovea. In 1959, this condition was described by Calbert Phillips, who noticed that

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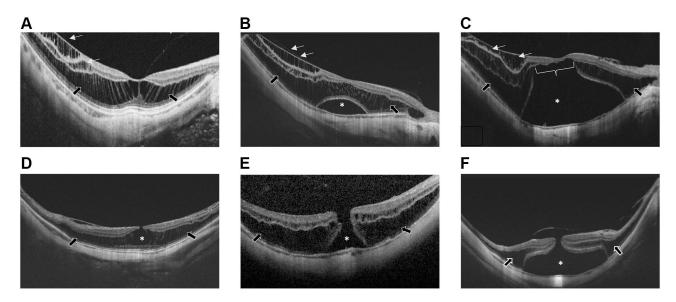


Figure 1 (A) Foveoschisis/maculoschisis/retinoschisis (FS/MS/RS). A separation of retinal layers, which remain connected by Müller cells stretched in multiple columnar structures appears in both inner retinal layers (white arrows, inner RS) and in outer retinal layers (black arrows, outer RS). (B) Foveal detachment (FD). Asterisk indicates the FD. White arrows shows the inner RS, black arrows the outer RS. (C) Retinal detachment (RD) (asterisk) associated with inner RS (white arrows) and outer RS (black arrows). White line indicates outer lamellar macular hole (O-LMH). (D) Lamellar macular hole (LMH). Asterisk indicates the partial foveal defect with intact outer retinal layers. Black arrows show the outer RS. (E) Full-thickness macular hole (FTMH). Asterisk indicates the FTMH associated with outer RS (black arrows). (F) Full-thickness macular hole with retinal detachment (MHRD). Asterisk indicates the MHRD associated with outer RS (black arrows).

localized posterior RD could occur over PS, without a detectable MH: he speculated that a condition such as FS might have explained the RD. <sup>38,39</sup> The first who used the term 'myopic foveoschisis' were Takano and Kishi, in 1999, who described the splitting of the macular inner retinal layers, separating the retina into a thinner outer layer and a thicker inner layer. 40 Baba et al, in 2003, found a correlation between the presence of FS and PS, assuming that FS could be due to scleral protrusion, overcoming the stretching capacity of the retina. 15 Benhamou et al described FS as a result of thickening in the outer retinal layer and characterized by perpendicular columns of tissue bridging the outer and inner layers. 14 The evolution was described as a progressive separation of retinal layers, which remained connected by Müller cells stretched in multiple columnar structures and appearing at OCT as "long, straight, highly reflective lines at the fovea and throughout the retinoschisis" area. 41 FS was accompanied by other OCT features such as ERMs, detachment of ILM, retinal microfolds, ellipsoid zone (EZ) line defects, paravascular microholes, LMHs, FTMHs, chorioretinopathy. 42

Several classifications have been proposed for FS. Shimada et al proposed 5 categories of outer FS based on the location and extension of it, ranging from no apparent FS (S0), extrafoveal FS (S1), only foveal FS (S2), foveal but not involving the entire macula FS (S3), FS with complete

macular involvement (S4).<sup>3</sup> The FS can involve different retinal layers: inner FS involves the inner plexiform layer (IPL), the ganglion cell layer (GCL), the retinal nerve fibre layer (RNFL); outer FS involves the outer plexiform (OPL) and the outer nuclear layers (ONL). Fujimoto et al and Ceklic et al classified the FS on the basis of the location of the splitting of the retina into inner, outer, inner and outer FS. 41,43

#### Foveal/Retinal Detachment

FS can evolve into FD. The progression tends to be slow. In some cases, FS may remain stable or even reduce, if the tractions are spontaneously relieved. 44 Shimada et al, by following 207 high myopic eyes for at least 24 months, described a decrease or resolution of the FS in 3.9% of eyes and a progression of MTM in 11.6% of eyes, noticing that eyes with a more extensive FS tended to progress more than eyes with a less extensive one.<sup>3</sup> In another study, the same author described 4 different OCT stages of the progression from FS to RD: stage 1, an irregularity of the external retinal layer thickness with focal elevation of the retina; stage 2, an outer LMH (O-LMH) that developed foveally or extrafoveally, associated with a small RD; stage 3, the O-LMH increased vertically and separated horizontally the columnlike structures within the RS layer, with an enlargement of RD; stage 4, the RD further enlarged and the RS resolved. 45

