# **ORIGINAL ARTICLE**

# Panretinal photocoagulation laser in diabetic retinopathy patients

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### ABSTRACT

Background: It was reported that corneal endothelial cell depletion is a complication of the retinal laser. In this study, the effect of panretinal laser on corneal endothelium was investigated. Method: A group consisting of 47 eyes of 47 Diabetic patients who need a panretinal photocoagulation laser was compared with another group of 47 diabetic patients who did not require a laser. The patient's age was chosen in the range of 50-70 and the tools included Ellex double frequency LASER, Specular microscopy, Pachymetry, and Condense Lens 90. Time course of coefficient of variation, endothelial cell density, hexagonal cells and central corneal thickness were evaluated. ANOVA, Dennett's, and Mann-Whitney test was used to compare the data. P-value of <0.05 was considered statistically significant. Results: We found no significant differences in age, gender between the two groups. There was no significant differences in hexagonal cells, coefficient of variation, endothelial cell density and central corneal thickness between the two groups at any time points. **Conclusion**: This study suggests that the "panretinal photocoagulation laser" conserves corneal endothelial cells of subsequent damage in diabetic retinopathy patients

Keywords: photocoagulation laser, corneal, endothelial cell, argon laser, double frequency laser.

# INTRODUCTION

Diabetes can have diverse impacts on several sections of the optic system. One can be visual disorders due to vascular and metabolic defects that can change the retina and the visual pathway. The formation of new vessels in the retina is the progress of retinopathy and causes proliferative diabetic retinopathy, so neovascularization happens in response to retinal ischemia, and new vessels grow on optical discs or elsewhere. Proliferative diabetic retinopathy (PDR) is the main cause of blindness in working-age adults and has affected many communities worldwide (1, 2).

Human corneal endothelial cells (HCECs) do not regenerate in vivo to a considerable point, holding in the G1-phase of the cell cycle(3). Panretinal photocoagulation laser (PRP) has been the standard remedy for patients with PDR since the 1970 (4). Although PRP reduces the risk of critical vision loss, its laser power can eradicate all retina layers such as ganglion cells and the retinal nerves, and hence cause visual field deficits. In such states, visual field testing can be lesser desirable to estimate the damage.(5).

Speculator microscopy is an optical reflection microscope that reflects Slit light to the cornea and allows it to be viewed. The device is designed so that it does not allow the observation of non-specular light waves, so the image that is seen is exclusively related to the light waves that are reflected speculatively (6). Some of the characteristics of abnormal corneal endothelial cells include the following: cell count < 1500 cell/mm<sup>2</sup>, severe polymegatism, severe pleomorphism, presence of guttata, presence of any abnormal formation such as coalescence, an abnormal defect in a cell, wide area of severe edema, presence of

bacteria or inflammatory cells and presence of ghost cells in the stroma. What is more common in people with diabetes is the increase in polymegatism, pleomorphism, and decreased cell hexagonally rate (7).

The laser beam needs to pass through anterior tissues such as the cornea, lens, and vitreous to reach the retina. The possibility of absorbing laser beam energy as it passes through the corneal tissue and damages the endothelial cells is predicted by a decrease in the number of cells, a decrease in the cells' hexagonally, or an increase in the thickness of the central part of the cornea(8-10). Due to the importance of corneal endothelium cells in maintaining corneal transparency, the laser beam's effect on corneal endothelium cells should be evaluated. Therefore, examining and comparing these cells before and after laser treatment can help clarify and provide other alternate methods for diabetic retinopathy if needed. This study aimed to compare variations in corneal endothelial cell indices with speculator microscopy examination before and after panretinal laser in diabetic retinopathy patients.

### **MATERIALS & METHOD**

Retinal photocoagulation typically involves using a laser pulse lasting 10-200ms, causing temporary hyperthermia several tens of degrees above normal body temperature. Currently, the most common laser used for photocoagulation is Nd: YAG=532 nm and Yellow semi conductive LASER= 577 nm. The tools and equipment used in this research include Ellex double frequency LASER (Germany), Specular microscopy (SP 3000P), Pachymetry (Topcon. Japan), Condense Lens 90(volk.US). This investigation was conducted as an intervention (Quasi-experiments). A group consisting of 50 eyes of 50 Diabetic patients who require a panretinal photocoagulation laser was compared with another group of 50 diabetic patients who did not need a laser. The patient's age was selected in the range of 50-70 years, and the study was explained to all members, and written consent was obtained from them. The subjects had the right to leave the examination whenever they wanted. Age, gender, history of eye disease, family history, visual acuity with and without refraction using the Snellen chart, corneal examination, anterior chamber, iris lens and anterior vitreous with slit biomicroscopy lamp, retinal examination with 95 mark volk lenses, microscopic specular and central corneal thickness (CCT), was conducted for all participants. In all four stages of the patient's examination, the physician was the same.

