Effect of Using Perfluorocarbon Liquids on Postoperative Retinal Displacement in Vitrectomy for Treatment of Primary Rhegmatogenous Retinal Detachment: A Comparative Study

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Abstract

Background: Perfluorocarbon liquids (PFCLs) are widely used for years as an adjuvant in vitrectomy for repair of retinal detachment. Studies have been made to find out the postoperative effects of using PFCLs on the retina. A new imaging modality, Fundus Autofluorescence (FAF) can describe retinal displacement after vitrectomy clearly, which we could not describe it before with other imaging modalities.

Aim of Study: To compare the incidence of retinal displacement after vitrectomy for repair of primary rhegmatogenous retinal detachment (RRD) with and without the use of PFCLs using FAF, and correlate them with final best corrected visual acuity (BCVA) and postoperative symptoms of distortion if present.

Patients and Methods: A comparative, prospective, interventional, randomized study. Thirty eyes of thirty patients having primary rhegmatogenous retinal detachment were divided into 2 groups. Group A: 15 eyes underwent pars plana vitrectomy without the use of PFCLs and group B: 15 eyes underwent pars plana vitrectomy with the use of PFCLs. Fundus Autofluorescence and amsler grid were done following absorption of intraocular gas.

Results: Retinal displacement evident on FAF was (53.3%) of cases in group A (without PFCL), and (40.0%) of cases in group B (with PFCL), respectively (p-value = 0.464). PFCL use has no statistical significance in the incidence of retinal displacement. There was no statistically significant difference in the final mean BCVA between the two groups. Postoperative distortion was detected in (85.7%) of eyes showing retinal displacement on FAF.

Conclusion: Retinal displacement is common following RRD repair and the use of PFCLs does not seem to affect the incidence of displacement.

Key Words: Rhegmatogenous retinal detachment – Retinal displacement – Pars plana vitrectomy – Perfluorocarbon liquids – Fundus autofluorescence.

Introduction

The selection of the surgical approach for repair of primary rhegmatogenous retinal detachment (RRD) is the most important indicator for postoperative functional and anatomical success [1]. The essential steps of any surgery depend on accurate identification, localization and treatment of all tears [2]. Currently the most two frequent surgical techniques are scleral buckling and pars plana vitrectomy (PPV), with pneumatic retinopexy being reserved for selected cases [3]. Pars plana vitrectomy has become more popular than scleral surgery in recent years, mainly for primary rhegmatogenous retinal detachment. Intra-surgery visualization and identification of retinal tears is broadly considered being superior to pre-surgery visualization and identification. Probably this is one among the main reasons for the tendency to use this technique as the first choice for repair of primary RRD. Yoshida et al., reported in a prospective and retrospective case series study that primary reattachment rates were of 75-100% with a final rate of 96-100% [4].

The well acclaimed use of perfluorocarbon liquids (PFCLs) in vitreoretinal surgery being due to their unique physical properties as they have a high specific gravity ranging from 1.76 to 2.03, low surface tension, low viscosity and optical clarity making them an ideal intraoperative adjuvant in vitreoretinal surgery mainly to relocate and flatten the detached retina, so that improving the efficiency and safety of the surgery especially in complicated cases [8].

Fundus Autofluorescence (FAF) has been considered as a novel imaging modality to identify a
new postoperative entity following retinal detachment surgery which is retinal displacement and various studies have been directed to correlate between it and the functional outcome of surgery in terms of visual acuity and postoperative symptoms [6]. FAF elegantly depicts this shift via lines of increased autofluorescence, which reflect the caliber and orientation of the adjacent retinal vessels. These lines indicate the original location of the retinal vessels, which have been displaced with the retina after retinal reattachment. Dell’Omo et al., named these lines as retinal vessel printings (RVPs) [7]. While Lee et al., named them as RPE vessel ghost [8]. The reason why lines of increased autofluorescence indicate the preoperative site of retinal vessels is unclear. Among the explanations of this phenomenon was that stated in a study by Shiragami et al., in which they suggested that the RPE cells under the retinal vessels were prevented from light irradiation for a long time and hence they maintained dark adaptation because they were shaded by the retinal vessels. After that, acute exposure to the excitation light, which was caused by translocation of the retinal vessels, may induce increased FAF signals [6]. On the other hand, Dell’Omo et al., proposed that a different composition and characteristics of fluorophores may exist between the RPE cells belonging to the shifted vessels’ prints (normally shaded by the overlying vessels) and the neighboring cells (normally exposed to light) [7].

