### Environ Health Perspect

### DOI: 10.1289/EHP9833

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### **Supplemental Material**

## Evaluation of Early Biomarkers of Atherosclerosis Associated with Polychlorinated Biphenyl Exposure: An *in Vitro* and *in Vivo* Study

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Figure S2. KEGG pathway analyses of genes targeted by differentially expressed miRNAs. High-throughput sequencing analysis of differentially expressed miRNAs in HUVECs with 5  $\mu$ M PCB29-pQ treatment for 24 h (n = 3). KEGG pathway enrichment in miRNA, the rich factor represents the degree of enrichment. The node size shows the number of selected genes, and the color scale represents -log (P value). The exact data are presented in Excel Table S4.

Figure S3. The expression of *HDAC7-AS1*, *MIR-7-5p* and Ago2 in HUVECs exposed to PCB29-pQ. RT-qPCR was performed to validate (a) lncRNAs and (b) miRNAs identified as differentially expressed by RNA sequencing assay.  $\beta$ -*Actin* was used as housekeeping genes for lncRNAs quantification and *U6* was used as housekeeping genes for miRNAs quantification, respectively. RP5-1057120.4 (termed as *HDAC7-AS1*) and *MIR-7-5p* were selected as target lncRNA and miRNA for further investigation (indicated with red boxes). Data are presented as mean  $\pm$  SD (n = 3). (c) Anti-Ago2 RNA binding protein immunoprecipitation assay was used to pull down endogenous RNAs associated with Ago2; IgG served as the control. Ago2 in proteins from Ago2-RIP assay was measured by western blotting. Data are presented as mean  $\pm$  SD (n = 3). P values were determined by two-way ANOVA, followed by Tukey's *post hoc* test. Data are graphed relative respective vehicle controls. The exact mean and SD value are presented in Table S20.

**Figure S4. The expression of** *HDAC7* **level in HUVECs transfected with** *HDAC7-AS1* **siRNA or pEZ-M61-HDAC7-AS1**. (a) Predicted relationship between ENST0000080059 (*HDAC7*) and ENST00000599515 (*HDAC7-AS1*) *via* RNAplex

(http://www.tbi.univie.ac.at/RNA/RNAplex.1.html) (b) After transfection with NC siRNA or *HDAC7-AS1* siRNA (25, 50, or 100 nM) for 48 h, the silencing efficiency of *HDAC7-AS1* siRNA in HUVECs was determined by RT-qPCR. Data are presented as mean  $\pm$  SD (n = 3). (c) After transfection with 2.5 µg pEZ-M61-NC or pEZ-M61-*HDAC7-AS1* (2.5, 5, or 7.5 µg) for 6 h, HUVECs were treated with 5 µM PCB29-pQ. *HDAC7-AS1* level was determined by RT-qPCR. Data are presented as mean  $\pm$  SD (n = 3). *HDAC7* expression of PCB29-pQ-exposed cells or control after (d) transfection with 25 nM NC siRNA or 25 nM *HDAC7-AS1* siRNA for 24 h or (e) transfection with 2.5 µg pEZ-M61-NC or 2.5 µg pEZ-M61-*HDAC7-AS1* for 6 h. Data are presented as mean  $\pm$  SD (n = 3). The housekeeping gene  $\beta$ -actin was used for quantification, and the primer information is shown Table S3. P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. For b and c, expression is shown relative to NC siRNA control and pEZ-M61-NC controls, respectively. The exact mean and SD values are presented in Table S21.

**Figure S5. The localization of LncRNA** *HDAC7-AS1* in HUVECs. HUVECs were treated with 5  $\mu$ M PCB29-pQ for 24 h. (n = 3). Fluorescent *in situ* hybridization (FISH) assay was used to investigate the localization of *HDAC7-AS1* in in the cells. *HDAC7-AS1* was stained red using an RNA probe, and the nucleus was stained blue with DAPI. Scale bar = 10  $\mu$ m.

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**Figure S7. Target genes TGF-β2 and PPME1 protein levels in HUVECs exposed to PCB29pQ with** *MIR-7-5p* **inhibitor or** *TGF-β2/PPME1* **siRNA**. After co-transfection with 100 nM NC inhibitor or *MIR-7-5p* **inhibitor** and NC siRNA or *TGF-β2/PPME1* siRNA (25 nM) for 24 h, HUVECs were treated with 5 µM PCB29-pQ for 24 h. (n = 3). Protein levels of (**a**) PPME1 and (**b**) TGF-β2 were detected by western blotting (upper panel). The relative protein expression levels (lower panel) were quantified by ImageJ software. Data are presented as mean ± SD (n = 3). β-Actin was used as an internal loading control. Data are graphed relative to the expression in cells exposed to the NC inhibitor and NC siRNA together. The exact mean and SD values are presented in Table S22. **Figure S8.** Luciferase analysis of the activity of *MIR-7-5p* bind to *TGF-β2, PPME1*, and *HDAC7-AS1* in HUVECs transfected with NC mimic or *MIR-7-5p* mimic. (a) HUVECs were transfected with NC mimic or *MIR-7-5p* mimic for 48 h, together with NC-3'UTR, *TGF-β2* (WT)-3'UTR, or *TGF-β2* (MUT)-3'UTR. (b) HUVECs were transfected with 50 nM NC mimic or 50 nM *MIR-7-5p* mimic for 48 h, together with NC-3'UTR, *PPME1* (WT)-3'UTR, or *PPME1* (MUT)-3'UTR. (c) HUVECs were transfected with 50 nM NC mimic or 50 nM *MIR-7-5p* mimic for 48 h, together with Luc-NC, Luc-*HDAC7-AS1-*WT, or Luc-*HDAC7-AS1-*MUT. For all panels, Gaussia luciferase activity was analyzed, corresponding to *TGF-β2, PPME1*, and *HDAC7-AS1* transcription. Data are presented as mean  $\pm$  SD (n = 3). P values were determined by two-way ANOVA, followed by Tukey's *post hoc* test. Data are graphed relative to the NC-mimic treated, NC-3'UTR controls. The exact mean and SD values are presented in Table S23.

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Figure S12. HE staining of aortic root cross-sections and TC and TG levels in *ApoE<sup>-/-</sup>* mice were intravenous (*i.v.*) injected with *AAV-HDAC7-AS1*. Male *ApoE<sup>-/-</sup>* mice were *i.v.* injected with AAV-*HDAC7-AS1 via* tail vein  $(4 \times 10^{10} \text{ particles/mouse})$  to create an *HDAC7-AS1* overexpressed mice model. Male *ApoE<sup>-/-</sup>* mice that received the AAV vector were used as AAV control mice. Control and *HDAC7-AS1* overexpressed *ApoE<sup>-/-</sup>* mice were fed a western high-fat diet for 12 weeks and tap water ad libitum. Mice were injected with 5 mg/kg body weight of PCB29-pQ or equal volumes of corn oil by *i.p.* injection once a week for 12 continuous weeks with the first two injections during the 1st week (3 days apart). (a) (Left panel) HE of the aortic root (Right panel) plaque area in the aortic root was quantified by ImageJ software. Data are presented as mean ± Standard Deviation (SD). Scale bar = 200 µm. (b) TC level and (c) LDL-C level. Data are presented as mean ±SD (n = 5). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. The exact mean and SD values are presented in Table S26.

**Figure S13. CAV1 phosphorylation and TGF-β, PPME1, and inflammatory factor levels in HUVECs exposed PCB29-pQ with** *CAV1* **siRNA.** (a) HUVECs were treated with 5 μM PCB29-pQ for 1, 3, and 6 h. (upper panel) Protein levels of p-CAV1 and CAV1 were detected by western blotting. β-Actin was used as an internal loading control. The relative protein expression levels (lower panel) were quantified by ImageJ software. Data are presented as mean ± SD (n = 3). After transfection with 25 nM NC siRNA or *CAV1* siRNA for 48 h, HUVECs were treated with 5 μM PCB29-pQ for 24 h. The siRNA information is shown in Table S6. (b) Protein levels of p-CAV1, CAV1, IL-1β, IL-6, and TNFα were detected by western blotting. β-Actin was used as an internal loading control. (c-h) p-CAV1, CAV1, IL-1β, IL-6, and TNFα were quantified by ImageJ software. Data are presented as mean ± SD (n = 3). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. Data are graphed relative to the NC siRNA group. The exact mean and SD values are presented in Table S27.

## Figure S14. p-CAV1 level in *ApoE<sup>-/-</sup>* mice that were *i.v.* injected with AAV-*HDAC7-AS1*.

Male  $ApoE^{-/-}$  mice were i.v. injected with AAV-*HDAC7-AS1 via* tail vein (4×10<sup>10</sup> particles/mouse) to create an *HDAC7-AS1* overexpressed mice model. Male  $ApoE^{-/-}$  mice that received the AAV vector were considered as AAV control mice. Control and *HDAC7-AS1* overexpressed  $ApoE^{-/-}$  mice were fed a Western high-fat diet for 12 weeks and tap water ad *libitum*. Mice were injected with 5 mg/kg body weight of PCB29-pQ or equal volumes of corn oil by *i.p.* injection once a week for 12 continuous weeks with the first two injections during the 1<sup>st</sup> week (3 days apart). (Left panel) The presence of p-CAV1 in aortic root cross-sections was detected by double immunostaining with the use of antibodies against p-CAV1 (red), endothelial cell marker CD31 (green), and nucleus marker DAPI (blue), respectively. Scale bar = 100 µm. White arrows represent the colocalization of p-CAV1 (red) and CD31 (green). (Right panel) Colocalization of p-CAV1 and CD31 was analyzed by Pearson's correlation coefficient. Data were presented as mean ± SD (n = 3). The exact mean and SD values are presented in Table S28.

**Figure S15. Gene type identification of** *ApoE* and *CAV1* knockout mice. DNA from the tail of offspring mice was extracted and analyzed by PCR and agarose gel electrophoresis. (a) Homozygous of *ApoE*: 245 bp; Heterozygous of ApoE: 245 bp & 155 bp; WT of ApoE: 155 bp. (b) Homozygous of *CAV1*: 410 bp; Heterozygous of *CAV1*: 690 bp & 410 bp; WT of *CAV1*: 690 bp. The PCR primer sequences for *ApoE* and *CAV1* mice genotyping are shown in Table S2.

Figure S16. Immunohistochemistry staining of CD68 and TC and LDL-C levels in  $ApoE^{-/-}$  and  $ApoE^{-/-}/CAVI^{-/-}$  mice.  $ApoE^{-/-}$  mice were crossed with  $CAVI^{-/-}$  mice to generate  $ApoE^{-/-}/CAVI^{-/-}$  mice.  $ApoE^{-/-}/CAVI^{-/-}$  mice were fed a Western high-fat diet for 12 weeks and tap water ad libitum. Mice were treated with 5 mg/kg body weight of PCB29-pQ or equal volumes of corn oil by *i.p.* injection. (a) (Left panel) Immunohistochemistry staining of CD68 detected macrophage infiltration located at the aortic wall. Scale bar = 200 µm. (Right panel) Quantification proportion of positive CD68 expressions area to total aortic section using was performed by ImageJ software. Data are presented as mean  $\pm$  SD (n = 3). Serum (b) TC level and (c) LDL-C levels. Data are presented as mean  $\pm$ SD (n = 5). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. The exact mean and SD values are presented in Table S29.

Figure S17. Adhesion molecules, pro-inflammatory cytokines and p65 protein expression levels in HUVEC exposed to PCB29-pQ. HUVECs were treated with 5 µM PCB29-pQ for 1, 3, or 6 h. (a) IL-1β, IL-6, TNF-α, ICAM-1, and VCAM-1 expressions in cell lysates were analyzed by western blotting. (**b-f**) IL-1 $\beta$ , IL-6, TNF- $\alpha$ , ICAM-1, and VCAM-1 expressions were quantified by ImageJ software. Data are presented as mean  $\pm$  Standard Deviation (SD) (n = 3). (g) Calcein-AM-loaded THP-1 cells (10<sup>6</sup> cells/ml) were added to HUVECs and then incubated for 3 h. The unbound cells were washed off, and attached fluorescent monocytes were visualized using an optical microscope. (Left panel) The fluorescence intensity of Calcein-AM was quantified using ImageJ software. Data are presented as mean  $\pm$  SD (n = 3). P value was determined by unpaired Student's t-test. (h) HUVECs were treated with 5 µM PCB29-pQ for 1, 3, or 6 h. IkBa and p-p65 levels were analyzed by western blotting analysis.  $\beta$ -Actin was used as an internal loading control. (i-j) IkBa and p-p65 levels were quantified by ImageJ software. Data are presented as mean  $\pm$  SD (n = 3). (k) HUVECs were pretreated with 5  $\mu$ M p65 inhibitor PDTC for 1 h, followed with 5  $\mu$ M PCB29-pQ exposure for 6 h. IL-1 $\beta$ , IL-6, and TNF- $\alpha$  in cell lysates were analyzed by western blotting. β-Actin was used as an internal loading control. (l-n) IL-1β, IL-6, and TNF- $\alpha$  protein expression levels were quantified by ImageJ software. Data were presented as mean  $\pm$  SD (n = 3). P values were determined by one-way ANOVA, followed by Tukey's *post* hoc test. The exact mean and SD values are presented in Table S30.