It was reassured that microscopic speculation is a noncontact method and would not cause any harm. Patients who have diabetic retinopathy and needed Panretinal laser photocoagulation were candidates for this treatment Patients with a history or active status of intraocular inflammation, patients with a history of any previous ocular surgery, patients with known Corneal endothelial Fuchs dystrophy, patients with known Iridocorneal endothelial syndrome, known Corneal opacity patients, patients with any glaucoma, Patients with reduced anterior chamber depth, who have used contact lenses in the past three months, who undergo intraocular surgery during these four months, who have previously undergone intraocular laser of any kind for any reason, Patients who have previously undergone photorefractive surgery, patients who have had an episode of corneal edema for any reason in their previous ophthalmology record, patients who have received an intraocular injection (in the anterior chamber or in vitro) of antibiotics, corton, ranibizumab or bevacizumab, were excluded from the study. In the current study, 3 patients were excluded from the study due to non-referral in one month after laser treatment, and for this reason, three patients were excluded from the control group.

#### Statistics

All statistical analyses were performed using SPSS 22. One-way analysis of variance (ANOVA) was performed for the analysis of the time course of changes with the Dunnett's test for multiple comparisons. The Mann-Whitney test was used to compare the data between the two groups. Unless otherwise indicated, the results are expressed as mean ±SD, p- value of p<0.05 was considered statistically significant.

### RESULTS

Demographics of the study population are listed in Table 1. We found no significant differences in age (p=0.43), gender (p=0.63) between the two groups. Best corrected visual acuity at baseline is also shown in Table 1 and as you can see, in terms of the visions, the two groups are significantly different. In the control group, all patients had mild to moderate diabetic retinopathy who did not receive laser treatment. In the treatment group, 33 patients, 70%, had Severe NPDR (*non-proliferative diabetic retinopathy*), and 14 patients, 30%, had early PDR retinopathy.

| Demographic                                  |             | Control Group | Treatment group | P value |
|--|-------------|---------------|-----------------|---------|
| Age (mean ±SD years)                         |             | 60.6 ± 4.97   | 59.38± 3.57     | 0.43    |
| Gender (% female)                            |             | 42.55         | 44.68           | 0.63    |
| Best corrected visual acuity BCVA <b>(%)</b> | 20/600      | 14.89         | 40.42           | 0.02    |
|  | 20/400      | 29.78         | 34.04           |         |
|  | 20/300      | 25.53         | 23.4            | 0.02    |
|  | 20/250      | 29.78         | 2.12            |         |
|  | Severe NPDR |               | 70              |         |
| Diabetic Retinopathy (%)                     | Early PDR   |               | 30              |         |
|  | Mild NPDR   | 100           |                 |         |
| Laser Spot                                   |             |               | 1629.89         |         |
| Laser Power                                  |             |               | 199.78          |         |

Table 1 Demographics Features of the study population

Time course of CV (coefficient of variation), ECD (endothelial cell density), HEX (hexagonal cells), and CCT (central corneal thickness) is shown in Table 2. The coefficient of variation was not statistically significant in control and treatment group p=0.7. There was no significant differences in hexagonal cells between the two groups at any time point's p=0.8. We found no significant

differences in central corneal thickness (p= 0.4) and endothelial cell density (p=0.2) between the two groups. We also detect no significant changes in coefficient of variation, endothelial cell density, hexagonal cells, and central corneal thickness 6 month after panretinal photocoagulation laser.

| Exam                       | Groups    | Before PRP | After 1 month | After 3 months | After 6 months | P value |  |
|----------------------------|-----------|------------|---------------|----------------|----------------|---------|--|
|                            | Control   | 40.3±3.5   | 40.2±3        | 40.1±3         | 40.3±3.2       |         |  |
| HEX (%)                    |           |            |               |                |                | 0.8     |  |
|                            | Treatment | 36.9±3.1   | 36.4±3        | 36.2±2.6       | 36.3±2.9       |         |  |
|                            |           |            |               |                |                |         |  |
| CV (%)                     | Control   | 37.6±1.9   | 37.7±1.6      | 37.6±1.5       | 37.8±2.1       | 0.7     |  |
|                            | Treatment | 37.5±2.9   | 37.9±2.9      | 37.7±2.3       | 37.6±3.0       |         |  |
| ССТ                        | Control   | 556.1±13.4 | 556.3±12.8    | 555.4±10.5     | 553.2±12.7     | 0.4     |  |
|                            | Treatment | 551.6±15.8 | 551.5±15.8    | 551.7±15.9     | 552.1±14.9     |         |  |
| ECD <sub>(cells/mm³)</sub> | Control   | 2637±186   | 2635±185      | 2624±184       | 2620±184       | 0.2     |  |
|                            | Treatment | 2771±176   | 2770±168      | 2779±161       | 2774±173       |         |  |

Table 2. ECD, CV and percentage of HEX before and after times in control and treatment group

CV, coefficient of variation; ECD, endothelial cell density; HEX, hexagonal cells; CCT, central corneal thickness.