To date, there have been many studies about retinal displacement after vitrectomy [7,8]. However, in our study we compared the incidence of retinal displacement after vitrectomy for repair of primary rhegmatogenous retinal detachment with and without the use of PFCLs, using FAF which is a valuable simple imaging technique, and correlate them with the final best corrected visual acuity (BCVA) and postoperative symptoms of distortion if present.

Material and Methods

This study is a comparative prospective interventional randomized study. It included 30 eyes of 30 patients undergoing primary vitrectomy for repair of primary rhegmatogenous retinal detachment (RRD). The primary outcome of the study was evaluation of presence or absence of retinal displacement using FAF. Secondary outcomes were postoperative visual acuity and symptoms of distortion like metamorphopsia, if present. Patients were recruited from the outpatient clinic of the Research Institute of Ophthalmology in Giza from the period of June 2017 till April 2018. It was approved by the Institutional Review Board and was conducted in compliance with principles of Helsinki declaration.

We included patients with primary RRD (with proliferative vitreoretinopathy (PVR) grade A or B) assigned for primary vitrectomy. Cases were divided randomly into two groups A and B; each included 15 eyes: Group A: Underwent pars plana vitrectomy without the use of PFCLs. Group B: Underwent pars plana vitrectomy with the use of PFCLs.

We excluded cases with PVR more than grade B, cases that had performed previous retinal detachment surgeries; pneumatic retinopexy, scleral buckle or PPV, cases with old standing retinal detachment, exudative or tractional retinal detachment, giant retinal tears, macular holes and pediatric cases.

Baseline evaluation:

Complete ocular examination was done including BCVA (with snellen chart, which was converted to logMAR for statistical analysis), intraocular pressure (IOP) using Goldmann applanation tonometry and anterior segment biomicroscopy. The retinal detachment in all cases was evaluated using slit lamp biomicroscopy using (Super Field lens; VOLK, Mentor, Ohio, USA) and indirect ophthalmoscopy determining extent and configuration of the retinal detachment, macular status, number and position of the breaks and grade of PVR.

Surgical procedure:

In Group A cases: PPV was done using 23-gauge system (Constellation system. Alcon, Fort Worth, TX). Posterior vitreous detachment were induced, followed by core vitrectomy and assisted by the use of triamcinolone acetonide to stain the vitreous in order to achieve near complete vitreous removal, followed by shaving of the vitreous base assisted by scleral indentation. Shift to fluid air exchange, followed by drainage of subretinal fluid through retinal breaks. Finally, laser was applied around all marked breaks and augmented by 360 degrees barrage if the retina condition required this. This was followed by closure of the sclerotomies then by the use of two syringes one filled with gas and the other acting as a chimney replacing the air with the non expansile concentration (20%) of the sulphur hexafluoride (SF6) gas. Periocular antibiotic steroid combination was given and the (IOP) was adjusted.

In Group B cases: The same steps applied except that after completion of the core vitrectomy step, PFCLs were infused over the optic disc to
displace the subretinal fluid to the periphery followed by complete shaving of the vitreous base. This was followed by more injection of PFCL and then either apply the laser under PFCL or replace the PFCL by air and apply laser under air. After complete removal of PFCL, all cases were tamponaded with gas intra-operatively. Patients were advised to take the face down position immediately following the operation. All patients were prescribed topical antibiotics and steroids after the operation.

Post-operative follow-up:

Patients were followed-up on the 1st day post-operatively then after 1 week, 2 weeks, 1 month and 3 months respectively, examination included: BCVA by refraction; slit lamp examination, anterior segment biomicroscopy, IOP measurement using Goldmann applanation tonometry, dilated fundus examination with binocular slit-lamp biomicroscopy (using 90 diopters lens) and indirect ophthalmoscopy (using 20 D lens). Symptoms of distortion were subjectively recorded using Amsler grid. Imaging (FAF) was typically performed following absorption of intraocular gas tamponade using Spectralis HRA+OCT (Heidelberg Engineering, Heidelberg, Germany). B-FAF (excitation wavelength at 488nm and barrier filter at 500nm) and red-free pictures (excitation wavelength at 508nm and barrier filter at 358nm) were obtained.

Image analysis:

Postoperatively, the presence of ghost vessels and their position (upward or downward) were analyzed in relation to the adjacent retinal vessels. According to the study published by Lee et al., which defined the ghost vessels as lines of increased autofluorescence running approximately parallel to adjacent retinal vessels, separated from them and closely reflecting their caliber and orientation. The presence of ghost vessels was taken as evidence of retinal displacement [8].

