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SUPPLEMENTAL INFORMATION

Retinal Organoid Chip: Engineering a Physiomimetic Oxygen Gradient for Optimizing Long Term Culture of Human Retinal Organoids

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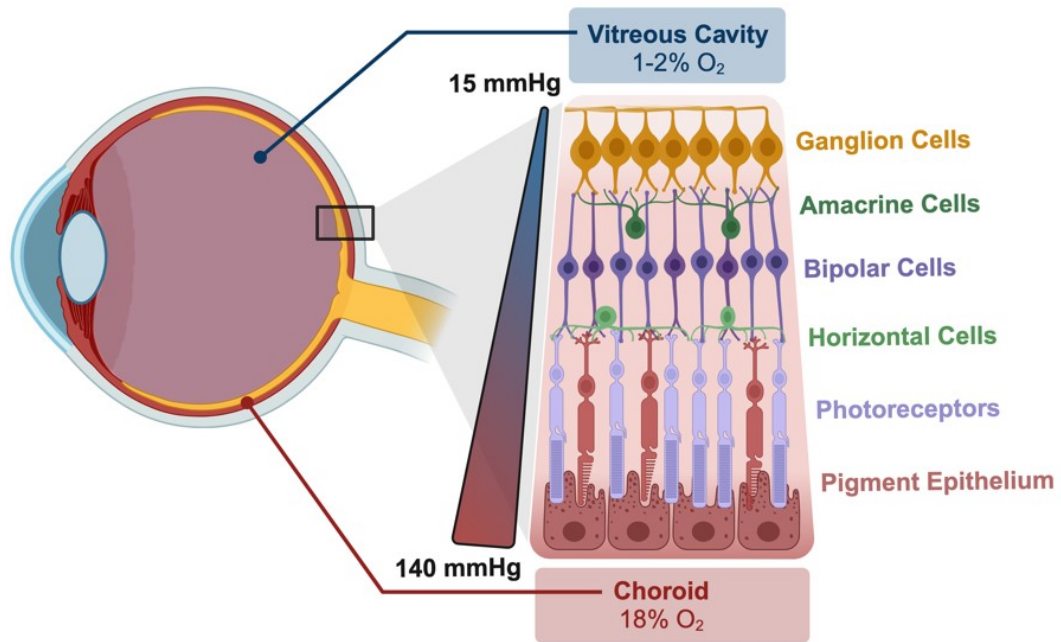
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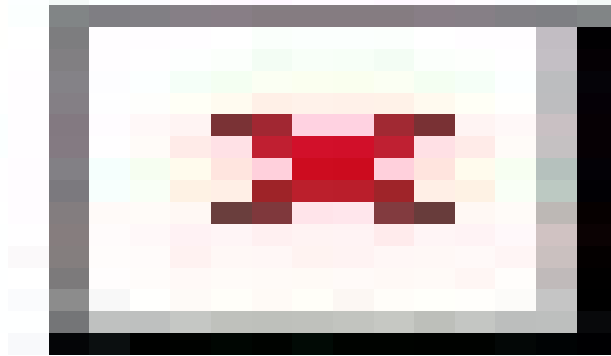
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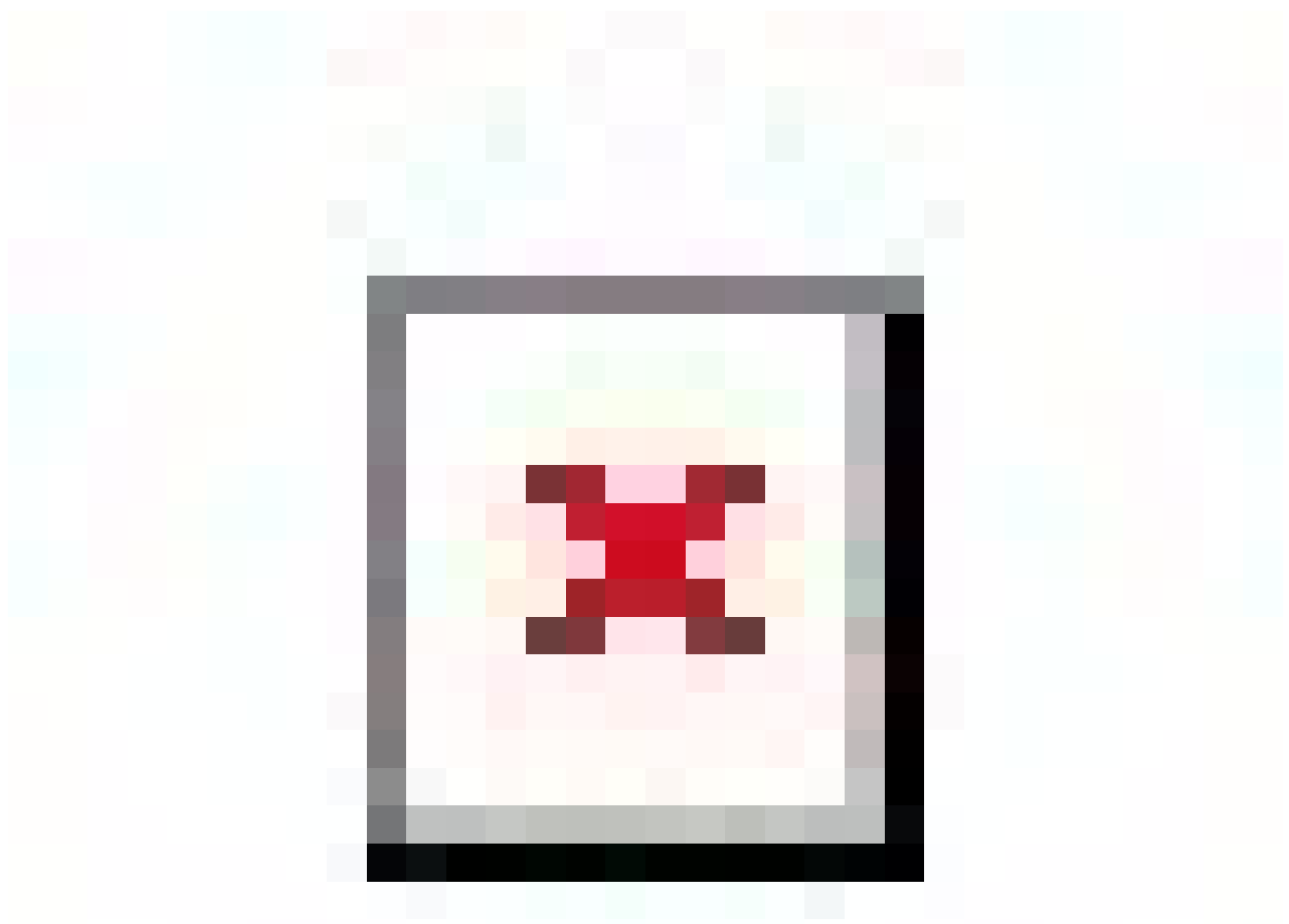
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27 **Supplementary Figure 1.** A schematic of the human eye anatomy highlights the retina, which is
 28 organized into distinct layers with various retinal cell phenotypes, situated between the hypoxic
 29 vitreous cavity and the oxygenated choroid. This positioning leads to the establishment of a steep
 30 oxygen gradient across the retinal tissue.



