Supplemental Online Content

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This supplemental material has been provided by the authors to give readers additional information about their work.

eTable 1. Baseline factors of participants/study eye alive through 5 years with eyes assigned to ranibizumab

| Baseline Factors | Participants Treated with Ranibizumab | | | | | |
|---|--|--|--|--|--|--|
| No. of Participants | 170 | | | | | |
| Participant Factors | | | | | | |
| Sex, N (%) | | | | | | |
| Female | 76 (44.7%) | | | | | |
| Male | 94 (55.3%) | | | | | |
| Age (yrs.) | | | | | | |
| Median (Q1, Q3) | 51 (44, 59) | | | | | |
| Mean ± SD | 51 ± 12 | | | | | |
| No. of study eyes enrolled, N (%) | | | | | | |
| 1 | 89 (52.4%) | | | | | |
| 2 | 81 (47.6%) | | | | | |
| Race/ethnicity, N (%) | | | | | | |
| White | 88 (51.8%) | | | | | |
| Hispanic | 42 (24.7%) | | | | | |
| Black/African American | 35 (20.6%) | | | | | |
| Other | 5 (2.9%) | | | | | |
| Diabetes type , N (%) | | | | | | |
| Туре 1 | 40 (23.5%) | | | | | |
| Туре 2 | 124 (72.9%) | | | | | |
| Uncertain | 6 (3.5%) | | | | | |
| Duration of diabetes (yrs.) | | | | | | |
| Median (Q1, Q3) | 18 (12, 25) | | | | | |
| Mean ± SD | 19 ± 11 | | | | | |
| Hemoglobin A1c (%) ^a | | | | | | |
| Median (Q1, Q3) | 8.6 (7.5, 10.3) | | | | | |
| Mean ± SD | 8.9 ± 2.2 | | | | | |
| Functional Comorbidity Index | | | | | | |
| Median (Q1, Q3) | 0 (0, 1) | | | | | |
| Mean ± SD | 0.8 ± 1.1 | | | | | |
| Fellow eye visual acuity (letter score) ^b | | | | | | |
| Median (Q1, Q3) | 76 (64, 83) | | | | | |
| Snellen equivalent, Median (Q1, Q3) | 20/32 | | | | | |
| | (20/50, 20/25) | | | | | |
| Study-Eye Factors | | | | | | |
| Visual acuity (letter score) ^b | | | | | | |
| Median (Q1, Q3) | 78 (72, 84) | | | | | |
| Snellen equivalent, Median (Q1, Q3) | 20/32 (20/40, 20/20) | | | | | |
| OCT central subfield thickness (Stratus equivalent, µm) ^c | | | | | | |

| | Participants Treated with |
|---|---------------------------|
| Baseline Factors | Ranibizumab |
| Median (Q1, Q3) | 222 (196, 265) |
| Mean ± SD | 257 ± 104 |
| Presence of DME ^c , N (%) | |
| No | 123 (73.2%) |
| Yes | 45 (26.8%) |
| Neovascularization on clinical examination ^d , N (%) | |
| NVD or NVE only | 108 (65.5%) |
| NVD and NVE | 57 (34.5%) |
| Lens status on clinical examination, N (%) | |
| Phakic | 18 (10.6%) |
| Posterior chamber intraocular lens | 152 (89.4%) |
| Diabetic retinopathy severity (ETDRS level) c,e, N (%) | |
| Severe NPDR or less (≤level 53) | 19 (11.3%) |
| Inactive PDR (level 60) | 0 |
| Mild PDR (level 61) | 26 (15.5%) |
| Moderate PDR (level 65) | 62 (36.9%) |
| High risk PDR (level 71 and 75) | 58 (34.5%) |
| Advanced PDR (level 81 and 85) | 3 (1.8%) |
| Prior treatment for DME, N (%) | |
| No | 130 (76.5%) |
| Yes | 40 (23.5%) |
| Prior focal/grid laser for DME, N (%) | |
| No | 142 (83.5%) |
| Yes | 28 (16.5%) |
| Prior anti-VEGF treatment for DME, N (%) | |
| No | 150 (88.2%) |
| Yes | 20 (11.8%) |

Abbreviations: OCT, optical coherence tomography; DME, diabetic macular edema; NVD, neovascularization of the disc; NVE, neovascularization elsewhere; ETDRS, Early Treatment Diabetic Retinopathy Study; PDR, proliferative diabetic retinopathy; VEGF, vascular endothelial growth factor.