Statistical analysis:

Data were collected, revised, coded and entered to the Statistical Package for Social Science (IBM SPSS) version 20. The quantitative data were presented as mean, standard deviations when their distribution found parametric while qualitative data were presented as number. The comparison between two independent groups with qualitative data was done by using Chi-square test. The comparison between two independent groups with quantitative data and parametric distribution was done by using Independent t-test. The confidence interval was set to 95% and the margin of error accepted was set to 5%. So, the p-value was considered significant as the following: p>0.05: Non-significant, p<0.05: Significant, p<0.01: Highly significant.

Results

This study included 30 eyes of 30 patients divided into 2 groups 15 eyes each: Group A: underwent PPV without the use of PFCLs, Group B: underwent PPV with the use of PFCLs. With a mean age of 56.27±11.82 for Group (A) patients and 49.13±13.09 for Group (B) patients. In Group (A) 9 eyes belonged to 9 females and the other 6 eyes belonged to 6 males, while in Group (B) 5 eyes belonged to 5 females and the other 10 eyes belonged to 10 males. The baseline characteristics of the studied groups are summarized in Table (1). Baseline characteristics of the two groups were homogeneous with no significant statistical difference between groups in terms of age and gender.

| Table (1): Baseline characteristics of the studied groups. |
|---------------------------------|---------------------------------|----------------|--|
|                                 | **Group A**                     | **Group B**                     | **P**-value | **Sig.** |
| No.= 15                         | No.= 15                         |                               |            |         |
| **Age:**                        |                                 |                               |            |         |
| Mean±SD                         | 56.27±11.82                     | 49.13±13.09                    | 0.128      | NS      |
| Range                           | 30-72                           | 25-73                          |            |         |
| **Gender:**                     |                                 |                               |            |         |
| Females                         | 9 (60.0%)                       | 5 (33.3%)                      | 0.143      | NS      |
| Males                           | 6 (40.0%)                       | 10 (66.7%)                     |            |         |
| Left                            | 6 (40.0%)                       | 10 (66.7%)                     |            |         |

SD = Standard Deviation of age mean.
NS = Not Significant.
P = Probability level.
PACL = Perfluorocarbon Liquids.

The primary anatomical success (defined as successful retinal reattachment after the first surgery) was achieved in all 15 eyes of Group A patients (100%) and in 14 of the 15 eyes of Group B patients (93.3%). On the other hand, the final anatomical success (defined as successful retinal reattachment regardless the number of procedures) was achieved in all 30 eyes of the studied groups.

After retinal reattachment retinal ghost vessels indicating retinal displacement were observed in 14 of the 30 eyes (46.6%) (8 eyes in Group A (53.3%) and 6 eyes in Group B (40.0%). There was no statistically significant difference between both groups (p-value = 0.464). The direction of retinal displacement in all of the eyes of the studied groups was downwards (100%) (Fig. 1) (Table 2).
Table (2): FAF findings in the studied groups.

<table>
<thead>
<tr>
<th>FAF Findings</th>
<th>Group A (Without PFCL)</th>
<th>Group B (With PFCL)</th>
<th>( p )-value</th>
<th>Sig.</th>
</tr>
</thead>
<tbody>
<tr>
<td>No. Retinal displacement:</td>
<td>No.</td>
<td>%</td>
<td>No.</td>
<td>%</td>
</tr>
<tr>
<td>No</td>
<td>7</td>
<td>46.7</td>
<td>9</td>
<td>60.0</td>
</tr>
<tr>
<td>Yes</td>
<td>8</td>
<td>53.3</td>
<td>6</td>
<td>40.0</td>
</tr>
<tr>
<td>Direction:</td>
<td>Down</td>
<td>8</td>
<td>100.0</td>
<td>6</td>
</tr>
</tbody>
</table>

NS = Not Significant.
P = Probability level.
PFCL = Perfluorocarbon Liquids.
NA = Not Applicable.
FAF = Fundus Autofluorescence.

Fig. (1): Post-operative FAF of a case of retinal displacement, note the hyperfluorescent lines above & parallel to the retinal vessels indicating downward displacement of the retina.

In Group (A) patients none of the 3 eyes having fovea sparing retinal detachments showed retinal displacement, while of the remaining 12 fovea-involving retinal detachments 8 eyes showed retinal displacement indicating that all cases (100%) of the group (A) eyes showing retinal displacement had a fovea-involving detachment. There was no statistically significant correlation between neither the location of breaks nor the detachment extent with the incidence of retinal displacement.

Unlike Group (A), in Group (B) patients 2 eyes having fovea-sparing retinal detachment showed retinal displacement and the remaining 4 eyes had fovea-involving detachments. However similarly to Group (A) patients, there was no statistically significant correlation between neither the location of breaks nor the detachment extent with the incidence of retinal displacement.