**Figure S18.** Adhesion molecules, pro-inflammatory cytokines and p65 protein expression levels in HUVEC exposed to PCB29-pQ with *CAV1* siRNA. HUVECs were transfected with NC siRNA or 25 nM *CAV1* siRNA for 48 h and then treated with 5 µM PCB29-pQ for 6 h. (a) IL-1β, IL-6, TNF-α, ICAM-1 and VCAM-1 expressions were analyzed by western blotting. β-Actin was used as an internal loading control. (b-f) IL-1β, IL-6, TNF-α, ICAM-1 and VCAM-1 expression levels were quantified by ImageJ software. Data were presented as mean ± Standard Deviation (SD) (n = 3). (g) IkBα, p-p65, and p65 expressions were analyzed by western blotting. β-Actin was used as an internal loading control. (h-i) IkBα, p-p65, and p65 expression levels were quantified by ImageJ software. Data were presented as mean ± SD (n = 3). (j) (Left panel) Immunofluorescence analysis of p65. Green staining represents the location of p65. Nuclei shown in blue were stained with DAPI. Scale bar = 10 µm. (Right panel) Co-localization of p-CAV1 and DAPI was analyzed by Pearson's correlation coefficient. Data were presented as mean ± SD (n = 3). P values were determined by one-way ANOVA, followed by Tukey's post hoc test. The exact mean and SD values are presented in Table S31. Figure S19. Inflammatory factors and ROS levels in HUVEC exposed to PCB29-pQ. HUVECs were pretreated with 40  $\mu$ M VC, 20  $\mu$ M VE or 5 mM NAC for 1 h, followed with 5  $\mu$ M PCB29-pQ exposure for 6 h. (n = 3). (a) p-p65, p65, IL-1 $\beta$ , IL-6, TNF $\alpha$ , ICAM-1 and VCAM-1 in cell lysates were analyzed by western blotting. (b-g) p-p65, p65, IL-1 $\beta$ , IL-6, TNF $\alpha$ , ICAM-1 and VCAM-1 and VCAM-1 expression levels were quantified by ImageJ software. Data were presented as mean  $\pm$  Standard Deviation (SD) (n = 3). (h) ROS levels were detected by DCFH-DA (10  $\mu$ M) probe. Data were presented as mean  $\pm$  SD (n = 3). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. In all graphs, data is normalized to the control cell groups. The exact mean and SD values are presented in Table S32.

Figure S20. p-CAV1 and CAV1 levels in HUVEC exposed to PCB29-pQ and antioxidants.

HUVECs were pretreated with 40  $\mu$ M VC, 20  $\mu$ M VE or 5 mM NAC for 1 h, followed by 5  $\mu$ M PCB29-pQ exposure for 6 h. HUVECs were pretreated with (a) 40  $\mu$ M VC, 20  $\mu$ M VE or 5 mM NAC, (b) 200 U/ml PEG-SOD, 500 U/ml PEG-CAT and 5 mM GSH-MEE for 1 h, followed with 5  $\mu$ M PCB29-pQ exposure for 6 h. (Left panel) p-CAV1 and CAV1 in cell lysates were analyzed by western blotting.  $\beta$ -Actin was used as an internal loading control. (Right panel) p-CAV1 and CAV1 expression levels were quantified by ImageJ software. Data were presented as mean  $\pm$  Standard Deviation (SD) (n = 3). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. All data is graphed relative to the control cell groups. The exact mean and SD values are presented in Table S33.

**Figure S21. Heatmap showing correlation scores between factors.** Plasma RNA was extracted by TRNzol universal reagent. CHD group (n = 77) contains patients with > 50% coronary artery stenosis, and the control group (n = 50) contains subjects with < 50% coronary artery stenosis. RT-qPCR analysis of *HDAC7-AS1*, *MIR-7-5p*, *TGF-β2*, *PPME1*, *IL-1β*, *IL6* and *TNFα* expressions. Spearman's rank correlation coefficient was used to assess the correlation between the two indicated factors. P value of < 0.05 was considered significant. The size and color of circle represents the correlation between two factors. X's mean no significance. Summary data can be found in Table S11.

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Table S26. HE staining of aortic root cross-sections and total cholesterol (TC) and lowdensity lipoprotein (LDL) levels in  $ApoE^{-/-}$  mice were *i.v.* injected with AAV-HDAC7-AS1.

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**Table S28.** Pearson's correlation coefficient for expression of **p-CAV1 and** CD31 in *ApoE*-/- mice that *were i.v.* injected with AAV-*HDAC7-AS1*.

Table S29. Immunohistochemistry staining of CD68, total cholesterol (TC), and low-density lipoprotein level (LDL) in *ApoE<sup>-/-</sup>* and *ApoE<sup>-/-</sup>/CAV1<sup>-/-</sup>* mice.

Table S30. Adhesion molecules, pro-inflammatory cytokines and *p65* protein expression level in HUVEC exposed to PCB29-pQ.

Table S31. Adhesion molecules, pro-inflammatory cytokines and *p65* protein expression levels in HUVEC exposed to PCB29-pQ with *CAV1* siRNA.

Table S32. Protein expression of inflammatory factors and ROS levels in HUVEC exposed to PCB29-pQ.

Table S33. p-CAV1 protein expression relative to CAV1 total protein in HUVEC exposed to PCB29-pQ and antioxidants.

Additional File- Excel Document



Figure S1. Functional analyses of genes targeted by differentially expressed miRNAs. Highthroughput sequencing analysis of differentially expressed miRNAs after 5  $\mu$ M PCB29-pQ treatment of HUVECs for 24 h (n = 3). Target gene of differentially expressed genes were screened and functional analyses performed. Gene ontology has three ontologies: molecular function, cellular component, and biological process. The exact data are presented in Excel Table S3.



Figure S2. KEGG pathway analyses of genes targeted by differentially expressed miRNAs. Highthroughput sequencing analysis of differentially expressed miRNAs in HUVECs with 5  $\mu$ M PCB29pQ treatment for 24 h (n = 3). KEGG pathway enrichment in miRNA, the rich factor represents the degree of enrichment. The node size shows the number of selected genes, and the color scale represents -log (P value). The exact data are presented in Excel Table S4.



Figure S3. The expression of *HDAC7-AS1*, *MIR-7-5p* and Ago2 in HUVECs exposed to PCB29pQ. RT-qPCR was performed to validate (a) lncRNAs and (b) miRNAs identified as differentially expressed by RNA sequencing assay.  $\beta$ -*Actin* was used as housekeeping genes for lncRNAs quantification and *U6* was used as housekeeping genes for miRNAs quantification, respectively. RP5-1057120.4 (termed as *HDAC7-AS1*) and *MIR-7-5p* were selected as target lncRNA and miRNA for further investigation (indicated with red boxes). Data are presented as mean  $\pm$  SD (n = 3). (c) Anti-Ago2 RNA binding protein immunoprecipitation assay was used to pull down endogenous RNAs associated with Ago2; IgG served as the control. Ago2 in proteins from Ago2-RIP assay was measured by western blotting. Data are presented as mean  $\pm$  SD (n = 3). P values were determined by two-way ANOVA, followed by Tukey's *post hoc* test. Data are graphed relative respective vehicle controls. The exact mean and SD value are presented in Table S20.



Figure S4. The expression of *HDAC7* level in HUVECs transfected with *HDAC7-AS1* siRNA or pEZ-M61-*HDAC7-AS1*. (a) Predicted relationship between ENST0000080059 (*HDAC7*) and ENST00000599515 (*HDAC7-AS1*) via RNAplex (http://www.tbi.univie.ac.at/RNA/RNAplex.1.html) (b) After transfection with NC siRNA or *HDAC7-AS1* siRNA (25, 50, or 100 nM) for 48 h, the silencing efficiency of *HDAC7-AS1* siRNA in HUVECs was determined by RT-qPCR. Data are presented as mean  $\pm$  SD (n = 3). (c) After transfection with 2.5 µg pEZ-M61-NC or pEZ-M61-*HDAC7-AS1* level was determined by RT-qPCR. Data are presented as mean  $\pm$  SD (n = 3). (c) After transfection with 5 µM PCB29-pQ. *HDAC7-AS1* level was determined by RT-qPCR. Data are presented as mean  $\pm$  SD (n = 3). *HDAC7* expression of PCB29-pQ-exposed cells or control after (d) transfection with 25 nM NC siRNA or 25 nM *HDAC7-AS1* siRNA for 24 h or (e) transfection with 2.5 µg pEZ-M61-NC or 2.5 µg pEZ-M61-*HDAC7-AS1* for 6 h. Data are presented as mean  $\pm$  SD (n = 3). The housekeeping gene *β-actin* was used for quantification, and the primer information is shown Table S3. P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. For b and c, expression is shown relative to NC siRNA control and pEZ-M61-NC controls, respectively. The exact mean and SD values are presented in Table S21.



Figure S5. The localization of LncRNA *HDAC7-AS1* in HUVECs. HUVECs were treated with 5  $\mu$ M PCB29-pQ for 24 h. (n = 3). Fluorescent *in situ* hybridization (FISH) assay was used to investigate the localization of *HDAC7-AS1* in in the cells. *HDAC7-AS1* was stained red using an RNA probe, and the nucleus was stained blue with DAPI. Scale bar = 10  $\mu$ m.



Figure S6. Volcano plots analyzing differential expression with mRNA sequencing (mRNA-Seq) in the control group and PCB29-pQ group. The abscissa represents the logarithmic values of two different groups, and the ordinate represents mRNAs differences (fold change  $\geq 2$  and P value < 0.05) between two groups. Red dots indicated PCB29-pQ group is higher relative to control group). Green dots indicated lower expression in PCB29-pQ group than control group (judgment standard is P value <0.05, and the difference multiple is more than 2). Black dots indicated no difference between two groups.



Figure S7. Target genes TGF-B2 and PPME1 protein levels in HUVECs exposed to PCB29-pQ

with *MIR-7-5p* inhibitor or *TGF-β2/PPME1* siRNA. After co-transfection with 100 nM NC inhibitor or *MIR-7-5p* inhibitor and NC siRNA or *TGF-β2/PPME1* siRNA (25 nM) for 24 h, HUVECs were treated with 5  $\mu$ M PCB29-pQ for 24 h. (n = 3). Protein levels of (**a**) PPME1 and (**b**) TGF-β2 were detected by western blotting (upper panel). The relative protein expression levels (lower panel) were quantified by ImageJ software. Data are presented as mean  $\pm$  SD (n = 3).  $\beta$ -Actin was used as an internal loading control. Data are graphed relative to the expression in cells exposed to the NC inhibitor and NC siRNA together. The exact mean and SD values are presented in Table S22.

b



Figure S8. Luciferase analysis of the activity of *MIR-7-5p* bind to *TGF-β2*, *PPME1*, and *HDAC7-AS1* in HUVECs transfected with NC mimic or *MIR-7-5p* mimic. (a) HUVECs were transfected with NC mimic or *MIR-7-5p* mimic for 48 h, together with NC-3'UTR, *TGF-β2* (WT)-3'UTR, or *TGF-β2* (MUT)-3'UTR. (b) HUVECs were transfected with 50 nM NC mimic or 50 nM *MIR-7-5p* mimic for 48 h, together with NC-3'UTR, *PPME1* (WT)-3'UTR, or *PPME1* (MUT)-3'UTR. (c) HUVECs were transfected with 50 nM NC mimic or 50 nM *MIR-7-5p* mimic for 48 h, together with Luc-NC, Luc-*HDAC7-AS1*-WT, or Luc-*HDAC7-AS1*-MUT. For all panels, Gaussia luciferase activity was analyzed, corresponding to *TGF-β2*, *PPME1*, and *HDAC7-AS1* transcription. Data are presented as mean  $\pm$  SD (n = 3). P values were determined by two-way ANOVA, followed by Tukey's *post hoc* test. Data are graphed relative to the NC-mimic treated, NC-3'UTR controls. The exact mean and SD values are presented in Table S23.



Figure S9. The tube forming ability in HUVECs exposed to PCB29-pQ and transfected with a *MIR-7-5p* inhibitor. (Left panel) After transfection with 100 nM NC inhibitor or 100 nM *MIR-7-5p* inhibitor for 48 h, followed by exposure to 5  $\mu$ M PCB29-pQ for 24 h, cells were plated on Matrigel to conduct tube formation assay. Scale bar = 200  $\mu$ m. (Right panel) Quantification of tube formation through measurement of branch point number with ImageJ software. Data were presented as mean  $\pm$  SD (n = 3). P values were determined by two-way ANOVA, followed by Tukey's *post hoc* test. The exact mean and SD values are presented in Table S24.



Figure S10. Apoptosis and proliferation rates in HUVECs exposed to PCB29-pQ and transfected

with a *MIR-7-5p* mimic or pEZ-M61-*HDAC7-AS1*. (a) Cell apoptosis was assessed by Annexin V-FITC/PI double staining with a flow cytometer. (n = 3). (b) Cell proliferation was measured by BrdU/PI double staining with a flow cytometer. (n = 3).



Figure S11. *HDAC7-AS1*, *MIR-7-5p*, *TGF-β2*, *PPME1* mRNA levels, TGF-β2 and PPME1 protein levels, and apoptosis or proliferation rates in HUVECs exposed to PCB29-pQ and transfected with a *MIR-7-5p* inhibitor or *HDAC7-AS1* siRNA. After co-transfection with 100 nM NC inhibitor

or 100 nM *MIR-7-5p* inhibitor and NC siRNA or *HDAC7-AS1* siRNA (25 nM) for 24 h, HUVECs were treated with 5  $\mu$ M PCB29-pQ for 24 h. RNAs expression of (**a**) *MIR-7-5p*, (**b**) *HDAC7-AS1*, (**c**) *TGF-β2* and (**d**) *PPME1* were detected by RT-qPCR. *β-actin* was used as a housekeeping gene, except U6 was used for *MIR-7-5p* quantification. The primer information is shown in Table S3-4. (**e**) Protein levels of TGF-β2 and PPME1 were detected by western blotting. β-Actin was used as an internal loading control. (**f**) TGF-β2, (**g**) PPME1 protein expression levels were quantified by ImageJ software. Data were presented as mean ± Standard Deviation (SD) (n = 3). (**h**) Cell viability was measured by CCK-8 kit. Data were presented as mean ± SD (n = 3). (**i**) Cell apoptosis was performed by Annexin V-FITC/PI assay with a flow cytometer. (**j**) Cell proliferation was performed by BrdU/PI staining. P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. Data are graphed relative to the cells exposed to NC inhibitor, NC siRNA, and vehicle control. The exact mean and SD values are presented in Table S25.



Figure S12. HE staining of aortic root cross-sections and TC and TG levels in  $ApoE^{-/-}$  mice were intravenous (*i.v.*) injected with AAV-HDAC7-AS1. Male  $ApoE^{-/-}$  mice were *i.v.* injected with AAV-HDAC7-AS1 via tail vein (4×10<sup>10</sup> particles/mouse) to create an HDAC7-AS1 overexpressed mice model. Male  $ApoE^{-/-}$  mice that received the AAV vector were used as AAV control mice. Control and HDAC7-AS1 overexpressed  $ApoE^{-/-}$  mice were fed a western high-fat diet for 12 weeks and tap water *ad libitum*. Mice were injected with 5 mg/kg body weight of PCB29-pQ or equal volumes of corn oil by *i.p.* injection once a week for 12 continuous weeks with the first two injections during the 1<sup>st</sup> week (3 days apart). (**a**) (Left panel) HE of the aortic root (Right panel) plaque area in the aortic root was quantified by ImageJ software. Data are presented as mean ± Standard Deviation (SD). Scale bar = 200 µm. (**b**) TC level and (**c**) LDL-C level. Data are presented as mean ±SD (n = 5). P values were

determined by one-way ANOVA, followed by Tukey's *post hoc* test. The exact mean and SD values are presented in Table S26.



