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1) Appendix Figure Legends S1 – S3

Appendix Figure Legend S1. Dual neutralization of VEGF-A and ANG-2 in JR5558 mice reduced the number of TUNEL-positive apoptotic nuclei in the photoreceptor layer.

A Bar/scatter graph of the number of TUNEL-positive cells in the retina of mice treated with 10 mg/kg IgG control, 5 mg/kg anti-VEGF-A, 5 mg/kg anti-ANG-2, 10 mg/kg anti-VEGF-A/ANG-2 or left untreated. Data shown as mean \pm SEM with n = 10 animals per group and * denotes all significant changes after one-sided ANOVA and Tukey's multiple t-test (Tukey-Kramer HSD). Anti-VEGF-A/ANG-2 (**, P = 0.007) and Anti-ANG-2 (*, P = 0.011) are significantly different from IgG control.

B Annotated images of representative examples of retina TUNEL staining of untreated (top left), IgG control (top middle;10 mg/kg), anti-VEGF-A (bottom left; 5 mg/kg), anti-ANG-2 (bottom middle; 5 mg/kg), and anti-VEGF-A/ANG-2 (bottom right; 10 mg/kg). For better visualization images were converted to black/white modus using PhotoShop CS6 with contrast enhancement applied equally to all images and non-tissue background was removed using Lasso tool. Scale bars = 500 μ m. SEM, standard error of the mean; ANOVA, analysis of variance.

Appendix Figure Legend S2. Coordinates of Iba1-Positive Cells on RPE/Choroid Flat Mounts

XY-coordinates of identified Iba1-positive cells per RPE/choroid were exported to GraphPad Prism and displayed as scatter plots to visualize distribution and clumping.

Appendix Figure Legend S3. Thumbnail images of annotated retinal flat mounts (TUNEL) and coordinates.

XY-coordinates of TUNEL-positive cells per retina were exported to GraphPad Prism and displayed as scatter plots to visualize distribution.



APPENDIX FIGURE S2 - COORDINATES OF IBA1-POSITIVE CELLS ON RPE/CHOROID FLAT MOUNTS

A) Group: Untreated

















APPENDIX FIGURE S2 - COORDINATES OF IBA1-POSITIVE CELLS ON <u>RPE/CHOROID FLAT MOUNTS</u> B) Group: IgG control

















APPENDIX FIGURE S2 - COORDINATES OF IBA1-POSITIVE CELLS ON <u>RPE/CHOROID FLAT MOUNTS</u> C) Group: Anti-VEGF-A















APPENDIX FIGURE S2 - COORDINATES OF IBA1-POSITIVE CELLS ON <u>RPE/CHOROID FLAT MOUNTS</u> D) Group: Anti-ANG-2

















APPENDIX FIGURE S2 - COORDINATES OF IBA1-POSITIVE CELLS ON <u>RPE/CHOROID FLAT MOUNTS</u> E) Group: anti-VEGF-A/ANG-2











X-Axis



APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES A) Group: Untreated





ID 911









APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES A) Group: Untreated





ID 941





APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES B) Group: IgG control















APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES B) Group: IgG control











APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES C) Group: Anti-VEGF-A













APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES C) Group: Anti-VEGF-A











APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES D) Group: Anti-ANG-2













Y-Axis



APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES D) Group: Anti-ANG-2









APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES E) Group: anti-VEGF-A/ANG-2















APPENDIX FIGURE S3 - THUMBNAIL IMAGES OF ANNOTATED RETINAL FLAT MOUNTS (TUNEL) AND COORDINATES E) Group: anti-VEGF-A/ANG-2