The mean post-operative BCVA was 0.70 ± 0.42 for Group (A) patients and 0.85 ± 0.42 for Group (B) patients during the first follow-up visit following the absorption of intra-ocular gas tamponade. BCVA was measured using the Snellen's chart and converted to LogMAR for statistical analysis. There was no statistically significant difference between both groups (\( p \)-value = 0.327).

All patients underwent screening for symptoms of distortion using Amsler grid test, 9 patients from Group (A) reported to have symptoms of distortion post-operatively in comparison to the 10 patients from Group (B) who also reported to have post-operative symptoms of distortion.

In the patients with evidence of retinal displacement on FAF, symptoms of distortion were present in 85.7% (12/14) eyes, on the other hand in those with symptoms of distortion there was evidence of retinal displacement in 63.1% (12/19) eyes this may indicate that retinal displacement is not the only cause of the symptoms of distortion (Table 3).

Table (3): Correlation between Retinal displacement on FAF and Symptoms of distortion in the studied groups.

<table>
<thead>
<tr>
<th>Visual Disturbances</th>
<th>No displacement</th>
<th>Displacement</th>
<th>( p )-value</th>
<th>Sig.</th>
</tr>
</thead>
<tbody>
<tr>
<td>No</td>
<td>9</td>
<td>56.3</td>
<td>2</td>
<td>14.3</td>
</tr>
<tr>
<td>Metamorphopsia</td>
<td>6</td>
<td>37.5</td>
<td>7</td>
<td>50.0</td>
</tr>
<tr>
<td>Metamorphopsia+</td>
<td>1</td>
<td>6.3</td>
<td>4</td>
<td>28.6</td>
</tr>
<tr>
<td>Micropsia</td>
<td>0</td>
<td>0.0</td>
<td>1</td>
<td>7.1</td>
</tr>
</tbody>
</table>

Discussion

PFCLs have become popular in retinal detachment surgery because of their physical properties (optical transparency, low viscosity, immiscibility with water) [9]. They allow the retina to reattach without the need for retinotomies and intraoperative stabilization of the retina in more complex situations, such as trauma with the presence of PVR.9,10 The major side effects of PFCL arise from their abnormal intraocular fate, in the vitreous cavity, subretinally or in the anterior chamber [9,10,11]. Benefits of the presented PFCL-free technique include, in addition to avoiding possible side effects of the lagging PFCL, better visibility of the anterior retina in the air-filled eye, which may additionally facilitate retinal reattachment [12].

In our study the primary anatomical success was achieved in all 15 eyes (100%) of Group (A) patients where PFCL wasn’t used, and in 14 of the
15 eyes of Group (B) patients (93.3%) where PFCL was used, which wasn’t statistically significant (p-value = 0.309), with both groups (100%) successfully achieving the final anatomical success.

Comparable results between both approaches have been documented in literature as regards the anatomical outcomes of PPV in cases having primary RRD with primary anatomical success rates ranging from 90.0% to 98.0% in cases with routine PFCL use [13,14,15]. It was 95.7% in the study published by Schneider et al., in which PFCL wasn’t used, with all of these mentioned studies achieving a near 100% final anatomical success.

(FAF) has been reported to be an accurate mean of assessing retinal displacement, and this was the technique of choice in several studies [16]. In the study published by Lee et al., they hypothesized that the use of PFCLs might theoretically reduce the incidence of postoperative retinal displacement evident on FAF by reducing the residual postoperative SRF based on the fact that most studies attribute the displacement to the effect of the residual SRF postoperatively acting as a downward force on the retina hence causing downward displacement of the retina following retinal detachment surgery [6,7,8].

In our study FAF showed retinal displacement (demonstrated by the hyperfluorescent lines running parallel to the retinal blood vessels) in 14 of the 30 eyes (46.6%). These results are consistent with the recent study conducted by Dell’Omo et al., which documented the displacement in 41.2% of cases [7]. This was slightly less than the older studies by Shiragami et al.; Lee et al.; Cobos et al. and Codenotti et al., which were 62.8%, 72%, 60%, 71.4% respectively [6,8,16,17].

Of the 14 cases that showed retinal displacement on FAF, PFCL was used in 6 eyes (42.8%) and not used in 8 eyes (57.1 %) hence indicating that its use has no statistical significance in the incidence of retinal displacement. This is consistent with the studies performed by Shiragami et al.; Dell’Omo et al. and Codenotti et al., who also found no statistically significant correlation between the use of PFCLs and reducing the incidence of retinal displacement [6,7,17].