Figure S13. CAV1 phosphorylation and TGF-β, PPME1, and inflammatory factor levels in HUVECs exposed PCB29-pQ with *CAV1* siRNA. (a) HUVECs were treated with 5 µM PCB29-pQ for 1, 3, and 6 h. (upper panel) Protein levels of p-CAV1 and CAV1 were detected by western blotting. β-Actin was used as an internal loading control. The relative protein expression levels (lower panel) were quantified by ImageJ software. Data are presented as mean  $\pm$  SD (n = 3). After transfection with 25 nM NC siRNA or *CAV1* siRNA for 48 h, HUVECs were treated with 5 µM PCB29-pQ for 24 h. The siRNA information is shown in Table S6. (b) Protein levels of p-CAV1, CAV1, IL-1β, IL-6, and TNFα were detected by western blotting. β-Actin was used as an internal loading control. (**c-h**) p-CAV1, CAV1, IL-1β, IL-6, and TNFα were quantified by ImageJ software. Data are presented as mean

 $\pm$  SD (n = 3). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. Data are graphed relative to the NC siRNA group. The exact mean and SD values are presented in Table S27.



**Figure S14.** p-CAV1 level in  $ApoE^{-L}$  mice that were *i.v.* injected with AAV-*HDAC7-AS1*. Male  $ApoE^{-L}$  mice were *i.v.* injected with AAV-*HDAC7-AS1 via* tail vein (4×10<sup>10</sup> particles/mouse) to create an *HDAC7-AS1* overexpressed mice model. Male  $ApoE^{-L}$  mice that received the AAV vector were considered as AAV control mice. Control and *HDAC7-AS1* overexpressed  $ApoE^{-L}$  mice were fed a Western high-fat diet for 12 weeks and tap water *ad libitum*. Mice were injected with 5 mg/kg body weight of PCB29-pQ or equal volumes of corn oil by *i.p.* injection once a week for 12 continuous weeks with the first two injections during the 1<sup>st</sup> week (3 days apart). (Left panel) The presence of p-CAV1 in a ortic root cross-sections was detected by double immunostaining with the use of antibodies against p-CAV1 (red), endothelial cell marker CD31 (green), and nucleus marker DAPI (blue), respectively. Scale bar = 100 µm. White arrows represent the colocalization of p-CAV1 (red) and CD31 (green). (Right panel) Co-localization of p-CAV1 and CD31 was analyzed by Pearson's correlation coefficient. Data were presented as mean  $\pm$  SD (n = 3). The exact mean and SD values are presented in Table S28.



**Figure S15. Gene type identification of** *ApoE* **and** *CAV1* **knockout mice.** DNA from the tail of offspring mice was extracted and analyzed by PCR and agarose gel electrophoresis. (**a**) Homozygous of *ApoE*: 245 bp; Heterozygous of *ApoE*: 245 bp & 155 bp; WT of *ApoE*: 155 bp. (**b**) Homozygous of *CAV1*: 410 bp; Heterozygous of *CAV1*: 690 bp & 410 bp; WT of *CAV1*: 690 bp. The PCR primer sequences for *ApoE* and *CAV1* mice genotyping are shown in Table S2.



Figure S16. Immunohistochemistry staining of CD68 and TC and LDL-C levels in  $ApoE^{-/-}$  and  $ApoE^{-/-}/CAVI^{-/-}$  mice.  $ApoE^{-/-}/CAVI^{-/-}$  mice were crossed with  $CAVI^{-/-}$  mice to generate  $ApoE^{-/-}/CAVI^{-/-}$  mice.  $ApoE^{-/-}$  or  $ApoE^{-/-}/CAVI^{-/-}$  mice were fed a Western high-fat diet for 12 weeks and tap water *ad libitum*. Mice were treated with 5 mg/kg body weight of PCB29-pQ or equal volumes of corn oil by *i.p.* injection. (a) (Left panel) Immunohistochemistry staining of CD68 detected macrophage infiltration located at the aortic wall. Scale bar = 200 µm. (Right panel) Quantification proportion of positive CD68 expressions area to total aortic section using was performed by ImageJ software. Data are presented as mean  $\pm$  SD (n = 3). Serum (b) TC level and (c) LDL-C levels. Data are presented as mean  $\pm$ SD (n = 5). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. The

Control

PCB29-pQ

PCB29-pQ

Control

exact mean and SD values are presented in Table S29.



Figure S17. Adhesion molecules, pro-inflammatory cytokines and p65 protein expression levels

in HUVEC exposed to PCB29-pQ. HUVECs were treated with 5  $\mu$ M PCB29-pQ for 1, 3, or 6 h. (a) IL-1 $\beta$ , IL-6, TNF- $\alpha$ , ICAM-1, and VCAM-1 expressions in cell lysates were analyzed by western blotting. (b-f) IL-1 $\beta$ , IL-6, TNF- $\alpha$ , ICAM-1, and VCAM-1 expressions were quantified by ImageJ

software. Data are presented as mean  $\pm$  Standard Deviation (SD) (n = 3). (g) Calcein-AM-loaded THP-1 cells (10<sup>6</sup> cells/ml) were added to HUVECs and then incubated for 3 h. The unbound cells were washed off, and attached fluorescent monocytes were visualized using an optical microscope. (Left panel) The fluorescence intensity of Calcein-AM was quantified using ImageJ software. Data are presented as mean  $\pm$  SD (n = 3). P value was determined by unpaired Student's *t*-test. (h) HUVECs were treated with 5  $\mu$ M PCB29-pQ for 1, 3, or 6 h. IkB $\alpha$  and p-p65 levels were analyzed by western blotting analysis.  $\beta$ -Actin was used as an internal loading control. (i-j) IkB $\alpha$  and p-p65 levels were quantified by ImageJ software. Data are presented as mean  $\pm$  SD (n = 3). (k) HUVECs were pretreated with 5  $\mu$ M p65 inhibitor PDTC for 1 h, followed with 5  $\mu$ M PCB29-pQ exposure for 6 h. IL-1 $\beta$ , IL-6, and TNF- $\alpha$  in cell lysates were analyzed by western blotting.  $\beta$ -Actin was used as an internal loading control. (I-n) IL-1 $\beta$ , IL-6, and TNF- $\alpha$  protein expression levels were quantified by ImageJ software. Data were presented as mean  $\pm$  SD (n = 3). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. The exact mean and SD values are presented in Table S30.



Figure S18. Adhesion molecules, pro-inflammatory cytokines and p65 protein expression levels in HUVEC exposed to PCB29-pQ with *CAV1* siRNA. HUVECs were transfected with NC siRNA or 25 nM *CAV1* siRNA for 48 h and then treated with 5 µM PCB29-pQ for 6 h. (a) IL-1β, IL-6, TNF- $\alpha$ , ICAM-1 and VCAM-1 expressions were analyzed by western blotting. β-Actin was used as an internal loading control. (b-f) IL-1β, IL-6, TNF- $\alpha$ , ICAM-1 and VCAM-1 expression levels were quantified by ImageJ software. Data were presented as mean ± Standard Deviation (SD) (n = 3). (g) IκB $\alpha$ , p-p65, and p65 expressions were analyzed by western blotting. β-Actin was used as an internal

loading control. (**h-i**) I $\kappa$ B $\alpha$ , p-p65, and p65 expression levels were quantified by ImageJ software. Data were presented as mean  $\pm$  SD (n = 3). (**j**) (Left panel) Immunofluorescence analysis of p65. Green staining represents the location of p65. Nuclei shown in blue were stained with DAPI. Scale bar = 10  $\mu$ m. (Right panel) Co-localization of p-CAV1 and DAPI was analyzed by Pearson's correlation coefficient. Data were presented as mean  $\pm$  SD (n = 3). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. The exact mean and SD values are presented in Table S31.



Figure S19. Inflammatory factors and ROS levels in HUVEC exposed to PCB29-pQ. HUVECs were pretreated with 40  $\mu$ M VC, 20  $\mu$ M VE or 5 mM NAC for 1 h, followed with 5  $\mu$ M PCB29-pQ exposure for 6 h. (n = 3). (a) p-p65, p65, IL-1 $\beta$ , IL-6, TNF $\alpha$ , ICAM-1 and VCAM-1 in cell lysates were analyzed by western blotting. (b-g) p-p65, p65, IL-1 $\beta$ , IL-6, TNF $\alpha$ , ICAM-1 and VCAM-1 expression levels were quantified by ImageJ software. Data were presented as mean  $\pm$  Standard Deviation (SD) (n = 3). (h) ROS levels were detected by DCFH-DA (10  $\mu$ M) probe. Data were presented as mean  $\pm$  SD (n = 3). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. In all graphs, data is normalized to the control cell groups. The exact mean and SD values are presented in Table S32.



Figure S20. p-CAV1 and CAV1 levels in HUVEC exposed to PCB29-pQ and antioxidants. HUVECs were pretreated with 40  $\mu$ M VC, 20  $\mu$ M VE or 5 mM NAC for 1 h, followed by 5  $\mu$ M PCB29-pQ exposure for 6 h. HUVECs were pretreated with (a) 40  $\mu$ M VC, 20  $\mu$ M VE or 5 mM NAC, (b) 200 U/ml PEG-SOD, 500 U/ml PEG-CAT and 5 mM GSH-MEE for 1 h, followed with 5  $\mu$ M PCB29-pQ exposure for 6 h. (Left panel) p-CAV1 and CAV1 in cell lysates were analyzed by western blotting.  $\beta$ -Actin was used as an internal loading control. (Right panel) p-CAV1 and CAV1 expression levels were quantified by ImageJ software. Data were presented as mean  $\pm$  Standard Deviation (SD) (n = 3). P values were determined by one-way ANOVA, followed by Tukey's *post hoc* test. All data is graphed relative to the control cell groups. The exact mean and SD values are presented in Table S33.

| C7-AS  | 1        | ×      | ×     | ×     |      | ×    | ×      | - 0.8 |
|--------|----------|--------|-------|-------|------|------|--------|-------|
| 0.34 m | niR-7-5p |        |       | ×     | ×    | ×    |        | - 0.6 |
| .05    | -0.58    | TGF-β2 | ×     | ×     | ×    | ×    | ×      | - 0.4 |
| .18    | -0.24    | 0.13   | PPME1 | ×     | ×    | ×    | ×      | - 0.0 |
| .15    | 0.07     | 0.17   | 0.04  | IL-1β |      |      |        | 0.(   |
| 0.33   | 0.2      | -0.05  | -0.04 | 0.24  | IL-6 |      |        | 0.4   |
| 0      | 0.14     | 0.11   | 0.05  | 0.44  | 0.31 | TNFα |        | 0.0   |
| 0.16   | 0.33     | -0.11  | 0.06  | 0.26  | 0.29 | 0.41 | hs-CRP | 0.8   |

**Figure S21. Heatmap showing correlation scores between factors.** Plasma RNA was extracted by TRNzol universal reagent. CHD group (n = 77) contains patients with > 50% coronary artery stenosis, and the control group (n = 50) contains subjects with < 50% coronary artery stenosis. RT-qPCR analysis of *HDAC7-AS1*, *MIR-7-5p*, *TGF-β2*, *PPME1*, *IL-1β*, *IL6* and *TNFα* expressions. Spearman's rank correlation coefficient was used to assess the correlation between the two indicated factors. P value of < 0.05 was considered significant. The size and color of circle represents the correlation between two factors. X's mean no significance. Summary data can be found in Table S11.

| Producers                            | Antibodies (Item No.)  | Dilution                  |  |
|--------------------------------------|--|---------------------------|--|
|                                      | Goat anti-rabbit IgG (H+L), oralite<br>488 conjugate (# SA00013-2)     | 1:500 (IF)                |  |
|                                      | Mouse TNF-α monoclonal antibody (#<br>60291-1-lg)                      | 1:1000 (WB)               |  |
| Proteintech Group, Inc.              | Rabbit ICAM-1 monoclonal antibody<br>(# 60299-1-lg)                    | 1:1000 (WB)<br>1:500 (IF) |  |
| (Wuhan, China)                       | Rabbit VCAM-1 monoclonal antibody<br>(# 66294-1-lg)                    | 1:1000 (WB)<br>1:500 (IF) |  |
|                                      | Rabbit CAV1 polyclonal primary<br>antibody (#66067-1-lg)               | 1:1000 (WB)               |  |
|                                      | Rabbit Ago2 polyclonal antibody (#<br>10686-1-AP)                      | 1:500 (RIP)               |  |
|                                      | HRP-conjugated Goat Anti-Rabbit<br>IgG secondary antibody (# D11058)   | 1:2000 (WB)               |  |
| Sangon Biotech Co. Ltd.              | HRP-conjugated Rabbit Anti-mouse<br>IgG secondary antibody (# D110098) | 1:2000 (WB)               |  |
| (Wuhan, China)                       | Rabbit β-actin polyclonal antibody (#<br>D110001)                      | 1:5000 (WB)               |  |
|                                      | Rabbit TGF-β2 antibody (# D262351)                                     | 1:1000 (WB)<br>1:500 (IF) |  |
| Wanlei Co. Ltd.<br>(Shenyang, China) | Rabbit IL-6 polyclonal antibody (#<br>WL02841)                         | 1:1000 (WB)               |  |

# Table S1. Sources of antibodies using in this study.
|   | Rabbit IL-1β polyclonal antibody (#<br>WL00891)  | 1:1000 (WB)               |  |
|---|--|---------------------------|--|
|   | Rabbit IκBα antibody (# WL00148)                 | 1:1000 (WB)               |  |
| Bioss Biotech Co                              | Rabbit p65 antibody (# bs-20159R)                | 1:1000 (WB)               |  |
| Ltd. (Beijing, China)                         | Rabbit p-p65 (pSer536) antibody (# bs-<br>0982R) | 1:1000 (WB)<br>1:500 (IF) |  |
| Servicebio (Wuhan,<br>China)                  | Rabbit vWF polyclonal antibody (#<br>GB11020)    | 1:500 (IF)                |  |
|   | CD31 polyclonal antibody (# GB11063-<br>2)       | 1:500 (IF)                |  |
| Cell Signaling<br>Technology (Boston,<br>USA) | Rabbit p-CAV1 (Tyr14) antibody (#<br>3251)       | 1:1000 (WB)<br>1:500 (IF) |  |
| CUSABIO (Wuhan,<br>China)                     | Rabbit PPME1 antibody (# CSB-<br>PA018501LA01HU) | 1:1000 (WB)<br>1:500 (IF) |  |
| Abcam (Cambridge,<br>MA, USA)                 | Rabbit CD68 antibody (#ab283654)                 | 1:500 (IHC)               |  |

# Table S2. PCR primer sequences for ApoE and CAV1 mice genotype.

|      | Primer Type       | 5' - 3'                |
|------|-------------------|------------------------|
| АроЕ | Common            | GCCTAGCCGAGGGAGAGCCG   |
|      | Wild type Forward | TGTGACTTGGGAGCTCTGCAGC |
|      | Mutant Forward    | GCCGCCCCGACTGCATCT     |
| CAV1 | Common            | CTTGAGTTCTGTTAGCCCAG   |
|      | Wild type Forward | GTGTATGACGCGCACACCAAG  |
|      | Mutant Forward    | CTAGTGAGACGTGCTACTTCC  |

Sequences are mouse unless otherwise specified.

| Step  | 1               | 2                 |          |        | 3               |
|-------|-----------------|-------------------|----------|--------|-----------------|
|       | CYCLE (1 cycle) | CYCLE (35 cycles) |          |        | CYCLE (1 cycle) |
| Temp. | 94 °C           | 94 °C             | 55-65 °C | 72 °C  | 72 °C           |
| Time  | 2 min           | 30 sec            | 30 sec   | 30 sec | 2 min           |

Table S3. PCR program for *ApoE* and *CAV1* mice genotype.