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### Lamellar Macular Hole

LMH is characterized by an irregular foveal contour, an inner retinal defect with or without intraretinal splitting and the absence of full-thickness retinal foveal defect, defined by an intact foveal photoreceptor layer. 46 In some cases, however, a disruption of outer retinal layers, external limiting membrane (ELM) and EZ has been described. 47-53 LMHs in non-myopic eyes are associated with two types of ERM: conventional (commonly detected in macular pucker), that tomographically appears as a highly reflective line overlying the RNFL and with tractional properties; atypical, a thick membrane delimited by a highly reflective line and filled by moderately reflective material, without tractional properties. 49 The same occurs in LMHs in myopic eyes. 54 These two types of ERM show a different histological composition.<sup>49</sup> Furthermore, there are two morphological types of LMH: LMH with intraretinal splitting (IR split LMH) and V-shaped LMH (V LMH). 54,55 There is still debate whether myopic LMH is a stable condition or not: dell'Omo et al, in a retrospective observational case series of 44 myopic eyes with LMH, found that these entities are frequently associated with ERMs and LMH-associated epiretinal proliferation (LHEP) and reported a morphological and functional stability over years. 56 By contrast, in a retrospective observational longitudinal study, Frisina et al studied 40 myopic eyes, affected by LMH and PS, and found that myopic LMHs, associated with atypical ERM, are a more severe entity than myopic LMH associated with conventional ERM and that myopic LMHs do not seem to be a stable condition 54,55 and evolve in FTMH. LMH can be further divided into inner LMH (I-LMH), with splitting of the inner foveal layers, and O-LMH, with splitting of photoreceptors.<sup>57</sup>

# Macular Hole and Macular Hole Retinal Detachment

Myopic FTMH is considered as one of the final stages of MTM and is associated with severe visual impairment. Ikuno et al observed in 44 eyes with myopic FS that the natural course of FS is from RS to MH to MHRD and that the foveal status is a prognostic factor for surgical success after vitrectomy and ILM peeling, with better outcomes in RS group and worse outcomes in MH group. <sup>58</sup> According to Ikuno et al, there are two types of myopic MH: the "flat type" and the "schisis type." The first one is morphologically similar to the idiopathic MH (iMH) and does not usually evolve to MHRD. The "schisis type" is

characterized by a high and rectilinear wall with an acute angle to the RPE line, i.e. acute-angled edge at the top shorter than the base diameter. The "schisis type" is associated with an high risk of MHRD.<sup>59</sup> Protective factors to MHRD are all those factors that reduce the subretinal forces: a reduced height of PS, dome-shaped macula (DSM) and greater choroidal and scleral thickness.<sup>60</sup>

# Natural History and the MSS Classification

Parolini et al elaborated the new MTM Staging System (MSS) with the aim of offering information on the complete nomenclature, the pathogenesis, the natural history and the management of the disease.<sup>57</sup> They collected OCT images of 281 eyes with MTM, found all the possible OCT types of retinal and foveal tractional changes, correlated it with best corrected visual acuity (BCVA) and age of the patients and studied the evolution of MTM by examining the changes in OCT of 126 eyes with MTM over 11 years. They avoided the term FS and choose to use MS because, in most cases, the schisis affected the whole macula and not only the fovea. The authors hypothesized that the myopic tractional changes in retina and fovea were not simply different types of MTM, but evolutive stages of the same disease. They explained the pathogenesis of MTM, describing different centrifugal forces that act on retina and fovea playing against the centripetal forces that hold the retina together, exerted by Müller cells, ELM and ILM. On the basis of these theories, Parolini et al described two evolution patterns, foveal and retinal ones. The retinal patterns, due to centrifugal forces perpendicular to the retinal plane, evolve from inner MS (I-MS) or inner and outer MS (IO-MS) in stage 1, to predominantly outer MS (O-MS) in stage 2, to MS with macular detachment (MS-MD) in stage 3, to macular detachment (MD) in stage 4. The foveal patterns, due to centrifugal forces tangential to the foveal plane, evolve from normal foveal profile in stage a, to I-LMH in stage b, to FTMH in stage c. The O-LMH and any epiretinal abnormalities are findings that can be associated with each pattern and are marked in the MSS, respectively, with "O" and "+". Parolini et al found that the mean evolution time of MTM gradually decreased from stage 1 to 2 (20 months), stage 2 to 3 (12 months) and 3 to 4 (3 months), with BCVA decreasing with increasing stages.<sup>57</sup> There is also a correlation between MTM stage and age.