Among the 14 eyes that showed retinal displacement on FAF the direction of shift was downwards in all the eyes showing displacement This is in agreement with the studies by Shiragami et al.; Lee et al.; Codenotti et al. and Pandya et al., which all reported that retinal displacement occurs in downwards direction with gas tamponade [6,8,17,18]. In contrast to the study by Dell’Omo et al., which stated that displacement with gas tamponade can occur both upwards or downwards but also stating that downward displacement is much more common [7].

Another important aspect that was to be taken in consideration was the functional outcome following retinal detachment surgery, no statistically significant difference in the final mean BCVA was found between the two groups. These results are in agreement with the results published by Schneider et al., who also reported comparable final BCVA results (without the use of PFCL) to studies where PFCL was used [19].

The mean BCVA on the first postoperative follow-up following the absorption of gas did not differ in eyes with or without displacement evident on FAF. This observation is in line with the studies published by Shiragami et al. and Dell’Omo et al., who both found no correlation between the incidence of retinal displacement and the final BCVA [6,7].

In addition to the final BCVA, symptoms of distortion especially metamorphopsia was detected in 12 eyes of the 14 eyes showing retinal displacement on FAF (85.7%), 7 of the 8 eyes with retinal displacement in Group (A) (87.5%) (with the displacement not being macular in the only patient not complaining of metamorphopsia) and in 5 of the 6 eyes (4 of whom with displacement involving the macula) with retinal displacement in Group (B) (83.3%). This is in accordance with the study published by Lee et al., which reported metamorphopsia in 83% of patients with macular shift (19/23) [8]. However, of the 19 cases complaining of metamorphopsia only 12 had evidence of displacement on FAF (63.1%) which is less than the same study by Lee et al. in which reported that in patients complaining of symptoms of distortion there was evidence of shift in 86% (19/22) [8]. On the other hand, our results are higher than the results published by Dell’Omo et al., who reported retinal displacement in only 18.1% (4/22) of the patients complaining post-operatively of metamorphopsia, hence we recommend that further research must be put in the post-operative follow-up of patients [7].

Our study was limited by the small sample size, although it does hold the advantage over other studies being randomized in terms of PFCLs use. Also, the group tamponaded with gas without adjuvant PFCLs use and the group tamponaded with gas together with adjuvant PFCL use differed
with respect to some baseline features like number of detached quadrants, foveal involvement and concomitant involvement of the upper and lower quadrants. Further researches including larger sample size are needed to evaluate causes of retinal displacement and its implications.

**Conclusion:**

In conclusion, retinal displacement is common following RRD repair and the use of PFCLs does not seem to affect the incidence of displacement. We found no correlation between the incidence of retinal displacement and the final best corrected visual acuity, but usually it is associated with symptoms of distortion.

**Disclosure:**

The author reports no conflicts of interest in this work.

**References**

تأخير استخدام سائل البيروفورميكيون على إزاحة الشبكية
بعد عملية استئصال الجسم الزجاجي لعلاج انفصال الشبكية الأولي:
دراسة مقارنة

الغرض: مقارنة حالات إزاحة الشبكية بعد استئصال الجسم الزجاجي لإصلاح انفصال الشبكية الأولي مع ودون استخدام سائل البيروفورميكيون باستخدام أشعة ضوئية قاع العين وربطها بأفضل حالة بصر وأعراض التشبه في الصورة بعد العملية الجراحية.

الأهداف: دراسة مقارنة، مترقبة، تتخلية، عشوائية. تم تقسيم ثلاثين عينا للثلاثين مريضاً يعانون من انفصال الشبكية الأولي إلى مجموعتين.

المجموعة A: 15 عين خضعت لعملية استئصال الجسم الزجاجي بدون استخدام سائل البيروفورميكيون والمجموعة B: 15 عين خضعوا لعملية استئصال الجسم الزجاجي باستخدام سائل البيروفورميكيون.

النتائج: إزاحة الشبكية كانت واضحة على أشعة ضوئية قاع العين بنسبة 37.5% في حالة مجموعة A و 40% في حالة مجموعة B على التوالي (p=0.0464). استخدم سائل البيروفورميكيون ليس له تأثير إحصائي واضح على نسبة حول إزاحة في الشبكية. لا يوجد فرق إحصائي واضح في متوسط حد الإفسار النهائي بين المجموعتين. التشبه ما بعد العملية تم إكتشافه في 88.5% من العين بواسطة أشعة ضوئية قاع العين.

الخلاصة: إزاحة الشبكية مشكلة شائعة بعد عمليات إزالة الجسم الزجاجي لإصلاح الانفصال الشبكي الأولي واستخدام سائل البيروفورميكيون.

لا يبدو أنه يؤثر على هذه النسبة.