| 1 | Choroidal Vascular Changes after Encircling Scleral Buckling for Rhegmatogenous Retinal | | |
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| 2 | Detachment | | |
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26 Abstract

Background/Objectives: There is an ongoing debate on whether encircling scleral buckling (SB)
procedure for the treatment of rhegmatogenous retinal detachment (RRD) may cause an impairment
in choroidal blood flow. The aim of this study was to compare choroidal vascularity index (CVI) and
subfoveal choroidal thickness (CT) between eyes that had undergone encircling SB with unoperated
fellow eyes (FEs).

Subjects/Methods: Thirty patients treated with encircling SB for unilateral RRD were included.
Demographic and clinical characteristics as well as Enhanced Depth Imaging – Optical Coherence
Tomography scans were retrospectively collected. Images were binarised using ImageJ software,
total choroidal area along with luminal and stromal area (respectively, TCA, LA and SA) were
segmented and the CVI was computed as the ratio of LA/TCA. In addition, CT was evaluated.

Results: The mean follow-up interval between surgery and examination was 25.5 ± 16.8 months. Choroidal thickness, TCA, LA, and SA were significantly increased in the operated eyes compared to FEs (respectively, $271.7\pm78.0\mu$ m vs 238.5 ± 83.4 , P=0.001; 1.804 ± 0.491 mm² vs 1.616 ± 0.496 , P=0.001; 1.199 ± 0.333 mm² vs 1.067 ± 0.337 , P<0.001 and 0.605 ± 0.171 mm² vs 0.550 ± 0.171 , P=0.001). Conversely, CVI did not significantly differ between the two groups (66.4 ± 3.6 vs. 65.9 ± 3.2 , P=0.490).

43 Conclusions: In conclusion, eyes treated with encircling SB for RRD presented increased LA, SA
44 and CT compared with FEs, but showed no difference in CVI.

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Key words: Encircling scleral buckling; Choroidal vascularity index; Choroidal Thickness; Choroid;
Rhegmatogenous Retinal Detachment.

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52 Introduction

53 Rhegmatogenous retinal detachment (RRD) is a sight-threatening condition caused by the 54 passage of fluid from the vitreous cavity into the subretinal space through one or more retinal breaks, 55 leading to the separation of the neurosensory retina from the underlying retinal pigment epithelium¹. 56 Encircling scleral buckling (SB) represents an effective and well-established surgical technique for 57 the treatment of selected cases of primary or recurrent RRD. The positioning of an encircling element 58 around the circumference of the eye provides the reduction of vitreoretinal traction and the closure of 59 the retinal defect(s), allowing retinal re-attachment¹. Although uncommon, complications such as 60 anterior and posterior segment ischemia and serous choroidal detachment could occur postoperatively 61 ². The mechanical force exerted by the encircling band induces changes in anatomical and functional 62 parameters of the eyeball, such as axial length, anterior chamber depth and corneal biomechanics ^{3–} 63 ⁵. Moreover, previous studies disclosed an impaired ocular circulation, reporting postoperative changes in both choroidal and retinal blood flow ⁶⁻¹⁴. These findings suggest that a subclinical 64 65 ischaemia may exists in more cases than reported.

The advancements in spectral coherence optical coherence tomography (SD-OCT) technology made it possible to visualize the choroidal structure in great detail, and allowed the accurate calculation of different quantitative parameters ¹⁵. In particular, choroidal thickness (CT) has been used to investigate vascular changes after encircling SB. Although there seems to be some agreement that the encircling element affects CT, the results are somewhat contradictory ^{13,16–19}.

71 The recent introduction of binarisation algorithms on enhanced depth imaging SD-OCT (EDI SD-

72 OCT) images allows to separately investigate the vascular and stromal choroidal components 20 .

In particular, choroidal vascularity index (CVI), which is the proportion of the luminal area (LA) to the cross-sectional choroidal area, represents a novel choroidal biomarker that has been used to monitor choroidal status in several chorioretinal disorders ^{21–25}. Compared to CT, CVI seems to be less influenced by biological variables such as axial length, intraocular pressure, and diurnal variation

77 ²⁶.

The aim of this study was to compare CVI between eyes that had undergone encircling SB for
RRD and fellow eyes (FEs), and to further investigate possible correlations with clinical parameters.