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## **Surgical Management**

The history of the ab externo approach for MTM begins long before the definition of MTM itself, in 1930. The idea was to reinforce the posterior sclera limiting the progressive myopic elongation of the eye. Different types of material, such as fascia lata, 45 donor sclera, 61,62 and Lyodura, derived from processed cadaver dura mater, 63 were used for this this purpose. Fifty years later, a modified posterior scleral reinforcement (PSR) technique was proposed by Snyder and Thompson and further developed by several authors. 64-67 Finally, the macular buckle (MB) technique was firstly described by Schepens, Okamura and Brockhurst in 1957<sup>68</sup> and for long time considered as the best surgical approach for the treatment of myopic MHRD. 68-77 The rationale of MB was to overcome the pathologic posterior bulging and stretching of the sclera, especially in eyes with a pronounced PS, which is an important risk factor for the development of MTM. 78,79 The aim of it is to relieve the extern traction due to PS. On the other hand, considering DSM as a protective factor for the occurrence of MTM, the MB provides, in fact, an iatrogenic DSM.

After a long time-length, in which the ab externo approach was overwhelmed by the ab interno approach, the MB, with the use of a sponge or a solid silicone exoplant, returned back in vogue in the 2000s, 80-82 showing a very high reattachment rate, but still technically challenging. Attention was pointed to facilitate this surgery, with the design of different types of buckle. A T-shaped semirigid rod-exoplant, made with silicone and reinforced with titanium and with an indenting head was published by Tanaka, Ando and Usui in 2005.83 An L-shaped buckle, made with a silicone sleeve to embrace a titanium stent (MRI compatible), that allowed a macular indentation by a soft silicone sponge and an anterior suture, was developed in 2009 by Parolini et al. MB was reported to assure an effective axial myopia control in 2009 by Ward, 84 to have better visual outcomes compared to PPV, 85 and, in comparison with ab interno approach, the gold standard to treat MHMD, according to a 16-year review on MB for MTM of Alkabes et al.86

Pars plana vitrectomy (PPV) was introduced in the 1980s for the first time as treatment for MHRD. 87–89 Different types of intravitreal tamponades combined to PPV were proposed, like gas and silicone oil (SO). A higher anatomical success rate was reached with SO in comparison to gas, but an unsatisfactory functional success was reported by several

authors. 90-93 The introduction of surgical maneuvers such as ILM peeling<sup>94–96</sup> and laser treatment around the edges of MH had encouraging anatomical results, reaching among 90% of success rate. 97,98 The laser treatment, however, limited the functional recovery. Subsequently, PPV began to be used also for the treatment of RS, MS and RD without MH. The limits of PPV are many. The power of the tamponade is restricted by the configuration of the PS, ILM peeling could weaken an already weak retina and is not easy to see with the atrophic changes in the myopic choroid. The success rate was extremely variable between different studies. Therefore, the use PPV in MTM remained debatable. 99-104

### Summary

After several studies, MTM has been recognized not just as a group of different foveal and retinal changes in a tractional myopic environment, but as a single evolutive pathology with different stages. The same happened previously to MM as a whole, when the different aspects were found to be different stages of the same disease, only years after its first description.

An important role was played by the OCT, that allowed, on one hand, to precisely define the ultrastructural characteristics of the various features of MTM and permitted, on the other hand, to follow eyes over time, discovering the progressive nature of the disease. MTM is a diagnostic challenge because, in early stages, is often asymptomatic and difficult to identify biomicroscopically. OCT allows to detect minimal macular changes in an asymptomatic stage. In light of these observations, the Authors of the current review recommend following eyes with PM over time using OCT, in order to identify early stages of MTM, and emphasize the need to standardise the follow-up time. The proposed MSS staging system, based on OCT findings, lead to a better understanding of MTM as a whole and offered precise indications on prognosis and therefore management.

The treatment of MTM is a complex and debated topic for different reasons. In the early stages of MTM, such as FS, the surgical approach is not considered by everyone to be the right choice. Some authors, assuming that MTM is a progressive pathology, surgically treat the FS. Others, based on the slow morphological and functional progression of MTM, prefer to wait and follow the patient over time. Even in the management of advanced MTM stages, i.e. the presence of a MH, a RD or the association of both in MHRD, controversies exist. The main concern is the type of surgical approach: ab externo or ab interno. In the last decade, several MB techniques have been proposed with

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encouraging results. On the other hand, the ab interno approach, PPV, allows to perform a series of maneuvers facilitating the treatment of RD and MH. The choice between macular buckle or PPV for the treatment of MTM, is not obvious. For the treatment choice, the key is the understanding of the pathogenesis, i.e. the combination of centrifugal forces which are either tangential and/or perpendicular to the macula. The tangential forces are mainly counteracted by PPV and maneuvers on the ILM, while the perpendicular forces are mainly counteracted by the MB. Therefore, the two techniques have to be selected on the base of the MTM stage, single or combined.

### **Disclosure**

The authors report no conflicts of interest for this work.

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