# Table S4. RT-qPCR primer sequences for lncRNAs and mRNAs.

| Gene                   | Sequence (5' to 3')      |
|------------------------|--------------------------|
| <i>LINC01547-</i> F    | AGGCCAAGAGACAACAGCGATTAC |
| <i>LINC01547-</i> R    | GCCAAGTGTGGACTCAGAGCTTC  |
| SLCO4A1-AS1- F         | CTGTCAGCCGCCTTCTTGCC     |
| <i>SLCO4A1-AS1-</i> R  | GCGAGGAGCATGTAGAACCTGTC  |
| NRSN2-AS1-F            | AATGAGCGGAGATTGTGCCACTG  |
| NRSN2-AS1-R            | CGACAGTACCTACAACAGCCACAC |
| FGD5-AS1-F             | GTTGGACCAAGAGGCAGCTC     |
| FGD5-AS1-R             | AGACCACCGAACAGTGACCA     |
| MCM3AP-AS1-F           | AGAAGCTCCTCGCATCAGATCCTC |
| MCM3AP-AS1-R           | CACATGCACCGTAACTGGAAGAGG |
| <i>TUG1-</i> F         | TAGCAGTTCCCCAATCCTTG     |
| <i>TUG1-</i> R         | CACAAATTCCCATCATTCCC     |
| <i>RP3-416H24.1-</i> F | GGCTTCATTGGCACCACCTACTC  |
| <i>RP3-416H24.1-</i> R | GCAGAGCCAGAACTGGAACATAGG |
| <i>АС002550.5-</i> F   | CTGTCAGCCGCCTTCTTGCC     |
| <i>AC002550.5-</i> R   | GCGAGGAGCATGTAGAACCTGTC  |
| HDAC7-AS1-F            | GCCTTCTAGCCACAAGCACACTC  |
| HDAC7-AS1-R            | GCGAAGCGGAAGCCTCTGTTC    |
| <i>SLFNL1-AS-</i> F    | AACTGAAGCACAGAGGCATAGCAC |
| <i>SLFNL1-AS-</i> R    | CCAGGAGGCAGAGGACAGACC    |
| HDAC7-F                | TGCACCACCACCTCTTCCTAGC   |
| HDAC7-R                | ACTTCGCTTGCTCTTGTCCTTGTG |
| <i>TGF-β2-</i> F       | GTGCCTGAACAACGGATTGA     |
| <i>TGF-β2</i> -R       | AAGGAGAGCCATTCGCCTTC     |
| (mouse) TGF-β2-F       | TCGACATGGATCAGTTTATGCG   |
| (mouse) TGF-β2-R       | CCCTGGTACTGTTGTAGATGGA   |
| PPME1-F                | CAGTCCTGCTCCTTCTGCAT     |
| PPME1-R                | TTTCACCATGACTTCGCAGA     |
| (mouse) PPME1-F        | AGTCAGAGCGGAGCCAAGAT     |
| (mouse) PPME1-R        | TCGAAAAGTATCCTTGCCAGTTT  |
| <i>IL-1β</i> -F        | TTCGACACATGGGATAACGAGG   |
| <i>IL-1β</i> -R        | TTTTTGCTGTGAGTCCCGGAG    |
| (mouse) IL-1β-F        | GCAACTGTTCCTGAACTCAACT   |
| (mouse) IL-1β-R        | ATCTTTTGGGGTCCGTCAACT    |
| <i>IL-6-</i> F         | ACTCACCTCTTCAGAACGAATTG  |
| <i>IL-6-</i> R         | CCATCTTTGGAAGGTTCAGGTTG  |
| (mouse) <i>IL-6-</i> F | TAGTCCTTCCTACCCCAATTTCC  |

| (mouse) <i>IL-6</i> -R    | TTGGTCCTTAGCCACTCCTTC  |
|---------------------------|------------------------|
| <i>TNF-а-</i> F           | GAGGCCAAGCCCTGGTATG    |
| <i>TNF-</i> а-R           | CGGGCCGATTGATCTCAGC    |
| (mouse) <i>TNF-α-</i> F   | CAGGCGGTGCCTAGTTCTC    |
| (mouse) <i>TNF-α-</i> R   | CGATCACCCCGAAGTTCAGTAG |
| <i>β-actin-</i> F         | CATGTACGTTGCTATCCAGGC  |
| <i>β-actin-</i> R         | CTCCTTAATGTCACGCACGAT  |
| (mouse) <i>β-actin-</i> F | GGCTGTATTCCCCTCCATCG   |
| (mouse) <i>β-actin-</i> R | CCAGTTGGTAACAATGCCATGT |
| <i>18s</i> -F             | GTAACCCGTTGAACCCCATT   |
| 18s-R                     | CCATCCAATCGGTAGTAGCG   |

Sequences are human unless otherwise specified.

## Table S5. RT-qPCR primer sequences for miRNAs.

| miRNA            | Forward (5' to 3')      |
|------------------|-------------------------|
| <i>MIR-7-5p</i>  | UGGAAGACUAGUGAUUUUGUUGU |
| <i>MIR-24-3p</i> | UGGCUCAGUUCAGCAGGAACAG  |
| <i>MIR-9-5p</i>  | UCUUUGGUUAUCUAGCUGUAUGA |

Sequences are human unless otherwise specified.

## Table S6. RT-qPCR program.

## RT-qPCR program (For mRNA and lncRNA)

| Step  | 1                | 2                    |                        |        |        | 3           |        |
|-------|------------------|----------------------|------------------------|--------|--------|-------------|--------|
|       | Hot-Start<br>DNA | PCR                  |                        |        | М      | elt Curve   |        |
|       | Polymerase       |                      |                        |        |        |             |        |
|       | Activation       |                      |                        |        |        |             |        |
|       | HOLD             | CYCI                 | LE (40 cycles          | 5)     | CYC    | LE (1 cycle | e)     |
|       |                  | Denature             | Denature Anneal Extend |        |        |             |        |
| Temp. | 95 ℃             | 95 °C 50-60 °C 72 °C |                        | 95 °С  | 60 °C  | 95 ℃        |        |
| Time  | 10 min           | 15 sec               | 30 sec                 | 30 sec | 15 sec | 60 sec      | 15 sec |

## RT-qPCR program (For miRNA)

| Step  | 1  | 2                             |        |          | 3             |               |       | 4        |       |
|-------|--|-------------------------------|--------|----------|---------------|---------------|-------|----------|-------|
|       | Hot-Start<br>DNA<br>Polymerase<br>Activation | Enrich low-abundance<br>miRNA |        |          | PC            | R             | М     | lelt Cur | ve    |
|       | HOLD   | CYCLE (5 cycles)              |        |          | CYCLE<br>cycl | (40-45<br>es) | CYC   | CLE (1 c | ycle) |
|       |  |                               |        | Denature | Anneal,       |               |       |          |       |
|       |  |                               | -      |          |               | Extend        |       | -        | -     |
| Temp. | 95 °С  | 94 °C                         | 63-    | 72 °C    | 94 °C         | 60 °C         | 94 °C | 60 °C    | 95 °С |
|       |  |                               | 65 °C  |          |               |               |       |          |       |
| Time  | 15 min                                       | 20 sec                        | 30 sec | 34       | 20            | 34            | 15    | 60       | 15    |
|       |  |                               |        | sec      | sec           | sec           | sec   | sec      | sec   |

## Table S7. siRNAs, mimic, and inhibitor target sequences.

| Producers                   | Item No.  | Gene                      | Target sequences             |
|-----------------------------|---|---------------------------|------------------------------|
|                             | # 101001  | $TCE \rho_2 $ DNA         | 5'-GCGGCCUAUUCGUUUAGAA-3'    |
|                             | # A01001  | <i>I GF-p2</i> SIKINA     | 3'-UUCUAAAGCAAUAGGCCGC-5'    |
|                             | # 101001  | <i>DDME1</i> «:DNIA       | 5'-GAAUGAAACUGGCAAGGAU-3'    |
|                             | # A01001  | PPME1 SIKINA              | 3'-AUCCUUGCCAGUUUCAUUC-5'    |
| Shanghai                    | # 101001  | HDAC7-AS1                 | 5'-GATGTTGAGTTGAAGTTGA-3'    |
| Gene<br>Pharma Co.,<br>Ltd. | # A01001  | siRNA                     | 3'-TCAACTTCAACTCAACATC-5'    |
|                             | na Co.,<br>td. # A01001 CAVI siRN<br>nghai,<br>ina) # B01001 MIR-7-5p m | CAVI siRNA                | 5'-CGACGACGUGGUCAACAUU-3'    |
|                             |   |                           | 3'-AAUCUUGACCACGUCGUCG-5'    |
| (Shanghai,                  |   |                           | 5'-UGGAAGACUAGUGAUUUUGUUG-   |
| China)                      |   | MIR-7-5p mimic            | 3'                           |
|                             |   |                           | 3'-CAACAAAAUCACUAGUCUUCCA-5' |
|                             |   |                           | 5'-                          |
|                             | # B03001  | <i>MIR-7-5p</i> inhibitor | AACAACAAAAUCACUAGUCUUCCA-    |
|                             |   |                           | 3'                           |

Sequences are human unless otherwise specified.

### Table S8. Patient characteristics and clinical results.

|                             | Control (n=50)    | CHD (n=77)        | Р     |
|-----------------------------|-------------------|-------------------|-------|
| Male, n (%)                 | 27 (54.0)         | 59 (76.6)         | 0.011 |
| Age, years, mean $\pm$ SD   | $62.18 \pm 10.78$ | $64.08 \pm 7.769$ | 0.252 |
| Cardiovascular risk factors |                   |                   |       |
| Diabetes, n (%)             | 17 (34.0)         | 32 (41.6)         | 0.457 |
| Hypertension, n (%)         | 33 (66.0)         | 56 (74.7)         | 0.319 |
| Hyperlipidemia, n (%)       | 41 (82.0)         | 68 (88.3)         | 0.435 |

| Current smoke, n (%)                               | 20 (40.0)         | 43 (55.8)         | 0.103 |
|--|-------------------|-------------------|-------|
| Body mass index, kg/m <sup>2</sup> , mean $\pm$ SD | $25.94 \pm 3.433$ | $25.88 \pm 3.082$ | 0.917 |
| Blood biochemical analysis                         |                   |                   |       |
| TG (mM), mean $\pm$ SD                             | $4.017\pm0.814$   | $4.112 \pm 1.113$ | 0.610 |
| TC (mM), mean $\pm$ SD                             | $1.900\pm1.608$   | $1.910 \pm 1.483$ | 0.971 |
| LDL-C (mM), mean ± SD                              | $2.207\pm0.532$   | $2.430\pm0.994$   | 0.151 |
| HDL-C (mM), mean ± SD                              | $1.065 \pm 0.423$ | $1.019 \pm 0.374$ | 0.523 |
| FBG (mM), mean ± SD                                | $5.644 \pm 1.325$ | $6.298 \pm 2.154$ | 0.061 |
| HbA1c (%), mean $\pm$ SD                           | $6.312 \pm 0.157$ | $6.626 \pm 1.493$ | 0.231 |
| hs-CRP (mg/L), mean $\pm$ SD                       | $2.622 \pm 3.526$ | $8.418 \pm 16.30$ | 0.036 |
| $ALT(U/L)$ , mean $\pm$ SD                         | $25.55 \pm 16.97$ | $23.46 \pm 16.64$ | 0.498 |
| $AST(U/L)$ , mean $\pm$ SD                         | $24.92 \pm 11.78$ | $29.03\pm44.01$   | 0.524 |
| Albumin (g/L), mean $\pm$ SD                       | $39.37\pm2.737$   | $38.58 \pm 2.640$ | 0.112 |
| T-BIL ( $\mu$ M), mean $\pm$ SD                    | $15.31 \pm 6.063$ | $14.05 \pm 6.475$ | 0.279 |
| Urea (mM), mean $\pm$ SD                           | $4.954 \pm 1.323$ | $5.498 \pm 1.880$ | 0.081 |
| Creatinine ( $\mu$ M), mean $\pm$ SD               | $64.91 \pm 13.32$ | $70.38 \pm 17.20$ | 0.061 |
| Uric acid ( $\mu$ M), mean $\pm$ SD                | $361.3 \pm 88.16$ | $348.4 \pm 95.36$ | 0.452 |

Data are presented as Mean  $\pm$  SD or n (%). CHD, coronary heart disease; TC, total cholesterol; TG, total triglyceride; LDL-C, low-density lipoprotein cholesterol; HDL-C, high-density lipoprotein cholesterol; FBG, fasting blood glucose; hs-CRP, high-sensitivity C-reactive protein; ALT, alanine aminotransferase; AST, aspartate aminotransferase; T-BIL, total bilirubin. To compare the differences between two groups with normally distributed variables, Student's *t* test was used. To test for differences between two groups with skewed variables, the nonparametric Mann-Whitney *U* test was used. In addition, categorical variables were showed as number with percent and analyzed by the  $\chi^2$  test.

