## a.

WT (12 month)

b.

ADtg (12 month)

C.

d.


Supplementary figure 1. Addtional representative images and quantifications for figure 1. a-b. Representative images of acellular, degenerated retinal capillaries from a. a 12-month-old wild type (WT) mouse or b. a 12-month-old $\mathrm{APP}_{\text {SWE }} / \mathrm{PS} 1_{\triangle E 9}$ (ADtg) mouse. Red arrows indicate the degenerated capillaries. Scale bar $=20 \mu \mathrm{~m}$. c-d. Numbers of degenerated retinal capillaries whem mice are stratified by mouse age in $\mathbf{c}$. WT or d. ADtg mice groups. Data from individual mice (circles) as well as group means $\pm$ SEMs are shown. Fold changes are shown in red. Black-filled circles represent males and clear circles represent females. *p < $0.05,{ }^{* * * *} p<0.0001$, by one-way ANOVA with Tukey's post-hoc multiple comparison test.

$$
\begin{aligned}
& P_{I}=0.3964, F_{I}=2.774 \\
& P_{A}=0.0568, F_{A}=17.1 \\
& P_{G}=0.0011, F_{G}=85.29
\end{aligned}
$$

$$
P_{l}=0.4369, F_{1}=0.6697
$$

$$
P_{G e}=0.2702, \mathrm{~F}_{\mathrm{Ge}}=1.403
$$

a.

b.


$$
P_{G}=0.0055, F_{G}=14.21
$$

C.


Supplementary figure 2. Raw data of quantification of retinal vascular PDGFR $\beta$. a. Quantitative analysis of raw PDGFR $\beta$-immunoreactive (IR) area in each microscopic field of isolated retinal microvasculature from wild type (WT) ( $\mathrm{n}=6$ ) and APPswE/PS1 ${ }^{\mathrm{EE9}}$ ADtg ( $\mathrm{n}=6$ ) mice. $\mathbf{b}$. Quantification of PDGFR $\beta-I R$ in the same mice cohort when mice are stratified by genotypes of WT and ADtg with b. age of mice by 8 month and 12 month or c. sex of mice. Data from individual mice (circles) as well as from groups are shown with means $\pm$ SEMs. Black-filled circles represent males and empty circles represent females; ${ }^{*} p<0.05$, ** $p<$ 0.01, by two-way ANOVA with Tukey's post-hoc multiple comparison test. Two group statistical analysis was performed using an unpaired 2-tailed Student t-test. Percentage changes are shown in red.
a.

b.

$\mathrm{A} \beta(4 \mathrm{G} 8)$ Lectin DAPI i.
I.
C.

$P_{1}=0.4570, F_{1}=0.6107$
f. $P_{A}=0.4439, F_{A}=0.6485$

$P_{1}=0.7994, \mathrm{~F}_{1}=0.0691$
$P_{G \theta}=0.8208, F_{G e}=0.0548$
$P_{l}=0.4342, \mathrm{~F}_{1}=0.6779$
$P_{G \theta}=0.4199, F_{G e}=0.7228$
d. $P_{G}=0.0066, F_{G}=13.26$ e. $P_{G}=0.0097, F_{G}=11.41$

g.


[Area, $\mu \mathrm{m}^{2} \times 10^{3}$ ]

k.

n.


CD31 Thio-S
$A \beta_{40}(11 A 50)$

m.