^a Unavailable for 6 participants.

^b Visual acuity letter scores indicate best-corrected visual acuity in the study eye following protocol-defined refraction. Visual acuity was measured using electronic Early Treatment for Diabetic Retinopathy Study (ETDRS) visual acuity testing; higher letter scores indicate better vision.

^c Unavailable for 2 participants.

^d Unavailable for 5 participants.

^e Determined by central reading center on 7 standard stereo field fundus photographs.

eTable 2. Multivariable analysis of baseline factors associated with a long lapse in care

| Baseline Study-Eye Factor | Odds Ratio (95% Confidence Interval) | P Value |
|---|---|---------|
| Visual acuity letter score (for every 5-letter decrease ^a) | 1.21 (1.03, 1.43) | .02 |
| Neovascularization clinical examination (NVD and NVE vs. NVD only or NVE only) | 2.19 (1.09, 4.38) | .03 |
| Prior focal/grid laser for DME (No vs. Yes) | 3.48 (1.38, 8.78) | .008 |

Abbreviations: DME, diabetic macular edema; NVD, neovascularization of the disc; NVE, neovascularization elsewhere.

A backward stepwise variable selection approach was used with entry selection criterion set at $P \le .10$ and stay criterion set at $P \le .05$.

^a A 5-letter difference in visual acuity letter score is equivalent to one Snellen line.

| eTable 3. Duration of follow-up at time of dropout and visual acuity at baseline |
|--|
| and the last completed visit |

| | Completed | Dropped Out in Year 4 or 5 | Dropped Out in Year 2 or 3 | Dropped Out in Year 1 | | |
|---|----------------|-------------------------------|-------------------------------|--------------------------|--|--|
| No. of Eyes | 120 | 14 | 21 | 15 | | |
| Baseline visual acuity letter score ^a | | | | | | |
| Mean ± SD | 77 ± 12 | 72 ± 12 | 74 ± 10 | 72 ± 13 | | |
| Snellen equivalent, Mean | 20/32 | 20/40 | 20/40 | 20/40 | | |
| Median (Q1, Q3) | 80 (75, 85) | 72 (64, 81) | 76 (70, 81) | 77 (68, 81) | | |
| Snellen equivalent, | 20/25 | 20/40 | 20/32 | 20/32 | | |
| Median (Q1, Q3) | (20/32, 20/20) | (20/50, 20/25) | (20/40, 20/25) | (20/50, 20/25) | | |
| Visual acuity letter score at last completed visit ^b | | | | | | |
| Mean ± SD | 79 ± 20 | 77 ± 13 | 74 ± 20 | 79 ± 11 | | |
| Snellen equivalent, Mean | 20/32 | 20/32 | 20/32 | 20/25 | | |
| Median (Q1, Q3) | 84 (78, 89) | 84 (64, 86) | 78 (74, 82) | 83 (75, 86) | | |
| Snellen equivalent, | 20/20 | 20/25 | 20/32 | 20/25 | | |
| Median (Q1, Q3) | (20/32, 20/16) | (20/50, 20/20) | (20/32, 20/25) | (20/32, 20/20) | | |
| Change in visual acuity letter score from baseline at last completed visit ^{a,b} | | | | | | |
| Mean ± SD | +1.5 ± 19.4 | +5.1 ± 12.3 | +0.3 ± 18.4 | +7.9 ± 7.0 | | |
| Median (Q1, Q3) | +4 (-2, +10) | +9 (0, +13) | +5 (+1, +7) | +5 (+3, +10) | | |

^a Visual acuity letter scores indicate best-corrected visual acuity in the study eye following protocol-defined refraction. Visual acuity was measured using electronic Early Treatment for Diabetic Retinopathy Study (ETDRS) visual acuity testing; higher letter scores indicate better vision. A 5-letter difference in visual acuity letter score is equivalent to one Snellen line.

^b Visual acuity was missing for 1 participant at the last completed visit. The last available visual acuity data was reported instead.