Table S9. MIR-7-5p binding sites for HDAC7-AS1 as identified by Mireap, miRanda, and

| miRNA                | Target<br>Gene | Score | Energy<br>(kcal/mol) | 3'-miRNA-5'-alignment-5'-utr-3'  |
|----------------------|----------------|-------|----------------------|--|
| HSA-<br>MIR-7-<br>5p | HDAC7-<br>AS1  | 119   | -13.6                | <pre>3' uuGUUG-UUUUAGUGAUCAGAAGGu 5'</pre>   |
| HSA-<br>MIR-7-<br>5p | HDAC7-<br>AS1  | 117   | -15.34               | <pre>3' uuGUUGUUUUAGUGA-UCAGAaggu 5'<br/>      :    :       <br/>5' acCACCAAGATCCTTCAGTCTgage 3'</pre>               |
| HSA-<br>MIR-7-<br>5p | HDAC7-<br>AS1  | 115   | -16.82               | <ul> <li>3' uuguUGU- UUUAGUGAUCAGAAGGu 5'</li> <li>     :     : </li> <li>5' cagtACATCAGCCACCAGTCTCTCa 3'</li> </ul> |
| HSA-<br>MIR-7-<br>5p | HDAC7-<br>AS1  | 114   | -7.86                | 3' uuguuguuuUAGUGAU-CAGAAGGu 5'<br>      :       <br>5' ctgttctctACCCCTGTGGCTTCCc 3'                                 |
| HSA-<br>MIR-7-<br>5p | HDAC7-<br>AS1  | 111   | -11.1                | <pre>3' uuguuGUUUUAGUGAUCAGAAGGu 5'<br/>  :::           :<br/>5' cacccCAGGGGCCCCAGCCTTCTa 3'</pre>                   |

TargetScan.

Mireap (v2.0) (<u>http://sourceforge.net/projects/mireap/</u>);

miRanda (v3.3a) (<u>http://cbio.mskcc.org/microrna\_data/miRanda-aug2010.tar.gz</u>);

TargetScan (v7.0) (http://www.targetscan.org/vert\_71/).

| Table S10. | . Negatively | correlated | miRNA-gene | pairs: MIR- | -7-5p and | l its target genes. |
|------------|--------------|------------|------------|-------------|-----------|---------------------|
|            |              |            |            |             |           |                     |

| miRNA    | Gene    | P value     | asso_sign (miRNA- |
|----------|---------|-------------|-------------------|
|          |         |             | Target gene)      |
|          | CTSK    | 0.000142241 | UP-DOWN           |
|          | CSMD3   | 1.89E-06    | UP-DOWN           |
|          | HSPBAP1 | 9.02E-05    | UP-DOWN           |
|          | YPEL2   | 0.000112022 | UP-DOWN           |
|          | GLS     | 6.58E-20    | UP-DOWN           |
|          | FAM110C | 0.000660921 | UP-DOWN           |
|          | PPME1   | 1.12E-53    | UP-DOWN           |
| MIR-7-5p | IFITM10 | 0.009667697 | UP-DOWN           |
|          | ABL2    | 7.28E-39    | UP-DOWN           |
|          | TGFB2   | 6.75E-12    | UP-DOWN           |
|          | SH2D5   | 2.27E-65    | UP-DOWN           |

| PTCH1   | 3.89E-09    | UP-DOWN |
|---------|-------------|---------|
| SLC5A3  | 2.57E-119   | UP-DOWN |
| KLHDC7A | 0.000498548 | UP-DOWN |
| SLC23A3 | 0.00724907  | UP-DOWN |
| TRIML2  | 1.47E-81    | UP-DOWN |

The R package edgeR (<u>http://www.bioconductor.org/packages/release/bioc/html/edgeR.html</u>) was used to identify differentially expressed transcripts across the samples or groups as evaluated using RNA-seq. mRNA and miRNA with a fold change  $\geq 2$  and P < 0.05 were considered as significant differentially expressed genes (DEGs).

Table S11. Results of correlation analysis between two indicated factors.

|           |                 | r        | Р      |
|-----------|-----------------|----------|--------|
| HDAC7-AS1 | <i>MIR-7-5p</i> | -0.3389  | 0.0026 |
|           | TGF-β2          | 0.04843  | 0.6758 |
|           | PPME1           | 0.1826   | 0.1119 |
|           | IL-1β           | 0.1533   | 0.1833 |
|           | IL-6            | -0.3336  | 0.003  |
|           | TNF-α           | 0.002393 | 0.9835 |
|           | hs-CRP          | -0.1602  | 0.1953 |
| MIR-7-5p  | TGF-β2          | -0.5835  | 0.001  |
|           | PPME1           | -0.24    | 0.0353 |
|           | <i>IL-1β</i>    | 0.07496  | 0.5171 |
|           | IL-6            | 0.201    | 0.0796 |
|           | TNF-α           | 0.1395   | 0.2264 |
|           | hs-CRP          | 0.3304   | 0.0063 |
| TGF-β2    | PPME1           | 0.1337   | 0.2463 |
|           | <i>IL-1β</i>    | 0.172    | 0.1346 |
|           | IL-6            | -0.04524 | 0.696  |
|           | TNF-α           | 0.113    | 0.3277 |
|           | hs-CRP          | -0.1098  | 0.3763 |
| PPME1     | <i>IL-1β</i>    | 0.03938  | 0.7338 |
|           | IL-6            | -0.04292 | 0.7109 |
|           | TNF-α           | 0.04688  | 0.6856 |
|           | hs-CRP          | 0.06204  | 0.618  |
| IL-1β     | IL-6            | 0.2419   | 0.0341 |
|           | TNF-α           | 0.4351   | 0.0001 |
|           | hs-CRP          | 0.2608   | 0.0331 |
| IL-6      | TNF-α           | 0.313    | 0.0056 |
| Ī         | hs-CRP          | 0.2887   | 0.0178 |
| TNFa      | hs-CRP          | 0.4069   | 0.0006 |

Spearman's rank correlation coefficient was used to assess the correlation between two indicated mRNAs.

| -              |       |             | Γ_    |                                 | ·            | T     |
|----------------|-------|-------------|-------|---------------------------------|--------------|-------|
| Variables      | OR    | 95% CI      | Р     | Adjusted OR <sup><i>a</i></sup> | 95% CI       | Р     |
| Gender (male)  | 2.647 | 1.233-5.683 | 0.013 | 2.266                           | 0.343-14.961 | 0.396 |
| Age            | 1.028 | 0.987-1.027 | 0.180 |                                 |              |       |
| Diabetes       | 2.020 | 0.981-4.159 | 0.056 | 1.397                           | 0.266-7.325  | 0.692 |
| Hypertension   | 1.527 | 0.708-3.793 | 0.280 |                                 |              |       |
| Hyperlipidemia | 1.263 | 0.606-2.632 | 0.533 |                                 |              |       |
| Current smoker | 2.020 | 0.981-4.159 | 0.056 | 0.897                           | 0.195-4.134  | 0.890 |
| BMI            | 1.009 | 0.947-1.076 | 0.781 |                                 |              |       |
| TG             | 1.016 | 0.801-1.288 | 0.898 |                                 |              |       |
| TC             | 1.163 | 0.810-1.668 | 0.413 |                                 |              |       |
| LDL-C          | 1.557 | 0.922-2.628 | 0.098 | 2.090                           | 0.735-5.944  | 0.167 |
| HDL-C          | 0.779 | 0.314-1.937 | 0.592 |                                 |              |       |
| FBG            | 1.235 | 0.984-1.550 | 0.068 | 1.277                           | 0.834-1.955  | 0.261 |
| HbA1c          | 1.063 | 0.917-1.213 | 0.415 |                                 |              |       |
| hs-CRP         | 1.211 | 1.085-1.351 | 0.011 | 1.169                           | 0.999-1.368  | 0.052 |
| ALT            | 0.993 | 0.972-1.014 | 0.506 |                                 |              |       |
| AST            | 1.005 | 0.991-1.019 | 0.517 |                                 |              |       |
| Albumin        | 0.885 | 0.770-1.016 | 0.083 | 0.869                           | 0.679-1.112  | 0.264 |
| T-BIL          | 0.973 | 0.919-1.030 | 0341  |                                 |              |       |
| Urea           | 1.286 | 0.982-1.686 | 0.068 | 1.313                           | 0.756-2.292  | 0.331 |
| Creatinine     | 1.027 | 1.002-1.052 | 0.035 | 1.019                           | 0.957-1.085  | 0.549 |
| Uric acid      | 1.000 | 0.996-1.003 | 0.912 |                                 |              |       |
| HDAC7-AS1      | 0.238 | 0.125-0.451 | 0.032 | 0.430                           | 0.157-1.177  | 0.101 |
| MIR-7-5p       | 1.551 | 1.223-1.966 | 0.003 | 2.168                           | 1.060-4.434  | 0.034 |
| TGF-β2         | 0.553 | 0.357-0.857 | 0.008 | 0.651                           | 0.265-1.599  | 0.349 |
| PPME1          | 0.644 | 0.438-0.947 | 0.025 | 0.379                           | 0.193-0.743  | 0.005 |
| ΙL-1β          | 2.722 | 1.677-4.416 | 0.001 | 1.599                           | 0.928-1.756  | 0.091 |
| ΤΝFα           | 1.179 | 0.961-1.446 | 0.114 |                                 |              |       |
| IL6            | 1.223 | 1.006-1.487 | 0.043 | 1.095                           | 0.771-1.555  | 0.613 |

Table S12. Univariable and multivariable logistic regression analysis for determining the

independent risk factors of CHD.

OR, Odds ratio; CI, confidence interval; CHD, coronary heart disease; TC, total cholesterol; TG, total triglyceride; LDL-C, low-density lipoprotein cholesterol; HDL-C, high-density lipoprotein cholesterol;

FBG, fasting blood glucose, fasting blood glucose; hs-CRP, high-sensitivity C-reactive protein; ALT,

alanine aminotransferase; AST, aspartate aminotransferase.

<sup>*a*</sup> Corresponds to adjustment for gender, diabetes, current smoker, LDL-C, FBG, albumin, hs-CRP, urea, creatinine, Uric acid, *HDAC7-AS1*, *MIR-7-5p*, *TGF-β2*, *PPME1*, *IL-1β* and *IL-6*.

Table S13. High-throughput lncRNA and miRNA sequence analysis and relevant qPCR results

|        |                  | (         | Control |         | PCI       | 329-pQ   |         |
|--------|------------------|-----------|---------|---------|-----------|----------|---------|
|        |                  |           | Mean    | SD      |           | Mean     | SD      |
|        |                  | Cq values | (normal | (normal | Cq values | (normali | (normal |
|        |                  |           | ized)   | ized)   |           | zed)     | ized)   |
|        |                  | 21.70     | -       |         | 19.71     | -        |         |
|        | HDAC7-<br>ASI    | 21.68     | 1.00    | 0.02    | 19.88     | 3.57     | 0.61    |
|        |                  | 21.64     |         |         | 19.40     |          |         |
|        |                  | 21.74     |         |         | 22.53     |          |         |
|        | RP3-<br>416H24 1 | 21.60     | 1.00    | 0.05    | 22.62     | 0.52     | 0.08    |
|        | 7101127.1        | 21.70     |         |         | 22.21     |          |         |
|        |                  | 21.57     |         |         | 19.81     |          |         |
| Figure | LINC01547        | 21.48     | 1.00    | 0.07    | 21.28     | 2.20     | 1.00    |
| lc     |                  | 21.69     |         |         | 20.06     |          |         |
|        |                  | 21.81     |         | 0.04    | 21.74     | 0.93     |         |
|        | TUG1             | 21.77     | 1.00    |         | 21.73     |          | 0.06    |
|        |                  | 21.70     |         |         | 21.57     |          |         |
|        |                  | 21.86     |         |         | 21.59     |          |         |
|        | MCM3AP-<br>ASI   | 21.83     | 1.00    | 0.01    | 21.72     | 0.97     | 0.08    |
|        |                  | 21.85     |         |         | 21.83     |          |         |
|        |                  | 21.86     |         |         | 24.44     |          |         |
|        | FGD5-AS1         | 21.92     | 1.00    | 0.02    | 24.34     | 0.17     | 0.02    |
|        |                  | 21.87     |         |         | 24.09     |          |         |

from HUVECs treated with 5 µM PCB29-pQ.

|           | H         | Bio-NC  |         | Bio-M     | 1IR-7-5p | IR-7-5pMeanSD(normali(normalzed)ized) |  |  |
|-----------|-----------|---------|---------|-----------|----------|---------------------------------------|--|--|
|           |           | Mean    | SD      |           | Mean     | SD                                    |  |  |
| Figure 1d | Cq values | (normal | (normal | Cq values | (normali | (normal                               |  |  |
|           |           | ized)   | ized)   |           | zed)     | ized)                                 |  |  |
|           | 21.89     |         |         | 17.98     | 15.31    | 3.72                                  |  |  |
|           | 21.89     | 1.00    | 0.01    | 17.97     |          |                                       |  |  |
|           | 21.87     |         |         | 17.40     |          |                                       |  |  |

Mean and SD values are the ones presented in Figure 1c-d. Data reported relative to control or Bio-

NC.