Supplementary figure 3. Supplementary images, quantifications and correlations for figure 3. a-b. Representative fluorescence images of isolated retinal microvasculature stained for $A \beta$ ( 4 G 8 , magenta or red, as indicated under the images), blood vessels (lectin, green), and nuclei (DAPI, blue) in a perfused 8 -month-old APP SWE $^{2} / P S 1_{\triangle E 9}$ (ADtg) mice. Arrows indicate vascular $A \beta$. c. Quantitative analysis of the $A \beta(4 G 8)$-immunoreactive (IR) area normalized by lectin area in each microscopic field of isolated retinal microvasculature from wild type (WT) ( $n=6$ ) or ADtg ( $n=6$ ) mice. d-e. Quantitative analysis of the d. $A \beta(4 G 8)-I R$ area or e. A $\beta(4 G 8)-I R$ area normalized by lectin area in each microscopic field of isolated retinal microvasculature from the same cohort separated by different sex of mice. f. Quantitative analysis of the $A \beta$ (4G8)-IR area normalized by lectin area separated by different mice age groups ( 8 months and 12 months) and genotypes (WT and ADtg) in the same cohort. $n=3$ for each group. $g-h$. Pearson's coefficient ( $r$ ) correlation between the retinal $A \beta$ (4G8)-IR area against $\mathbf{g}$. degenerated capillaries or $h$. PDGFR $\beta$-IR area in ADtg mice ( $n=6$ ) of this cohort. i-j. Representative fluorescence images of isolated retinal microvasculature stained for $A \beta_{40}$ (11A50-B10, magenta), blood vessels (lectin, green), and nuclei (DAPI, blue) in a perfused 8-month-old ADtg mice. Arrows indicate vascular A $\beta$. k-n. Representative fluorescence images of retinal cross-section for thioflavin-S (Thio-S, green), A $3_{40}$ (11A50-B10, red) and blood vessels (CD31, blue) in a perfused 8-month-old $\mathbf{k}-\mathbf{m}$. WT or $\mathbf{n}$. ADtg mice. Data from individual mice (circles) as well as from groups are shown with means $\pm$ SEMs. Black-filled circles represent males and empty circles represent females; ${ }^{*} p<0.05$, ${ }^{* *} p<$ 0.01, by two-way ANOVA with Tukey's post-hoc multiple comparison test. Two group statistical analysis was performed using an unpaired 2 -tailed Student t-test. Fold changes are shown in red.


Supplementary figure 4. Additional data for figure 4. a-c. Densitometric analysis of western blot protein bands of a. claudin-1, b. ZO-1, and c. pNF-кB p65 with normalization, separated by mice age ( 4,8 , and 12 months) and genotype (WT and ADtg) in the same mice cohort as Figure 4. $\mathrm{n}=8$ for each group. d-f. Densitometric analysis of western blot protein bands of $\mathbf{d}$. Claudin-1, e. ZO-1 and f. pNF-кB p65 in the same mice cohort ( $\mathrm{n}=12$ for each group) separated by sex. Data from individual mouse (circles) as well as groups are shown as means $\pm$ SEMs. Black circles represent males and clear circles represent females. *p < 0.05, ** $p<$ 0.01, by one-way or two-way ANOVA with Tukey's post-hoc multiple comparison test. Two group statistical analysis was done by an unpaired 2-tailed Student t-test, and is shown in parenthesis. Percentage and fold changes are shown in red. g-h. Pearson's coefficient (r) correlation between retinal degenerated capillaries (Degen Caps) and the densitometric analysis of western blot protein bands of $\mathbf{g}$. ZO-1, or $\mathbf{h}$. pNF-кB p65 in the same mice cohort of figure 4 ( $n=48$ ). i-k. Pearson's coefficient ( $r$ ) correlation between retinal 4G8-immunoreactive area and the densitometric analysis of western blot protein bands of $\mathbf{i}$. claudin-1, $\mathbf{j}$. ZO-1, or $\mathbf{k}$. pNF-кB p65 in a subset of the APPSWE/PS1 ${ }_{\triangle E 9}$ (ADtg) mice cohort in figure 4 ( $\mathrm{n}=6$ ).
a.

## ADtg 16M


b.

ADtg 8M


Supplementary figure 5. Additional representative images for figure 5. a-b. Representative images of noninvasive retinal microvascular imaging after intraperitoneal fluorescein injection in a. 16-month-old APPSWE/PS1 1 E9 (ADtg) and b. 8-month-old ADtg mice.

