

Black patients sustain vision loss while White and South Asian patients gain vision following delamination or segmentation surgery for tractional complications associated with proliferative diabetic retinopathy

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Abstract

Purpose This retrospective comparative case series aims to determine whether patient ethnicity (White versus South Asian versus Black) is related to the outcome of surgical treatment for traction complications of severe proliferative diabetic retinopathy (PDR).

Setting Moorfields Eye Hospital London, UK.

Methods All patients who underwent vitrectomy with, delamination and/or segmentation for PDR over a 5-year period (2009–2014) were reviewed retrospectively. Patients were divided into White, South Asian or Black groups, and their age, gender, HbA1C and type of diabetes were recorded. A total of 484 patients (253 White, 117 South Asian, 114 Black) were included. Twenty-one patients were excluded due to inadequate documentation.

Outcomes LogMAR Visual acuity (converted from Snellen) (VA), was recorded pre-operatively and ~6 months post surgery (range 5–8 months). Surgical outcome was classified according to the type and duration of tamponade required post-operatively.

Results Pre-operative VA and HbA1C values were similar across all three ethnic groups ($P=0.64$ and 0.569 , respectively). Change in VA (mean \pm SD) was 0.41 ± 0.78 , 0.14 ± 0.76 and -0.26 ± 0.57 in White, South Asian and

Black patient groups respectively ($P<0.001$). Multiple regression analysis showed that post-op VA was significantly related to race and pre-op VA only (both $P<0.001$). The Black patient group were more likely to require silicone oil tamponade ($P<0.001$) and long-term retention of silicone oil ($P<0.001$) than the White and South Asian patient groups.

Conclusions This study demonstrates that Black patients on average lose vision following delamination surgery for traction complications of PDR while White and South Asian patients gain vision. The same group is also at higher risk of retaining silicone more than 6 months after surgery. This difference remains even when corrected for glycaemic control. The higher risk of visual loss and long-term retention of silicone oil in black patients requires further investigation. If these results are confirmed, surgeons should consider their patients' ethnicity before proceeding with surgical treatment of diabetic tractional detachment.

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Introduction

Vision loss is one of the most feared complications amongst the diabetic population.

Although the risk of retinopathy can be reduced with good glycaemic and blood pressure (BP) control^{1–4} and, ocular complications may be treatable with laser,⁵ surgery⁶ and pharmacologically,⁷ visual loss from diabetic retinopathy (DR) remains prevalent. Severe proliferative diabetic retinopathy (PDR) poses the most serious threat to sight. Complications of PDR such as intraocular haemorrhage, fibrous proliferation, traction retinal detachment or traction retinoschisis^{8,9} can lead to profound vision loss.¹⁰ The prevalence of PDR is estimated at between 13 and 50% after 15–20 years of diabetes duration in patients needing insulin^{11,12} with a risk of becoming legally blind after 5–10 years if left untreated.¹³

Vitreoretinal surgery, including vitrectomy, with or without delamination and segmentation, is recognised as the principal treatment for late complications of PDR such as intraocular haemorrhage, traction retinal detachment (involving or threatening the macula) and, uncontrolled progressive fibro-vascular proliferation.^{6,14} The main aims of surgery are to clear the media opacities (usually blood) and divide or remove as many as possible of the adhesions causing retinal traction. Although several studies have evaluated the outcomes, complications and benefits of surgery for the complications of PDR,^{6,14–16} differences in outcome relative to ethnic origin have not been investigated.¹⁷ This study sets out to explore the impact of the ethnicity of patients—White versus Black versus South Asian, on the treatment outcomes of the subgroup of patients who undergo vitrectomy with delamination and/or segmentation surgery for the traction complications of severe PDR.