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33 **Supplementary Figure 2.** Dental Pulp Stem Cells (DPSC) reprogrammed for pluripotency were tested
34 on pluripotency markers SOX2, OCT4 and SSEA4 with immunofluorescence staining. Both original
35 DPSC derived iPSC line and a genetically modified iPSC line were found to be pluripotent.



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37 **Supplementary Figure 3.** *The workflow of RBPMS quantification from immunofluorescent images*
38 *using ImageJ, as described in the materials and methods.*

Supplementary Table 1: Computational Modeling Parameters

| | |
|------------------------------------|---|
| Physics & Study Type | Laminar Flow; Stationary Transport of Diluted Species; Time-dependent |
| Oxygen Diffusion Coefficients | Water: $3.0 \cdot 10^{-9} \text{ m}^2/\text{s}$ PFA: $4.56 \cdot 10^{-8} \text{ m}^2/\text{s}$ |
| Retinal Organoid Consumption Rates | Oxygen: $-0.034091 \text{ mol}/(\text{m}^3 \cdot \text{s})$ Glucose: $1.0278\text{E-}11 \text{ kg/s}$ |
| Boundary Conditions Oxygen | $C_{\text{water}} = 0.245 \text{ mol}/\text{m}^3$ $C_{\text{sodium sulfate channel}} = 0 \text{ mol}/\text{m}^3$ |
| Fluid Flow | $V = 50 \text{ }\mu\text{L}/\text{hour}$ |

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Supplementary Table 2: Retinal Organoid Culture Media

| Neural Induction Media (50 mL) | | |
|--|-------------------------------|------------------------------------|
| 48.5 mL | Gibco™ DMEM:F12 | Thermo Fisher Scientific #11320033 |
| 0.5 mL | Non-Essential Amino Acids | VWR #16777-186 |
| 0.5 mL | Gibco™ GlutaMAX™ Supplement | Thermo Fisher Scientific #35050061 |
| 50 µL | Heparin Solution | Stemcell Tehcnologies # 07980 |
| 0.5 mL | Gibco™ N-2 Supplement | Thermo Fisher Scientific #17502048 |
| Retinal Differentiation Media (50 mL) | | |
| 24 mL | Gibco™ DMEM:F12 | Thermo Fisher Scientific #11320033 |
| 24 mL | Gibco™ DMEM | Thermo Fisher Scientific #11965092 |
| 0.5 mL | Non-Essential Amino Acids | VWR #16777-186 |
| 0.5 mL | Gibco™ GlutaMAX™ Supplement | Thermo Fisher Scientific #35050061 |
| 1.0 mL | Gibco™ B27™ Supplement | Thermo Fisher Scientific #12587010 |
| 0.5 mL | Gibco™ Antibiotic-Antimycotic | Thermo Fisher Scientific #15240096 |

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