# Table S14. The expression of *HDAC7-AS1*, *MIR-7-5p* and *TGF-β2/PPME1* in HUVECs exposed

# to 5 µM PCB29-pQ.

|           |                 | Control   |                              |                        | PCB29-pQ  |                          |                        |
|-----------|-----------------|-----------|------------------------------|------------------------|-----------|--------------------------|------------------------|
|           |                 | Cq values | Mean<br>(norm<br>alized<br>) | SD<br>(normalize<br>d) | Cq values | Mean<br>(normal<br>ized) | SD<br>(normali<br>zed) |
|           |                 | 21.91     |                              |                        | 22.76     |                          |                        |
| Figure 2a | TGF <b>-</b> β2 | 21.93     | 1.00                         | 0.01                   | 23.02     | 0.48                     | 0.06                   |
|           |                 | 21.92     |                              |                        | 22.68     |                          |                        |
|           |                 | 22.17     |                              |                        | 22.46     |                          |                        |
|           | PPME1           | 22.09     | 1.00                         | 0.03                   | 22.36     | 0.71                     | 0.04                   |
|           |                 | 22.13     |                              |                        | 22.50     |                          |                        |

|                               |        | (                   | Control                      |                        |                     | PCB29-pQ                 |                        |  |
|-------------------------------|--------|---------------------|------------------------------|------------------------|---------------------|--------------------------|------------------------|--|
| Figure 2b<br>(Right<br>panel) |        | Grayscale<br>values | Mean<br>(norm<br>alized<br>) | SD<br>(normalize<br>d) | Grayscale<br>values | Mean<br>(normal<br>ized) | SD<br>(normali<br>zed) |  |
|                               | TGF-β2 | 49800.52            |                              | 0.30                   | 13613.11            | 0.23                     | 0.05                   |  |
|                               |        | 49779.34            | 1.00                         |                        | 13624.50            |                          |                        |  |
|                               |        | 49799.71            |                              |                        | 13598.56            |                          |                        |  |
|                               | PPME1  | 63958.41            | 1.00                         | 0.20                   | 43513.44            | 0.54                     | 0.09                   |  |

|  | 63928.43 |  | 43484.88 |  |
|--|----------|--|----------|--|
|  | 63944.37 |  | 43491.22 |  |

|           |        | N     | C mimic                      |                        | MIR   | R- <i>7-5p</i> mimic     |                        |
|-----------|--------|-------|------------------------------|------------------------|-------|--------------------------|------------------------|
|           |        | Cq    | Mean<br>(norm<br>alized<br>) | SD<br>(normalize<br>d) | Cq    | Mean<br>(normal<br>ized) | SD<br>(normali<br>zed) |
| Figure 2c | TGF-β2 | 25.54 | 1.00                         | 0.03                   | 26.88 | 0.49                     | 0.06                   |
|           |        | 25.45 |                              |                        | 27.13 |                          |                        |
|           |        | 25.53 |                              |                        | 27.20 |                          |                        |
|           | PPME1  | 25.69 | 1.00                         | 0.03                   | 26.90 | 0.68                     | 0.04                   |
|           |        | 25.77 |                              |                        | 26.73 |                          |                        |
|           |        | 25.70 |                              |                        | 26.79 |                          |                        |

|                     |        | N                   | C mimic                      |                        | <i>MIR-7-5p</i> mimic |                          |                        |
|---------------------|--------|---------------------|------------------------------|------------------------|-----------------------|--------------------------|------------------------|
|                     |        | Grayscale<br>values | Mean<br>(norm<br>alized<br>) | SD<br>(normalize<br>d) | Grayscale<br>values   | Mean<br>(normal<br>ized) | SD<br>(normali<br>zed) |
| Figure 2d<br>(Right |        | 60469.01            |                              |                        | 29146.83              |                          |                        |
|                     | TGF-β2 | 60473.45            | 1.00                         | 0.20                   | 29101.17              | 0.41                     | 0.08                   |
| panel)              |        | 60443.78            |                              |                        | 29128.81              |                          |                        |
|                     | PPME1  | 52334.05            |                              |                        | 52342.9               | 0.32                     | 0.06                   |
|                     |        | 52312.78            | 1.00                         | 0.17                   | 52253.98              |                          |                        |
|                     |        | 52288.09            |                              |                        | 52338.05              |                          |                        |

|           | NC mimic               | e + pEZ-N<br>NC | M61- | pEZ-M61-N<br>m         | NC + <i>MII</i><br>nimic | R-7-5p | <i>MIR-7-5p</i> 1<br>M61-HI | nimic + j<br>DAC7-AS | pEZ-<br>S1 |
|-----------|------------------------|-----------------|------|------------------------|--------------------------|--------|-----------------------------|----------------------|------------|
|           | luciferase<br>activity | Mean            | SD   | luciferase<br>activity | Mean                     | SD     | luciferase<br>activity      | Mean                 | SD         |
| Figure 2f | 29.48                  |                 |      | 25.60                  |                          |        | 30.35                       |                      |            |
|           | 29.86                  | 29.81           | 0.30 | 25.05                  | 25.34                    | 0.28   | 33.00                       | 31.46                | 1.38       |
|           | 30.08                  |                 |      | 25.37                  |                          |        | 31.02                       |                      |            |

|           | NC mimic               | e + pEZ-N<br>NC | <i>A</i> 61- | pEZ-M61-N<br>m         | NC + MIP | R-7-5p | <i>MIR-7-5p</i> 1<br>M61-HI | mimic + j<br>DAC7-AS | pEZ-<br>S1 |
|-----------|------------------------|-----------------|--------------|------------------------|----------|--------|-----------------------------|----------------------|------------|
|           | luciferase<br>activity | Mean            | SD           | luciferase<br>activity | Mean     | SD     | luciferase<br>activity      | Mean                 | SD         |
| Figure 2g | 8.11                   |                 |              | 5.26                   |          |        | 6.22                        |                      |            |
|           | 7.52                   | 7.81            | 0.30         | 5.36                   | 5.36     | 0.09   | 6.38                        | 6.35                 | 0.12       |
| 7.80      |                        |                 |              | 5.45                   |          |        | 6.45                        |                      |            |

Mean and SD values are the ones presented in Figure 2a, b, c, d and f. Expression is relative to control or NC mimic for data from 2a, b, c, and d. For Figure 2F, data is expressed as luciferase activity (a.u.)

Table S15. Endothelial injury and atherogenesis in *ApoE<sup>-/-</sup>* mice exposed to PCB29-pQ, and apoptotic rate in HUVECs exposed to PCB29-pQ.

|                           |                                 | Contro            | ol         |      | PCB29             | -pQ   |       |
|---------------------------|---------------------------------|-------------------|------------|------|-------------------|-------|-------|
|                           |                                 | Individual values | Mean       | SD   | Individual values | Mean  | SD    |
|                           |                                 | 13.01             |            |      | 54.48             |       |       |
| Figure 3a<br>panel: % pla | (Right<br>aue area)             | 15.08             | 14.89      | 1.80 | 48.96             | 45.47 | 11.17 |
|                           | 1 )                             | 16.59             |            |      | 32.98             |       |       |
| Figure 3h (R              | ight nanel·                     | 0.50              |            |      | 0.94              |       |       |
| Pearson's co              | orrelation                      | 0.15              | 0.33       | 0.18 | 0.99              | 0.98  | 0.03  |
| coeffic                   | ient)                           | 0.33              |            |      | 1.00              |       |       |
|                           | NC                              | 99.07             |            |      | 84.14             |       |       |
|                           | inhibitor<br>+ NC               | 109.28            | 100.0<br>0 | 5.85 | 75.19             | 81.21 | 5.21  |
|                           | siRNA                           | 91.65             |            |      | 84.30             |       |       |
|                           | NC                              | 115.81            |            |      | 95.23             |       |       |
| Figure 3d                 | MIR-7-                          | 105.25            | 111.1<br>7 | 5.40 | 92.29             | 97.03 | 5.85  |
| viability, %)             | inhibitor                       | 112.46            |            |      | 103.57            |       |       |
|                           | MIR-7-                          | 63.92             |            |      | 49.93             |       |       |
| 5p<br>inhibitor           | Sp<br>inhibitor<br>+ $TGF_{-}R$ | 57.62             | 60.10      | 3.36 | 55.44             | 50.27 | 5.01  |
|                           | 2  siRNA                        | 58.76             |            |      | 45.44             |       |       |

| MIR-7-<br>5p   | 64.3 | 3        |      | 45.79 |       |      |
|----------------|------|----------|------|-------|-------|------|
| inhibito +     | 54.7 | 74 57.59 | 5.86 | 52.60 | 48.14 | 3.87 |
| PPME.<br>siRNA | 53.7 | 70       |      | 46.03 |       |      |

Mean and SD values are the ones presented in Figure 3a, b and d.

Table S16. Measures of endothelial injury in cells and protein and mRNA expression of *TGF-β2* and *PPME1* in HUVECs transfected with

|           | NC min | nic + pEZ- | M61-NC  | NC mimic<br>I | NC mimic + pEZ-M61-NC +<br>PCB29-pQ |         |        | mic + pEZ $-ASI + PC$ | -M61-<br>B29-pQ | MIR-7-5p mimic + pEZ-<br>M61-HDAC7-AS1 + |         |          |  |
|-----------|--------|------------|---------|---------------|-------------------------------------|---------|--------|-----------------------|-----------------|--|---------|----------|--|
|           |        |            |         |               |                                     |         |        |                       |                 |  | PCB29-p | Q        |  |
|           | Cq     | Mean       | SD      | Cq            | Mean                                | SD      | Cq     | Mean                  | SD              | Cq                                       | Mean    | SD       |  |
|           | values | (normal    | (normal | values        | (normal                             | (normal | values | (normal               | (normal         | values                                   | (normal | (normali |  |
|           |        | ized)      | ized)   |               | ized)                               | ized)   |        | ized)                 | ized)           |  | ized)   | zed)     |  |
|           | 23.01  |            |         | 21.70         |                                     |         | 25.50  |                       |                 | 16.74                                    |         |          |  |
| Figure 4a | 23.09  | 1.00       | 0.03    | 21.67         | 2.11                                | 0.24    | 25.60  | 0.29                  | 0.02            | 16.46                                    | 102.90  | 11.57    |  |
| _         | 23.06  |            |         | 21.41         |                                     |         | 25.70  |                       |                 | 16.73                                    |         |          |  |
|           | 23.17  |            |         | 23.38         |                                     |         | 20.23  |                       |                 | 19.76                                    |         |          |  |
| Figure 4b | 22.92  | 1.00       | 0.09    | 23.38         | 0.60                                | 0.02    | 20.27  | 11.72                 | 0.29            | 19.71                                    | 12.25   | 0.44     |  |
|           | 23.06  |            |         | 23.45         |                                     |         | 20.30  |                       |                 | 19.66                                    |         |          |  |
|           | 22.92  |            |         | 23.59         |                                     |         | 24.30  |                       |                 | 24.35                                    |         |          |  |
| Figure 4c | 22.95  | 1.00       | 0.02    | 23.63         | 0.47                                | 0.00    | 24.42  | 0.63                  | 0.03            | 24.17                                    | 0.51    | 0.07     |  |
| _         | 22.90  |            |         | 23.63         |                                     |         | 24.35  |                       |                 | 23.98                                    |         |          |  |
|           | 23.12  |            |         | 23.01         |                                     |         | 23.93  |                       |                 | 23.99                                    |         |          |  |
| Figure 4d | 23.07  | 1.00       | 0.02    | 23.10         | 0.78                                | 0.02    | 23.97  | 0.94                  | 0.02            | 23.76                                    | 0.66    | 0.10     |  |
| _         | 23.09  | 1          |         | 23.06         | 1                                   |         | 23.94  | 1                     |                 | 24.20                                    | 1       |          |  |

a HDAC7-AS1 overexpression vector or a MIR-7-5p mimic exposed to 5 µM PCB29-pQ.

|           | NC min       | nic + pEZ- | M61-NC  | NC mimic + pEZ-M61-NC + |         |         | NC min   | nic + pEZ | -M61-   | <i>MIR-7-5p</i> mimic + pEZ- |          |          |
|-----------|--------------|------------|---------|-------------------------|---------|---------|----------|-----------|---------|------------------------------|----------|----------|
|           |              |            |         | PCB29-pQ                |         |         | HDAC7-   | ASI + PCI | B29-pQ  | M6                           | 1-HDAC7- | -ASI +   |
|           |              |            |         |                         |         |         |          |           |         |                              | РСВ29-р  | Q        |
|           | Graysc       | Mean       | SD      | Grayscal                | Mean    | SD      | Grayscal | Mean      | SD      | Grays                        | Mean     | SD       |
|           | ale          | (normal    | (normal | e values                | (normal | (normal | e values | (normal   | (normal | cale                         | (normal  | (normali |
|           | values       | ized)      | ized)   |                         | ized)   | ized)   |          | ized)     | ized)   | values                       | ized)    | zed)     |
| Figure 4f | 26963.<br>37 | 1.00       | 0.20    | 16137.55                | 0.61    | 0.12    | 28054.02 | 1.03      | 0.11    | 40 <u>5</u> 4.<br>58         | 0.15     | 0.03     |

|           | 26947. |      |      |          |      |      |          |      |      | 4076. |      |      |
|-----------|--------|------|------|----------|------|------|----------|------|------|-------|------|------|
|           | 55     |      |      | 16115.34 |      |      | 28059.10 |      |      | 37    |      |      |
|           | 26987. |      |      |          |      |      |          |      |      | 4068. |      |      |
|           | 28     |      |      | 16118.57 |      |      | 28038.02 |      |      | 09    |      |      |
|           | 34607. |      |      |          |      |      |          |      |      | 4518. |      |      |
|           | 94     |      |      | 2472.08  |      |      | 79170.54 |      |      | 18    |      |      |
| Figuro 4g | 34653. | 1.00 | 0.25 |          | 0.07 | 0.01 |          | 0.40 | 0.08 | 4527. | 0.12 | 0.02 |
| Figure 4g | 02     | 1.00 | 0.23 | 2490.17  | 0.07 | 0.01 | 79273.68 | 0.40 | 0.08 | 56    | 0.15 | 0.05 |
|           | 34649. |      |      |          |      |      |          |      |      | 4534. |      |      |
|           | 21     |      |      | 2488.51  |      |      | 79264.95 |      |      | 10    |      |      |

|           | NC min  | nic + pEZ- | M61-NC  | NC mimic + pEZ-M61-NC + |         |         | NC min    | nic + pEZ | -M61-   | <i>MIR-7-5p</i> mimic + pEZ- |         |          |  |
|-----------|---------|------------|---------|-------------------------|---------|---------|-----------|-----------|---------|------------------------------|---------|----------|--|
|           |         |            |         | PCB29-pQ                |         |         | HDAC7-    | AS1 + PC  | B29-pQ  | M6                           | 1-HDAC7 | -ASI +   |  |
|           |         |            |         |                         |         |         |           |           |         |                              | PCB29-p | Q        |  |
|           | Individ | Mean       | SD      | Individu                | Mean    | SD      | Individu  | Mean      | SD      | Indivi                       | Mean    | SD       |  |
|           | ual     | (normal    | (normal | al values               | (normal | (normal | al values | (normal   | (normal | dual                         | (normal | (normali |  |
|           | values  | ized)      | ized)   |                         | ized)   | ized)   |           | ized)     | ized)   | values                       | ized)   | zed)     |  |
|           | 98.67   |            |         | 73.49                   |         |         | 90.16     |           |         | 78.20                        |         |          |  |
| Figure 4h | 99.32   | 100.00     | 1.77    | 75.34                   | 72.17   | 4.00    | 100.47    | 93.44     | 6.10    | 76.08                        | 76.38   | 1.69     |  |
|           | 102.01  |            |         | 67.68                   |         |         | 89.69     |           |         | 74.85                        |         |          |  |