Methods

Patient selection and inclusion

This work describes a retrospective (2009–2014) comparative case series from a tertiary referral centre in the United Kingdom. A search of all surgical bookings on the electronic patient record database using the keywords ‘diabetes’ and/or ‘vitrectomy’ and/or ‘delamination’ and/or ‘segmentation’ was performed to identify all of the diabetic patients who underwent vitrectomy surgery (including ‘with delamination and segmentation’) during this period. Manual checking of each patient’s case note was performed to confirm the actual diagnoses and surgical procedures performed. Where delamination and/or segmentation were specifically recorded as part of the operation, the subject was included in our study. Cases of vitreous haemorrhage that only required vitrectomy and laser were excluded. The study received prospective internal review board approval (Research and

Development, Moorfields Eye Hospital NHS Foundation Trust. Reference number: ROAD 14/038).

Demographics

Ethnicity was classified according to the hospital record of self-reported ethnicity at registration for each patient. Patients were placed into three main groups: White, Black, or South Asian. The subgroups pooled into each main group were White (White British, White Irish, White any other background), South Asian (Indian, Pakistani, Afghan, Bangladeshi, Nepalese, Bhutanese, Maldivian) and Black (African, Caribbean, Black any other background). Cases where the ethnicity was not accurately recorded or not declared were excluded from the study. Mixed race or Filipino/Oriental racial groups were also excluded from the study but were minimal in number.

Other details collected were age, gender and type of diabetes. All Black patients were tested for the presence of Haemoglobin S and those that were positive were excluded from the study as their retinal pathology was treated as non-diabetic (6 patients were excluded for this reason).

Functional outcome

Visual acuity (VA), using the patients’ habitual best correction, was measured in Snellen and converted to logMAR. The VA was recorded pre-operatively and at the follow-up visit closest to 6 months post-operatively in both eyes (range 5–8 months), this will be referred to as the 6 months visit. The sub-form acuity measures namely ‘counting fingers’ (CF), ‘hand motion’ (HM), ‘light perception’ (LP) and ‘no light perception’ (NLP) received the following logMAR values: CF = 1.6, HM = 2.0, LP = 2.5, NLP = 3.0.¹⁸

Surgical outcome

The surgical outcome for each patient was recorded according to the type and duration of retinal tamponade required post-operatively. These were classified into three groups being (1) gas only or no tamponade (2) Silicone Oil (SO) tamponade used but removed <6 months post-operatively, and (3) SO used and retained ≥6 months.

Glycaemic control

HbA1c was recorded for each patient within 1 month of surgery.

Bilateral disease

In patients with bilateral traction pathology, where both eyes required vitrectomy and delamination surgery one eye was randomly chosen (<http://www.randomizer.org/>) for inclusion in the study in order to maintain sample independence.

Statistical analysis

All statistical analyses were performed using SPSS 23 for Mac (SPSS, Chicago, IL, USA). Descriptive statistics were presented as mean \pm SD. Histograms were plotted for each of the three variables—age, VA, and HbA1c levels to assess the normality of their sample distributions. After consultation with an independent statistician and, as normality of the sample distribution could not be established for the above three variables, non-parametric tests were chosen for the analysis. The level of statistical significance is set as $P < 0.05$. A multiple regression analysis was performed to predict the post-operative VA from age, pre-operative VA, HbA1c level, diabetes type (1 or 2), and ethnicity.

Results

The electronic search identified 1376 diabetic patients who underwent vitrectomy over the study period. Of these, 502 patients (668 eyes) were identified as having delamination or segmentation surgery as described above. Eighteen patients (27 eyes, 3.5%) were excluded because of inadequate documentation in the electronic database and case notes, or because they did not fit clearly into the three main racial groups we were analysing. Data from only one eye of the remaining 484 patients were analysed.

Demographics

Of the 484 patients, 253 were White, 117 were South Asian, and 114 were Black. Overall, 157 (32%) patients had bilateral surgery. As a proportion of the total number of patients within each racial group the number of patients having bilateral surgeries were 93/253 (37%) White, 33/117 (28%) South Asian and 31/114 (27%) Black.