81 Subjects and Methods:

82 Study Design and Patients

83 This retrospective cross sectional study included patients treated for RRD at a single tertiary-referral 84 center (S.Orsola-Malpighi University Hospital, Bologna, Italy) between January 2016 and December 85 2019. The study was performed in accordance with the principles of the Declaration of Helsinki and 86 was approved by the local Institutional Review Board. Written informed consent was obtained from 87 all subjects included in the study. Consecutive patients underwent uneventful SB for RRD in one eye 88 were screened for enrolment. SB was performed by the same surgeon (P.G.T.) in phakic patients with 89 RRD due to single retinal break or small confluent multiple breaks without significant lens 90 opacification. Eyes with SB were included as the study eyes and the FEs served as controls. Exclusion 91 criteria were any previous ocular surgery except for SB in one eye, history of retinal diseases (e.g. 92 choroidal neovascularization, diabetic retinopathy, retinal dystrophy and central serous 93 chorioretinopathy), glaucoma, spherical equivalent ≥ 6 diopters (D), anisometropia ≥ 1.5 D, poor 94 image quality and missing data from medical records.

95 The following data were extrapolated from medical records: age, sex, preoperative status of the
96 macula, postoperative BCVA in logMAR, postoperative fundus examination and spectral domain
97 EDI-OCT scan.

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99 Encircling Scleral Buckling Procedure

100 A 360-degree limbal conjunctival peritomy incision was made and traction sutures were placed 101 beneath the insertions of the exposed rectus muscles to facilitate positioning the globe. A 2.5 mm 102 wide silicone band (No. 240) was passed around the circumference of the globe and beneath the rectus 103 muscles at a distance of 14 mm from the limbus. The band was anchored with single interrupted

104 suture with bites parallel to the limbus placed in the center of each quadrant, the ends of the band 105 were then joined in the opposite quadrant of the retinal break(s) with a silicone sleeve. In all cases, 106 drainage procedure was performed by a sclerotomy just below the retinal break(s) then sutured by a 107 single scleral stitch. Ab externo cryotherapy was done in the retinal break(s) location. In order to 108 increase the buckling effect an adjunctive biconvex silicone 9 mm wide element (No. 279) was placed 109 beneath the band above the retinal break(s). In all patients included, their extension ranged from 3 to 110 5 clock hours. The ends of the encircling silicone band were then pulled until the desired buckle effect 111 was reached. A paracentesis was done and SF6 injection was performed 4 mm posterior to the limbus. 112 There were no intraoperative complications. At the end of the procedure, indirect ophthalmoscopy 113 revealed retinal reattachment in all patients.

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115 Image Analysis

116 The SD-OCT scans were obtained in all patients using the Spectralis HRA-OCT (Heidelberg 117 Engineering, Heidelberg, Germany). Each session was performed at approximately the same time of 118 the morning, to avoid diurnal variations. Macular OCT images were acquired with EDI mode using 119 a volume scan of 30 X 10 degrees centred on the fovea, with 100 frames averaged in each scan. The 120 OCT scan passing through the central foveal region was chosen for the analysis. The choroid was 121 defined as the space between the outer border of the retinal pigment epithelium and the choroidal-122 scleral junction. The subfoveal CT was measured manually by two independent examiners (F.B. and 123 P.R.R.) using the caliper function tool of the image analysis software. The mean of the two 124 measurements was used for the analysis. The OCT images were binarised and segmented by the same 125 examiners using the public domain software ImageJ 1.51s (National Institutes of Health, Bethesda, MD), with a semiautomated method previously described ^{20,22}. Briefly, the OCT image was opened 126 127 in ImageJ and the polygon tool was used to select the region of interest across the entire length of the 128 OCT scan. The upper boundary of the region of interest was traced along the choroidal-retinal 129 pigment epithelium junction and the lower boundary along the choroidal-scleral junction to identify

130 the total choroidal area (TCA) (Figure 1, part A). After conversion to an 8-bit image, Niblack's 131 autolocal threshold was applied to binarised the image and demarcate the LA and stromal area (SA) 132 (Figure 1, part B). The image was converted back to a red, green, blue image, and the colour threshold 133 tool was used to select the dark pixels, representing the LA (Figure 1, part C). The TCA and LA were 134 measured. The SA was calculated by subtracting LA from TCA. The CVI, defined as the LA divided 135 by the TCA, was then computed. The choroidal parameters calculation was performed separately by 136 2 investigators, both blinded for patients' characteristics (F.B. & M.P.), and the mean value for each 137 parameter calculated was used for the statistical analysis.