Mean and SD values are the ones presented in Figure 4a (relative MIR-7-5p mRNA level), b (relative HDAC7-AS1 mRNA level), c (relative TGF-

 $\beta$ 2 mRNA level), d (relative *PPME1* mRNA level), f (relative TGF- $\beta$ 2/ $\beta$ -actin protein level), g (relative PPME1/ $\beta$ -actin protein level), and h (cell viability, %). Results presented relative to NC mimic + pEZ-M61-NC.

|                               |                          | AAV-NC | l<br>, | AAV-N                   | NC + PCB2 | 29-pQ | AAV                    | -HDAC7-A | 4 <i>S1</i> | AA                       | V- <i>HDAC7</i><br>PCB29-p | - <i>AS1</i> +<br>•Q |
|-------------------------------|--------------------------|--------|--------|-------------------------|-----------|-------|------------------------|----------|-------------|--------------------------|----------------------------|----------------------|
|                               | Individ<br>ual<br>values | Mean   | SD     | Individu<br>al values   | Mean      | SD    | Individu<br>al values  | Mean     | SD          | Indivi<br>dual<br>values | Mean                       | SD                   |
| Figure 5a<br>(Right<br>panel) | 10.87<br>12.58<br>14.44  | 12.63  | 1.79   | 21.48<br>26.69<br>29.17 | 25.78     | 3.93  | 10.69<br>10.79<br>7.45 | 9.64     | 1.90        | 6.61<br>8.77<br>9.65     | 8.35                       | 1.57                 |
| Figure 5c                     | 0.67<br>0.66<br>0.76     | 0.70   | 0.06   | 0.27<br>0.23<br>0.27    | 0.26      | 0.02  | 0.95<br>0.92<br>0.93   | 0.93     | 0.01        | 0.71<br>0.78<br>0.86     | 0.78                       | 0.07                 |

Table S17. Atherosclerosis and inflammation in ApoE<sup>-/-</sup> mice exposed to PCB29-pQ accompanied by AAV-*HDAC7-AS1* treatment.

|           |        | A AV NC |         |        |           | 20 0    | ۸ ۸۲   |                   | 151     | AA     | V-HDAC7 | -ASI +   |
|-----------|--------|---------|---------|--------|-----------|---------|--------|-------------------|---------|--------|---------|----------|
|           |        | AAV-NC  | ,       | AAV-1  | NC + PCD. | 29-pQ   | AAV    | - <i>HDAC</i> /-/ | 431     |        | РСВ29-р | Q        |
|           | Cq     | Mean    | SD      | Cq     | Mean      | SD      | Cq     | Mean              | SD      | Cq     | Mean    | SD       |
|           | values | (normal | (normal | values | (normal   | (normal | values | (normal           | (normal | values | (normal | (normali |
|           |        | ized)   | ized)   |        | ized)     | ized)   |        | ized)             | ized)   |        | ized)   | zed)     |
|           | 24.72  |         |         | 26.83  |           |         | 22.80  |                   |         | 24.16  |         |          |
| Figure 5d | 25.47  | 1.00    | 0.29    | 27.68  | 0.25      | 0.10    | 22.30  | 17.06             | 4.81    | 24.78  | 7.19    | 2.05     |
|           | 24.86  |         |         | 27.83  |           |         | 21.96  |                   |         | 24.88  |         |          |
|           | 25.56  |         |         | 22.63  |           |         | 28.07  |                   |         | 27.65  |         |          |
| Figure 5e | 24.70  | 1.00    | 0.29    | 23.57  | 5.79      | 1.81    | 28.21  | 0.29              | 0.05    | 28.35  | 0.96    | 0.45     |
|           | 25.09  |         |         | 22.71  |           |         | 28.53  |                   |         | 26.97  |         |          |
|           | 24.46  |         |         | 28.62  |           |         | 25.99  |                   |         | 27.41  |         |          |
| Figure 5f | 24.87  | 1.00    | 0.24    | 27.88  | 0.14      | 0.05    | 24.62  | 2.59              | 1.29    | 26.66  | 1.34    | 0.42     |
| -         | 25.20  |         |         | 27.62  |           |         | 24.33  |                   |         | 26.46  |         |          |
|           | 24.56  |         |         | 26.09  |           |         | 24.60  |                   |         | 26.84  |         |          |
| Figure 5g | 24.24  | 1.00    | 0.27    | 26.22  | 0.33      | 0.02    | 23.50  | 2.65              | 1.27    | 26.31  | 0.93    | 0.22     |
|           | 23.76  |         |         | 26.00  |           |         | 24.60  |                   |         | 26.91  |         |          |
| Figure 5h | 24.74  | 1.00    | 0.71    | 21.78  | 4.57      | 1.52    | 28.29  | 0.24              | 0.14    | 26.04  | 1.13    | 0.55     |

|           | 23.31                                    |              |            | 22.76                                 |   |            | 26.74                                 |              |            | 25.84                                    |                 |                |
|-----------|--|--------------|------------|---------------------------------------|---|------------|---------------------------------------|--------------|------------|--|-----------------|----------------|
|           | 24.25                                    |              |            | 22.25                                 |   |            | 27.92                                 |              |            | 27.46                                    |                 |                |
|           | 24.01                                    |              |            | 22.69                                 |   |            | 28.17                                 |              |            | 26.14                                    |                 |                |
| Figure 5i | 24.14                                    | 1.00         | 0.14       | 23.26                                 | 3.72  | 1.54       | 26.68                                 | 0.32         | 0.15       | 26.44                                    | 1.18            | 0.13           |
|           | 24.39                                    |              |            | 22.05                                 |   |            | 27.09                                 |              |            | 26.33                                    |                 |                |
|           | 23.25                                    |              |            | 23.05                                 |   |            | 30.36                                 |              |            | 26.23                                    |                 |                |
| Figure 5j | 25.77                                    | 1.00         | 0.70       | 22.58                                 | 4.65  | 1.83       | 25.97                                 | 0.90         | 0.88       | 27.98                                    | 0.74            | 0.58           |
|           | 24.07                                    |              |            | 21.93                                 |   |            | 24.86                                 |              |            | 28.05                                    |                 |                |
|           |  |              |            |                                       |   | 00 mO      | A A37                                 |              | 151        | AA                                       | V-HDAC7         | - <i>AS1</i> + |
|           |  | AAV-NC       |            | AAV-N                                 | $\mathbf{V} + \mathbf{F} \mathbf{C} \mathbf{D}$ | 29-pQ      | AAV                                   | -IIDAC/-     | 451        |  | PCB29-n         | 0              |
|           |  |              |            |                                       |   |            |                                       |              |            |  | reber           | Y Y            |
|           | Individ                                  |              |            | Individu                              |   |            | Individu                              |              |            | Indivi                                   | <u>1002</u> , p | ~              |
|           | Individ<br>ual                           | Mean         | SD         | Individu<br>al values                 | Mean  | SD         | Individu<br>al values                 | Mean         | SD         | Indivi<br>dual                           | Mean            | SD             |
|           | Individ<br>ual<br>values                 | Mean         | SD         | Individu<br>al values                 | Mean  | SD         | Individu<br>al values                 | Mean         | SD         | Indivi<br>dual<br>values                 | Mean            | SD             |
|           | Individ<br>ual<br>values<br>0.57         | Mean         | SD         | Individu<br>al values<br>0.94         | Mean  | SD         | Individu<br>al values<br>0.17         | Mean         | SD         | Indivi<br>dual<br>values<br>0.77         | Mean            | SD             |
| Figure 5k | Individ<br>ual<br>values<br>0.57<br>0.42 | Mean<br>0.43 | SD<br>0.13 | Individu<br>al values<br>0.94<br>0.94 | Mean<br>0.95                                    | SD<br>0.02 | Individu<br>al values<br>0.17<br>0.19 | Mean<br>0.17 | SD<br>0.02 | Indivi<br>dual<br>values<br>0.77<br>0.72 | Mean<br>0.71    | SD<br>0.08     |

Mean and SD values are the ones presented in Figure 5a (plaque area, % of whole aorta), c (Pearson's correlation coefficient), d (relative *HDAC7-AS1* mRNA level), e (relative *MIR-7-5p* mRNA level), f (relative *TGF-\beta2* mRNA level), g (relative *PPME1* mRNA level), h (relative *IL-1\beta* mRNA level), i (relative *IL6* mRNA level), (relative *TNF* $\alpha$  mRNA level), and k (Pearson's correlation coefficient). Data in panels 5d through 5J are normalized to AAV-NC.

| Table S18. | Inflammation a | and atherogen | lesis in Apol | E <sup>-/-</sup> and Apol | <i>E-/-CAV1-/-</i> mice | exposed to PCB29-pQ. |
|------------|----------------|---------------|---------------|---------------------------|-------------------------|----------------------|
|            |                |               |               |                           |                         |                      |

| Eigung 6g     |                   | ApoE-/- |    | ApoE <sup>-/-</sup> /CAV1 <sup>-/-</sup> |      |    |  |
|---------------|-------------------|---------|----|--|------|----|--|
| (Right panel) | Individual values | Mean    | SD | Individual values                        | Mean | SD |  |

|          | 18.02 |       |      | 7.57  |       |      |
|----------|-------|-------|------|-------|-------|------|
| Control  | 18.86 | 18.86 | 0.84 | 4.54  | 6.78  | 1.97 |
|          | 19.70 |       |      | 8.24  |       |      |
|          | 22.60 |       |      | 10.90 |       |      |
| PCB29-pQ | 27.59 | 25.78 | 2.76 | 10.03 | 10.94 | 0.94 |
|          | 27.15 |       |      | 11.90 |       |      |

|                            |          |            | ApoE-/- |      |            | ApoE-/-/CAV1-/- |      |  |
|----------------------------|----------|------------|---------|------|------------|-----------------|------|--|
|                            |          | Individual | Mean    | SD   | Individual | Mean            | SD   |  |
|                            |          | values     | Ivicali | 30   | values     | Wieall          | 50   |  |
|                            |          | 16.00      |         | 2.53 | 10.00      |                 |      |  |
|                            |          | 14.00      |         |      | 9.00       | 9.13            |      |  |
|                            |          | 20.00      | - 14.88 |      | 11.00      |                 |      |  |
|                            | Control  | 15.00      |         |      | 8.00       |                 | 1.73 |  |
|                            |          | 16.00      |         |      | 12.00      |                 |      |  |
| Figure 6b<br>(Right panel) |          | 12.00      |         |      | 8.00       |                 |      |  |
|                            |          | 13.00      |         |      | 8.00       |                 |      |  |
|                            |          | 13.00      |         |      | 7.00       |                 |      |  |
|                            |          | 25.00      |         |      | 16.00      | -               |      |  |
|                            |          | 20.00      |         |      | 12.00      |                 |      |  |
|                            |          | 19.00      |         |      | 11.00      |                 |      |  |
|                            |          | 17.00      | 22.50   | 1 20 | 18.00      | 12.05           | 2.66 |  |
|                            | PCB29-pQ | 28.00      | 25.50   | 4.38 | 10.00      | 15.25           | 2.00 |  |
|                            |          | 29.00      |         |      | 13.00      |                 |      |  |
|                            |          | 24.00      |         |      | 14.00      |                 |      |  |
|                            | -        | 26.00      |         |      | 12.00      |                 |      |  |

|                      |          | Ap   | $poE^{-/-}$ | $ApoE^{-/-}/c$ | CAV1 <sup>-/-</sup> |
|----------------------|----------|------|-------------|----------------|---------------------|
| Figure 6c (p-p65/p65 |          | Mean | SD          | Mean           | SD                  |
| ratio)               | Control  | 1.00 | 0.16        | 0.23           | 0.09                |
|                      | PCB29-pQ | 2.33 | 0.86        | 0.96           | 0.17                |

|                     |                  |          |          | Ap                                       | oE-/- |          |                | ApoE-    | -/CAV1-/ | -           |
|---------------------|------------------|----------|----------|--|-------|----------|----------------|----------|----------|-------------|
| Figure 6c (p-C      | AV1/CAV1         |          | Mear     | 1  |       | SD       | Me             | an       |          | SD          |
| ratio               | )                | Control  | 1.00     |  |       | 0.11     | 0.             | )8       |          | 0.02        |
|                     |                  | PCB29-pQ | 3.75     |  |       | 0.92     | 0.07           |          |          | 0.01        |
|                     |                  |          |          |  |       |          |                |          | -        |             |
|                     |                  | Con      | trol     |  |       |          |                | PCB29-pQ |          |             |
|                     | A                | poE-/-   | ApoE-/-  | ApoE <sup>-/-</sup> /CAV1 <sup>-/-</sup> |       |          | <i>poE</i> -/- |          | ApoE     | -/-/CAV1-/- |
| Figure 6c (p-       | 18025.72         | 15823.21 | 7120.71  | 771                                      | 0.74  | 17526.13 | 17479.4        | 5 85     | 50.24    | 9882.45     |
| p65)                | 18017.23         | 15808.01 | 7137.93  | 772                                      | 6.64  | 17416.06 | 17442.1        | 1 85     | 93.80    | 9883.14     |
| Grayscale<br>values | cale 17994.76 15 |          | 7124.79  | 7124.79 7711.38                          |       | 17585.55 | 17606.1        | 8 85     | 72.30    | 9853.36     |
| Figure 6c           | 21915.41         | 17645.41 | 18709.33 | 1714                                     | 44.99 | 13269.30 | 10481.2        | 4 113    | 73.95    | 11423.09    |
| (p65)               | 21890.29         | 17623.68 | 18726.23 | 1710                                     | 50.83 | 13420.54 | 10315.4        | 0 114    | 02.96    | 11403.68    |
| Grayscale<br>values | 21920.01         | 17614.19 | 18723.14 | 17145.52                                 |       | 13415.86 | 10358.8        | 1 114    | 03.80    | 11389.21    |
| Figure 6c (p-       | 9486.03          | 8721.17  | 484.76   | 840                                      | 0.55  | 21084.55 | 25457.1        | 7 18     | 37.09    | 507.38      |
| CAV1)               | 9484.11          | 8723.01  | 516.13   | 880                                      | 0.38  | 21127.93 | 25430.5        | 9 18     | 24.28    | 422.53      |
| Grayscale<br>values | 9504.05          | 8703.10  | 478.96   | 857                                      | 7.29  | 21099.89 | 25474.2        | 5 20     | 03.54    | 621.80      |
| Figure 6c           | 19233.40         | 13857.08 | 3048.81  | 596                                      | 4.90  | 20226.02 | 20341.4        | 2 45     | 73.50    | 5367.20     |
| (CAV1)              | 19238.14         | 13878.08 | 3088.71  | 600                                      | 4.51  | 20239.88 | 20364.7        | 3 44     | 63.46    | 5460.55     |
| Grayscale<br>values | 19217.17         | 13861.90 | 3071.12  | 597                                      | 9.90  | 20269.11 | 20407.6        | 4 46     | 63.11    | 5260.69     |