The mean age (years) \pm standard deviation (SD) for all patients was 51.9 ± 14.4 . This was 48.9 ± 15.6 , 54.5 ± 11.7 , and 56.1 ± 12.7 in the White, South Asian and Black groups respectively. The White group was significantly younger than South Asian and Black groups ($P = 0.001$ and $P < 0.001$ respectively, Mann–Whitney U -test). There were 213 females (44.0%) and 271 males (56.0%) in total. Gender was not associated with racial group (% male by

racial group was: 54.9% for White, 57.3% for South Asian, and 57.0% for Black; P -value from χ^2 -test = 0.887).

Type 1 diabetes was present in 159 (33%) patients, while 325 patients (67%) had type 2 diabetes. Type 1 diabetes was more common in the White patient group (% type 1 by racial group was: 45.9% for White, 17.1% for South Asian, 20.2% for Black; P -value from χ^2 -test < 0.001). Within the type 2 group, the mean age (years) \pm SD was: 60.2 ± 10.0 , 57.7 ± 9.0 , and 60.4 ± 9.0 for White, South Asian, and Black groups, respectively. There were no statistically significant differences in the age distribution of the type 2 diabetic patients across the three ethnic groups ($P = 0.89$, Kruskal–Wallis test).

As type 1 diabetes typically present in much younger patients, giving rise to the younger mean age of in the White group of patients, we performed subgroup comparisons of only the type 2 diabetic patients across the three ethnic groups.

Glycaemic control

HbA1c values were available in 91% of the patients. The mean and standard deviation for each racial group are listed in Table 1. There were no significant differences in the distribution of HbA1c across the three groups ($P = 0.569$, Kruskal–Wallis test). Pearson's correlation analysis indicated that there was no significant association between peri-operative HbA1c level and the final VA at 6 months post-operatively (Pearson's correlation = 0.097).

Functional outcomes

Visual acuity was available for all patients pre-operatively and at 6 months post-operatively (as defined) in both eyes. The median pre-operative VA with interquartile range (logMAR), median post-operative VA with interquartile range (logMAR) and the mean change in VA \pm SD after surgery for patients in each ethnic group is recorded in Table 1. The same parameters for type 2 diabetic patients are also recorded for each ethnic group in Table 1.

The percentages of patients from each racial group with sub-form vision (including combined CF, HM, PL, and NPL), both pre and post-operative are documented in Table 1.

Multiple regression analysis

A multiple regression was performed to predict the post-op BCVA from age, pre-operative BCVA, per-operative HbA1c level, Diabetes type (1 or 2), and ethnicity. Collectively these variables significantly predicted post-op BCVA ($F(5, 436) = 39.1$, $P < 0.001$, $R^2 = 0.31$). Of these,

Table 1 Median with interquartile range (IQR) of pre-operative and post-operative Visual Acuity (VA)^a and percentage of patients requiring silicone oil (SO) tamponade for the three racial groups

Parameter	White	South Asian	Black	P-value	White Vs South Asian	White Vs Black	South Asian Vs Black
All pre-op VA median (IQR)	1 (0.18–2.5)	1 (0.18–2.5)	1 (0.18–2.5)	P = 0.643 ^b	—	—	—
All post-op VA median (IQR)	0.6 (–0.8–3)	0.78 (0–3)	1 (0–3)	P < 0.001 ^b	P < 0.001 ^c	P < 0.001 ^c	P = 0.001 ^c
Change in VA (mean ± SD)	0.41 ± 0.78	0.14 ± 0.76	–0.26 ± 0.57	P < 0.001 for all 3 groups ^d	—	—	—
T2 pre-op VA ^b median (IQR)	1 (0.18–2.5)	1 (0.18–2.5)	1 (0.3–2.5)	P = 0.554 ^b	—	—	—
T2 post-op VA ^b median (IQR)	0.6 (0–3)	0.78 (–0.08–1.93)	1.3 (0–3)	P < 0.001 ^b	P = 0.016 ^c	P < 0.001 ^c	P < 0.001 ^c
T2 change VA ^c (mean ± SD)	0.41 ± 0.75	0.11 ± 0.75	–0.31 ± 0.57	P < 0.001 for all 3 groups ^d	—	—	—
HbA1C (mmol/mol) (mean ± SD)	103.1 ± 34.1	107.0 ± 37.1	107.9 ± 37.7	P = 0.569 across the 3 groups ^b	—	—	—
SO used at surgery (%)	16	17	43	P < 0.001 ^e	P = 0.94 ^e	P < 0.001 ^e	P < 0.001 ^e
SO retained ≥ 6 months (%)	7	9	30	P < 0.001 ^e	P = 0.94 ^e	P < 0.001 ^e	P < 0.001 ^e
Pre-op BCVA (CF, HM, PL, NPL) (%) ^f	37	37	35	—	—	—	—
Post-op BCVA (CF, HM, PL, NPL) (%) ^f	19	32	46	—	—	—	—