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139 Statistical Analysis

The R studio software, version 1.2.5042, (http://www.r-project.org) was used for data analysis. Values are expressed as mean \pm standard deviation. The Shapiro-Wilk test was used to determine normality of data. The Student's t-test was used to compare normally distributed continuous variables between operated eyes (OEs) and FEs, whereas the Wilcoxon signed-rank test was used for not normally distributed variables. The correlations of choroidal parameters with demographic and clinical parameters were examined using the Pearson correlation analysis with a Bonferroni correction for multiple comparisons. A *P* value < 0.05 was considered statistically significant.

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148 **Results**

Thirty-nine patients treated with SB for RRD were initially identified. Of these, 9 patients did not satisfy the inclusion/exclusion criteria and were excluded from the final analysis. In particular, 2 presented high myopia, 3 had previous eye surgery in both eyes, 1 had glaucoma, 1 was lost to follow up and 2 presented poor OCT scans quality. Finally, 30 patients were enrolled in the study. Mean age was 61.7 ± 6.1 years (range 51-75 years), and 19 of the patients (63.3%) were male. Eighteen patients (60%) presented macula off RRD at the time of the procedure. The mean time interval between the surgery procedure and the follow-up visit was 25.5 ± 16.8 months (range 4–59 months). Mean BCVA 156 was 0.05 ± 0.07 logMAR in the OEs. Macular pucker was present in 9 of the OEs (30%) at follow-157 up visit. Choroidal parameters in eyes treated with encircling SB for RRD and FEs are reported in Table 1. Choroidal thickness, TCA, LA, and SA were significantly increased in the OEs compared 158 159 with FEs (respectively, $271.7 \pm 78.0 \,\mu\text{m}$ vs. 238.5 ± 83.4 , P = 0.001; $1.804 \pm 0.491 \,\text{mm}^2$ vs. $1.616 \pm$ 160 $0.496, P = 0.001; 1.199 \pm 0.333 \text{ mm}^2 \text{ vs. } 1.067 \pm 0.337, P < 0.001 \text{ and } 0.605 \pm 0.171 \text{ mm}^2 \text{ vs. } 0.550$ ± 0.171 , P = 0.001). On the contrary, CVI did not significantly differ between the two groups (66.4) 161 \pm 3.6 % vs. 65.9 \pm 3.2, P = 0.490). No significant correlations between choroidal parameters and 162 163 demographical and clinical parameters, including follow-up period, were found (always P > 0.05).

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165 Discussion

In the present study we investigated the choroidal structure in eyes treated with encircling SB after a relatively long term follow up. The evaluation was made using a previously validated technique, which allows to evaluate separately the stromal and the luminal choroidal components ²⁰. We found that both LA and SA were increased in the OEs compared with FEs. Interestingly, the CVI, that represents the ratio of LA to TCA, was not different between the two groups, suggesting that both LA and SA increased without changing their proportions.

172 It has been speculated that the mechanical force exerted by encircling and buckling elements 173 could determine a venous drainage obstruction, inducing an increased vascular resistance and thus, a 174 decreased ocular blood flow ⁶. Previous studies investigated ocular blood flow after SB procedure 175 using different techniques, including ocular pulse measurements, bidirectional laser Doppler, laser speckle flowmetry and color Doppler imaging, disclosing an impaired chorioretinal circulation ⁶⁻¹⁴. 176 177 However, mixed results on its changes over time have been reported. Some studies showed that the reduction of chorioretinal blood flow still persisted after a long term follow-up and improved only 178 after the removal of the encircling band ^{6,9–11}. Our results could support this hypothesis, since the 179 180 increase in LA had previously been interpreted as an indirect marker of blood flow stasis secondary to an impaired circulation ²⁴. Conversely, others reports showed a post-operative reduction in 181

182 choroidal blood flow that returned to baseline values 3 to 6 months after surgery ^{12,14}. It is difficult to 183 compare the results of these studies because they differ in terms of surgical technique, follow-up 184 period and measurement methods.

