**Supplemental Table 1.** Generalized Least Squares Model-Predicted 48-mo WMH and Fold-Differences by Treatment

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| **Treatment group** | **Predicted WMH at 48m (95% CI)** | **Treatment:**  **PBO fold-difference (95% CI)** |
| PBO | 1.96 (1.86 to 2.06) | --- |
| oCEE | 2.07 (1.96 to 2.19) | 1.04 (0.99 to 1.09) |
| tE2 | 2.04 (1.94 to 2.15) | 1.03 (0.98 to 1.08) |