Abbreviations: SO, silicone oil; T2, type 2 diabetes; ‘—’, no statistical analysis carried out. ^aVisual acuity with habitual correction recorded as Snellen and converted to logMAR. ^bKruskal–Wallis test. ^cMann–Whitney U-test. ^dPaired T-test. ^eχ²-test performed to compare association between SO tamponade requirement across all 3 ethnic groups. ^fAs a percentage of the total of the specific racial group.

only two variables showed statistically significant predictive value for the outcome of post-op BCVA. They are pre-op VA and ethnicity ($P < 0.001$ for both).

Outcomes of patients without long-term silicone oil tamponade

Subgroup analysis looking at the VA outcome in patients who did not require long-term silicone oil tamponade showed that the median pre and post-operative VA (InterQuartile Range) was 1 (0.18–2.50) and 0.48 (–0.8 to 3) for the White group ($n = 235$, 93% of White patients), 1 (0.18–2.50) and 0.6 (0–3) for the South Asian group ($n = 107$, 91% of South Asian patients), and 0.78 (0.18–1.83) and 1 (0–3) for Black group ($n = 80$, 70% of Black patients). No further statistical analysis was performed as this subgroup represented an inherently biased cohort selected for better outcome. However, the trend indicates that more White and South Asian patients are likely to be tamponade free in the long term, with better VA than Black patients.

Surgical outcome

The proportions of patients requiring SO tamponade at the end of surgery and those with SO retained after the 6 month’s visit are recorded in Table 1.

Discussion

Vision loss secondary to DR remains a major public health issue for most countries. Ethnic differences in the incidence, prevalence, severity and response to treatment in DR, as well as other ophthalmic conditions such as glaucoma and cataract have been demonstrated.^{19–23} In this study, we have shown that Black patients on average lose vision while White and South Asian patients gain vision following surgical delamination or segmentation for tractional retinal detachment or tractional schisis secondary to PDR. We have also observed that ethnicity affects the anatomical outcomes following surgery in that silicone oil is used in surgery and retained after 6 months more frequently in Black patients, than South Asian or White patients.

In terms of PDR we have observed within our study, that the proportion of White, South Asian and Black patients were 52.3, 24.2, and 23.5% while the respective proportion of these groups in the London population are 59.7, 16.9, and 11.2%.²⁴ This indicates that a higher proportion of South Asian and Black patients had severe PDR requiring surgery, which compounds the finding that the Black patient group has a much poorer visual and surgical outcomes after surgery. The proportions may

also reflect the higher representation of type 2 diabetes in these two ethnic groups.²⁵

The mean age of the patients in our series was lower in the White group, though this likely reflects the higher prevalence of type 1 diabetes in White patients compared to South Asian (up to 3 times) and Black (up to 6 times) patients.²⁶ When only type 2 diabetic patients were analysed, the age distribution was comparable for the three groups, but the Black group still had a significantly worse visual outcome. Importantly, a multiple regression analysis showed that mean age, type of diabetes and HbA1c alone across the three ethnic groups were insufficient to explain the differences in their visual outcome. It also confirmed that race and pre-operative VA was significantly predictive of final VA.