185 Recently, CT has been evaluated in patients after SB as a surrogate marker of choroidal blood 186 flow ^{13,16–19}. Previous studies indicated that segmental SB procedure, characterised by the placement 187 of a local buckling element, induces a temporary increase of subfoveal CT that returns to baseline 188 values 1 to 3 months after surgery ^{13,18,19}. Conversely, the use of encircling band in addition to the 189 local buckling element seems to lead to a long term increase in CT ^{17,27}.

190 Several factors might be related with the development of choroidal changes, including the 191 material of the buckling element and its extent (length, width and thickness), along with the use of cryotherapy, which can induce a post-operative inflammatory response ^{13,19}. We found that OEs 192 193 showed an increased CT along with a greater TCA compared with FEs. These results are consistent 194 with Odrobina and collaborators who disclosed an increased CT in patients treated with a 3.5 mm encircling SB after a mean follow-up of 22 months ¹⁷. These findings suggest that different 195 196 mechanisms might occur depending on whether an encircling band has been used or not. In fact, it is 197 reasonable to hypothesize that the presence of an encircling element may induce an impairment of 198 choroidal circulation over a longer period of time and may reduce the possibility of choroidal tissue 199 to return to baseline status.

The absence of change in CVI between OEs and FEs could be explained in two different ways: on the one hand, also the stromal component of the choroid might be affected by the placement of the encircling band; on the other hand, an adaptation process of the stromal tissue might occur following vascular changes. However, these hypotheses remain speculative and a prospective study is required to evaluate when stromal change occur.

Interestingly, none of the patients included presented any complication related with vascular impairment following the surgery. Moreover, all of them experienced an overall good prognosis in terms of visual acuity. According to a previous report, although an alteration of the choroidal blood flow seems to be common after encircling SB procedure, patients generally do not experience longterm visual complications. These complications could be related with an individual susceptibility or with the degree of tightness of the encircling band 6 .

This study has some limitations that should be taken into account. The main limit is related to its retrospective design, which hampered the evaluation of choroidal parameters before and after encircling SB, as well as the monitoring of their trend over time. In addition, the algorithm used for the calculation of choroidal parameters does not allow the evaluation of the different choroidal layers, namely the choriocapillaris, the medium choroidal vessel layer and the large choroidal vessel layer, thus it was not possible to determine their respective degree of involvement.

In conclusion, eyes treated with encircling SB for RRD show increased luminal and stromal areas along with a greater choroidal area, compared to FEs. Furthermore, OEs show no difference in CVI compared to FEs, suggesting that the increase in stromal and vascular components maintained their proportion unchanged. These results support the theory that encircling SB induces subclinical changes in choroidal circulation and provide a deeper characterisation of this phenomenon.

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312 Figure Legends

Figure 1. Choroidal vascularity index calculation with binarisation of OCT images in a representative eye treated with encircling scleral buckling for rhegmatogenous retinal detachment. A. Choroidal boundaries are traced to identify the total choroidal area (orange lines). B. The image is binarised with Niblack's auto-local threshold. C. The colour threshold tool is used to select the dark pixels, representing the luminal area (yellow lines). The choroidal vascularity index is obtained dividing luminal area by total choroidal area.

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Table 1. Choroidal parameters in eyes treated with encircling scleral buckling for rhegmatogenous retinal detachment and fellow eyes

| Parameter | Operated Eye | Fellow Eye | Р |
|------------------------|-------------------|-------------------|---------|
| Subfoveal CT (µm) | 271.7 ± 78.0 | 238.5 ± 83.4 | 0.001 |
| TCA (mm ²) | 1.804 ± 0.491 | 1.616 ± 0.496 | 0.001 |
| LA (mm ²) | 1.199 ± 0.333 | 1.067 ± 0.337 | < 0.001 |
| SA (mm ²) | 0.605 ± 0.171 | 0.550 ± 0.171 | 0.001 |
| CVI (%) | 66.4 ± 3.6 | 65.9 ± 3.2 | 0.490 |

320 CT: choroidal thickness; TCA: total choroidal area; LA: luminal area; SA: stromal area; CVI:

321 choroidal vascularity index. Significant P values (<0.05) are in bold.

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