## DISCUSSION

The current study evaluates if there is a correlation between panretinal photocoagulation laser and changes in the condition of corneal endothelial cells; it is tried to clarify whether changes in the density of endothelial cells occur due to panretinal photocoagulation laser: and in 6 months how are these changes? The number and hexagonal value of endothelial cells, the average Coefficient of Variation, and the mean cornea central thickness in laser group, during six months had no significant change.

Associated complications of panretinal photocoagulation were detected generally in the initial years of it. The frequent complications stated were choroidal detachment, exudative retinal detachment, anterior chamber narrowing, transient glaucoma, and ocular edema (11). Visual variations reported outright after panretinal photocoagulation include transient photophobia, impaired accommodation, and myopia. Reduced visual field and night vision deficiencies are typical subject magnitude (12, 13).

In the present study, our results demonstrated that the mean CCT loss was 0.36% in control group and increased 0.1% in treatment after 6 months indicating that panretinal photocoagulation laser provided a long-term safety in the term of CCT. Further the mean ECD loss was 0.64% in control and increased 0.1% in treatment after 6 months, indicating that panretinal photocoagulation laser provided almost long-term safety in the term of ECD and CCT. Previous investigations on the change in ECD after Lasers were summarized in Table 3 (10, 14-16).

| Author(Year)          | Eyes | Laser type                              | Follow-<br>up | Mean ECD<br>loss (%) | Mean CCT<br>loss (%) | CV (%)        | HEX<br>(%)    |
|-----------------------|------|---|---------------|----------------------|----------------------|---------------|---------------|
| Nurgul Ornek<br>2018  | 18   | Selective laser<br>trabeculoplasty      | 1 Month       | 1.3                  | 0.22                 | 40.1-<br>37.9 | 54-57         |
| Kanagaratnam<br>2020  | 244  | Selective laser<br>trabeculoplasty      | 1 Month       | 0.56                 | + 0.18               | 38.5-<br>38.8 | 50.3-<br>51.2 |
| Kanagaratnam<br>2020  | 132  | Retinal<br>photocoagulation<br>laser    | 1 Month       | 0.39                 | +0.36                | 38.1-<br>39.4 | 50.6-<br>50.4 |
| Pardos 1981           | 40   | Argon laser photocoagulation            | 1.5<br>Month  | 2.7                  |                      |               |               |
| Chandler 2003         | 60   | Trans corneal iridal photocoagulation   | 3 weeks       | 1.2                  | + 4.5                |               |               |
| Present Research 2020 | 47   | Panretinal<br>photocoagulation<br>laser | 6 Month       | + 0.1                | + 0.1                | 37.5-<br>37.6 | 36.9-<br>36.3 |

Table 3. Previous studies reporting central ECD and CCT loss, CV and percentage of HEX after laser.

CV, coefficient of variation; ECD, endothelial cell density; HEX, hexagonal cells; CCT, central corneal thickness.

The cornea's cellular stress levels are often indicated by the cell surface changes, such as the cells abnormally and the divergence from their hexagonal form (17, 18). We found no significant variations in morphologic parameters such as HEX or CV neither in control eyes nor in treatment eyes, suggesting that panretinal photocoagulation laser does not induce a notable change in polymegatism polymorphism even six months post-laser. Moreover, the

CV in our study was slightly higher, and the HEX was slightly lower than before the treatment. However, the information about CV and HEX should be described and interpreted by discretion and couldn't be accurately compared among methods employed in the current and previous studies. Pardos et al. 1981 stated that the mean ECD loss was 2.7% at six weeks after Argon laser photocoagulation (16). On the other similar to the present study, Kanagaratnam et al. 2020 revealed mean CCT increased 0.36 month after after one Retinal photocoagulation laser (15).

Previously, it has been reported that argon laser photocoagulation decreases endothelial cell density (16, 19), whereas, in another study, retinal photocoagulation protected against endothelial cell damage (15).Kanagaratnam .et al in a cohort study, revealed that Retinal Photocoagulation seems to have a more limited impact on the corneal endothelium, with the only alteration of note an accretion in polymegatism among pre retinal photocoagulation and post retinal photocoagulation (15).

#### CONCLUSION

This study suggests that the "panretinal photocoagulation laser" conserves corneal endothelial cells of subsequent damage in diabetic retinopathy patients. It seems that performing argon panretinal laser, which is the most common type of retinal laser performed in diabetic patients, does not damage the number and hexagonally of corneal endothelial cells and does not change the CCT. This study was six months that cannot generalize this result for a long time, and probably more studies are needed.

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