HbA1c level was not predictive of outcome independently of race and this is consistent with data from diabetic screening. Retinal screening has demonstrated a reduction in sight threatening DR in London,²⁷ with non-attendance to screening not influenced by ethnicity despite the disparity in socio-economic status.²⁸ In our cohort of patients there was also no significant difference in HbA1c levels between the different ethnic groups. The poorer visual outcomes observed in the South Asian and Black patients, therefore, could not be explained by differences in the systemic control of diabetes alone.

In many surgical treatments, the operative outcome is partly determined by the technical difficulty of the procedure. For diabetic delamination surgery, we have used the type and duration of tamponade required as a *defacto* indicator of the degree of difficulty and anatomical outcome of surgery. Silicone oil tamponade is mainly used in technically difficult cases in which retinal re-attachment is threatened by residual retinal traction.²⁹ In a previous study, Yorston *et al*¹⁷ showed that the use of long-acting tamponades, that is, silicone oil, was more likely in eyes with extensive areas of retinal adhesions. They noted this surgical feature more frequently in Black patients than in patients of other ethnicities. In our study, Black patients showed a markedly higher rate of SO tamponade and retention of SO long term (≥ 6 months) than in the other groups. It is possible that Black patients have stronger vitreo-retinal adhesions thereby rendering surgical delamination of their eyes technically more difficult. In keeping with the worse surgical outcomes, the post-operative visual acuity was also significantly worse in Black patients relative to the other two groups. In a subgroup analysis of visual acuity outcomes in patients who did not have silicone oil at 6 months there was a trend for black and South Asian patients to do worse. We did not examine this statistically as there was a sampling

bias in the choice of good outcome but it suggests black patients may do worse after surgery independent of silicone.

In our study, a potential confounding factor in terms of visual outcome is the extent and timing of previous retinal laser treatment. Unfortunately, we could not get reliable information about timing and extent of laser in this cohort of patients to analyse its effect. Furthermore, the extent of fibrovascular tissue could also be more extensive in black patients and therefore influence the visual and anatomical outcomes of surgery. Unfortunately, the source data in this retrospective study did not have the quality and consistency to allow us to make these assessments, which leave potential confounding variables unaddressed and we acknowledge identifies a limitation to the study.

Another limitation of our study is the retrospective design, which may represent a selection bias. Also, not all patients had peri-operative HbA1c measurements, though this data was available for 91% of the patients and was equally distributed between the three ethnic groups. We were not able to control for all major risk factors for the progression of diabetes and microvascular diseases, such as the presence of systemic hypertension and renal disease. Nevertheless, we feel that these factors may be less important, as we are evaluating a group of patients who already manifest a late complication of diabetes.

This study aimed to identify whether the outcome of surgical delamination or segmentation for traction complications of PDR was affected by the patients' ethnicity. Whilst black ethnicity may provide a protective influence on diabetes-associated macro-vascular disease,³⁰ our data strongly suggests that Black diabetic patients have significantly worse surgical and visual outcomes following diabetic delamination and segmentation surgery than White and South Asian patients. As poor surgical and subsequent functional outcomes lead to profound visual loss, we feel this ethnic difference needs to be highlighted so that further research can be performed in this area. Guthrie *et al*³¹ describe diabetic vitrectomy in E. Africa, and noted the beneficial effect of pre-treatment with Bevacizumab in a predominantly black population. It may be that use of a similar technique in the UK might lead to better results in Black patients. Currently, treatment and consenting discussions for severe PDR do not differentiate between different racial groups but given the potentially worse visual and surgical outcomes with long-term retention of silicone oil in black patients this distinction should be made.

Summary

What was known before

- Several studies demonstrated the role of ethnicity in the incidence and development of DR as well as other ophthalmic diseases, but until now racial differences in visual functionality after surgical treatment of severe PDR have never been demonstrated.

What this study adds

- This study shows a clear difference in visual and anatomic outcomes of surgical treatment of severe PDR based on ethnicity.

Conflict of interest

The authors declare no conflict of interest.

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