|                               |                  |            | $ApoE^{-/-}$ |      | ApoE <sup>-/-</sup> /CAV1 <sup>-/-</sup> |         |      |  |  |  |
|-------------------------------|------------------|------------|--------------|------|--|---------|------|--|--|--|
| Figure 6d<br>(Right<br>panel) |                  | Individual | Moon         | SD   | Individual                               | Moon    | SD   |  |  |  |
|                               |                  | values     | Ivicali      | 3D   | values                                   | Ivicali | 3D   |  |  |  |
|                               | Control PCB29-pQ | 0.29       |              |      | 0.09                                     |         |      |  |  |  |
|                               |                  | 0.19       | 0.19         | 0.10 | 0.15                                     | 0.09    | 0.06 |  |  |  |
|                               |                  | 0.10       |              |      | 0.04                                     |         |      |  |  |  |
|                               |                  | 0.96       | 0.86         | 0.00 | 0.59                                     | 0.59    | 0.02 |  |  |  |
|                               |                  | 0.84       | 0.80         | 0.09 | 0.57                                     | 0.38    | 0.02 |  |  |  |

| 0.78 |
|------|
|------|

|                  |          |            | ApoE <sup>-/-</sup> |      | ApoE-/-/CAVI-/- |       |      |  |  |
|------------------|----------|------------|---------------------|------|-----------------|-------|------|--|--|
|                  |          | Individual | Mean                | SD   | Individual      | Mean  | SD   |  |  |
|                  |          | values     | Wiedii              | 50   | values          | wiean | 3D   |  |  |
| Figure 6e        |          | 0.18       |                     |      | 0.05            |       |      |  |  |
| (Right<br>panel) | Control  | 0.10       | 0.23                | 0.15 | 0.28            | 0.13  | 0.13 |  |  |
|                  |          | 0.40       |                     |      | 0.06            |       |      |  |  |
|                  |          | 0.99       |                     |      | 0.53            |       |      |  |  |
|                  | PCB29-pQ | 1.00       | 0.93                | 0.11 | 0.45            | 0.49  | 0.04 |  |  |
|                  |          | 0.81       |                     |      | 0.49            |       |      |  |  |

|           |          |            | ApoE <sup>-/-</sup> |        | ApoE-/-/C  | 'AV1-/- |       |
|-----------|----------|------------|---------------------|--------|------------|---------|-------|
|           |          | Individual | Mean                | SD     | Individual | Mean    | SD    |
|           |          | values     | Weam                | 50     | values     | Wiean   | 50    |
|           |          | 440.40     |                     |        | 263.67     |         |       |
| Figure 6f | Control  | 183.36     | 365.54              |        | 202.68     | 272.87  | 60.05 |
|           |          | 296.01     |                     | 157.76 | 364.26     |         |       |
|           |          | 312.06     |                     |        | 244.65     |         |       |
|           |          | 595.89     |                     |        | 289.08     |         |       |
|           |          | 639.27     |                     |        | 563.34     |         |       |
|           |          | 942.63     |                     |        | 501.93     |         |       |
|           | PCB29-pQ | 594.96     | 760.11              | 174.41 | 487.67     | 523.07  | 39.67 |
|           |          | 702.90     |                     |        | 493.50     | 1       |       |
|           |          | 470.79     |                     |        | 568.89     |         |       |

| Figure 6g |         |                   | ApoE <sup>-/-</sup> | ApoE <sup>-/-</sup> /CAV1 <sup>-/-</sup> |                   |        |        |  |
|-----------|---------|-------------------|---------------------|--|-------------------|--------|--------|--|
|           |         | Individual values | Mean                | SD                                       | Individual values | Mean   | SD     |  |
|           | Control | 1468.8            | 1226.24             | 108 68                                   | 878.90            | 010.03 | 205.04 |  |
|           | Control | 986.7             | 1550.54             | 408.08                                   | 675.60            | 919.95 | 203.04 |  |

|          | 1042.9  |         |         | 1214.60 |         |        |
|----------|---------|---------|---------|---------|---------|--------|
|          | 1196.8  |         |         | 815.50  |         |        |
|          | 1986.5  |         |         | 1015.05 |         |        |
|          | 2130.9  |         |         | 1775.85 |         |        |
|          | 2056.95 |         |         | 1877.80 |         |        |
| PCB29-pQ | 1983.00 | 2135.64 | 135.701 | 1673.90 | 1791.71 | 142.14 |
|          | 2343.9  |         |         | 1645.00 |         |        |
|          | 2163.45 |         |         | 1986.00 |         |        |

Mean and SD values are the ones presented in Figure 6a (plaque area, % of whole aorta), b (plaque area, % of whole aorta), c (ratio of p-p65/p65 and pCAV1/CAV1 protein), d (Pearson's correlation coefficient), e (Pearson's correlation coefficient), f (*IL-6* (pg/ml)), and g (TNF $\alpha$  (pg/ml)). Data in figure 6c normalized to control ApoE<sup>-/-</sup> mice.

Table S19. Plasma HDAC7-AS1 (a), MIR-7-5p (b), TGF-β2 (c), PPME1 (d), IL-1β (e), IL-6 (f), and TNFα (g) levels in patients with CHD.

| Figu    | re 7a | Figu    | re 7b | Figu    | re 7c | Figu    | re 7d | Figu    | ire 7e | Figu    | re 7f | Figure  | e 7g |
|---------|-------|---------|-------|---------|-------|---------|-------|---------|--------|---------|-------|---------|------|
| Control | CHD    | Control | CHD   | Control | CHD  |
| 24.40   | 17.03 | 26.95   | 21.46 | 17.38   | 32.35 | 23.17   | 22.83 | 28.05   | 26.75  | 26.46   | 26.33 | 25.47   | 28.4 |
|         |       |         |       | 11.00   | 02.00 |         |       |         |        |         |       |         | 0    |
| 24.38   | 16.70 | 21.71   | 18.75 | 27.06   | 31.23 | 25.94   | 21.26 | 26.89   | 27.72  | 26.31   | 25.75 | 25.08   | 29.5 |
|         |       |         |       | 27.00   | 51.25 |         |       |         |        |         |       |         | 4    |
| 24.73   | 17.50 | 23.92   | 19.60 | 27.22   | 27.70 | 21.96   | 20.26 | 27.59   | 26.80  | 26.93   | 24.60 | 26.37   | 27.0 |
|         |       |         |       | 21.22   | 27.70 |         |       |         |        |         |       |         | 9    |
| 26.39   | 18.01 | 24.19   | 21.27 | 25.71   | 20.52 | 25.84   | 22.09 | 30.89   | 26.19  | 27.07   | 26.05 | 26.03   | 29.1 |
|         |       |         |       | 23.71   | 30.32 |         |       |         |        |         |       |         | 0    |
| 26.66   | 18.62 | 23.32   | 20.38 | 26.01   | 27.00 | 22.52   | 32.47 | 25.92   | 27.88  | 23.76   | 25.85 | 26.42   | 25.0 |
|         |       |         |       | 26.91   | 27.98 |         |       |         |        |         |       |         | 2    |

| 24.63 | 29.38 | 24.02 | 18.65 | 22.69 | 28.15 | 22.00 | 32.32 | 29.56 | 23.34 | 26.87 | 23.90 | 26.55 | 23.1      |
|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-----------|
| 20.82 | 26.60 | 24.21 | 19.90 | 26.20 | 28.91 | 22.73 | 31.89 | 27.86 | 29.65 | 24.99 | 26.18 | 24.61 | 28.8<br>4 |
| 23.75 | 27.35 | 26.49 | 20.30 | 21.52 | 32.17 | 24.02 | 28.22 | 29.81 | 26.42 | 29.28 | 26.11 | 26.27 | 27.1      |
| 23.73 | 25.62 | 25.65 | 20.36 | 26.50 | 30.99 | 21.59 | 31.79 | 25.57 | 26.88 | 29.21 | 27.14 | 28.76 | 29.1<br>0 |
| 23.48 | 27.79 | 25.69 | 21.38 | 21.88 | 28.96 | 24.75 | 31.06 | 30.67 | 29.34 | 26.95 | 24.87 | 26.84 | 27.3<br>0 |
| 21.54 | 17.03 | 22.34 | 22.38 | 24.32 | 28.93 | 24.20 | 31.08 | 28.53 | 26.98 | 26.79 | 25.66 | 25.37 | 27.9<br>9 |
| 25.02 | 18.47 | 27.15 | 18.97 | 22.97 | 30.76 | 25.56 | 33.10 | 28.05 | 26.14 | 26.17 | 22.90 | 27.85 | 29.5<br>6 |
| 25.71 | 27.46 | 26.63 | 22.45 | 22.97 | 27.62 | 24.36 | 31.48 | 27.66 | 27.62 | 28.52 | 24.70 | 26.99 | 23.6      |
| 27.24 | 24.41 | 22.03 | 19.47 | 23.41 | 29.54 | 24.97 | 33.91 | 28.62 | 22.97 | 25.47 | 20.60 | 25.51 | 21.4      |
| 26.00 | 26.54 | 22.93 | 21.68 | 27.37 | 22.61 | 22.12 | 31.18 | 28.34 | 24.86 | 27.30 | 23.71 | 26.18 | 24.0<br>4 |
| 25.59 | 27.31 | 23.93 | 21.62 | 19.07 | 21.69 | 22.73 | 33.09 | 28.44 | 27.58 | 24.79 | 23.96 | 27.19 | 21.2<br>8 |
| 22.93 | 26.28 | 23.83 | 19.80 | 24.84 | 20.37 | 24.34 | 28.62 | 29.27 | 27.66 | 28.34 | 25.85 | 25.64 | 24.5<br>2 |
| 24.91 | 24.47 | 23.21 | 19.05 | 24.31 | 20.61 | 24.50 | 27.25 | 29.48 | 27.54 | 25.62 | 25.07 | 25.77 | 19.5<br>5 |
| 27.63 | 28.80 | 24.73 | 21.78 | 24.79 | 21.91 | 23.89 | 33.00 | 27.57 | 26.11 | 25.50 | 22.57 | 27.13 | 22.9<br>1 |
| 25.98 | 27.10 | 26.02 | 20.82 | 24.55 | 20.34 | 24.25 | 30.97 | 29.76 | 28.82 | 25.57 | 25.63 | 25.43 | 24.4<br>2 |
| 24.31 | 25.62 | 26.22 | 20.33 | 23.14 | 31.55 | 21.64 | 29.10 | 29.07 | 29.52 | 26.27 | 24.04 | 25.36 | 26.4<br>0 |

| 24.56 | 26.77 | 22.76 | 19.64 | 24.07 | 21.17 | 23.40 | 27.67 | 29.37 | 28.93 | 27.40 | 23.99 | 27.00 | 25.6<br>5 |
|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-----------|
| 25.97 | 25.95 | 23.12 | 18.46 | 26.98 | 19.34 | 24.75 | 29.08 | 28.37 | 25.95 | 26.61 | 23.82 | 27.34 | 22.4      |
| 22.61 | 26.23 | 25.47 | 16.88 | 21.48 | 31.40 | 24.52 | 30.81 | 25.08 | 27.64 | 27.63 | 23.43 | 27.00 | 23.9      |
| 23.53 | 28.28 | 26.95 | 17.52 | 28.94 | 30.97 | 23.50 | 31.62 | 30.07 | 25.54 | 25.23 | 24.00 | 25.12 | 23.4<br>7 |
| 20.89 | 26.15 | 31.41 | 19.84 | 30.12 | 27.30 | 23.47 | 28.22 | 28.83 | 27.22 | 25.25 | 24.97 | 24.37 | 25.6      |
| 25.70 | 26.94 | 28.53 | 21.62 | 25.47 | 30.87 | 24.79 | 32.74 | 27.76 | 28.90 | 24.64 | 28.41 | 26.23 | 24.6      |
| 23.08 | 18.17 | 25.56 | 22.58 | 22.05 | 30.14 | 21.02 | 20.98 | 26.73 | 27.65 | 23.51 | 25.26 | 24.57 | 21.11     |
| 24.53 | 17.10 | 27.49 | 18.32 | 21.63 | 30.16 | 22.93 | 20.44 | 24.76 | 28.33 | 25.76 | 25.69 | 26.09 | 25.4<br>1 |
| 22.56 | 16.96 | 21.82 | 20.76 | 25.13 | 32.18 | 23.42 | 19.50 | 29.77 | 29.08 | 25.28 | 26.64 | 26.62 | 26.5<br>6 |
| 24.80 | 17.17 | 25.41 | 22.99 | 27.80 | 30.56 | 21.78 | 19.56 | 26.22 | 27.77 | 28.06 | 24.81 | 28.60 | 24.7      |
| 23.42 | 16.98 | 28.78 | 21.06 | 26.59 | 32.99 | 24.27 | 19.06 | 30.13 | 28.13 | 26.37 | 25.42 | 24.48 | 23.3      |
| 24.93 | 17.58 | 25.71 | 21.40 | 25.39 | 30.26 | 25.43 | 17.99 | 28.87 | 25.10 | 24.59 | 25.31 | 26.69 | 25.7      |
| 24.80 | 16.43 | 23.40 | 22.40 | 24.45 | 32.17 | 25.03 | 20.08 | 26.36 | 28.20 | 24.82 | 25.48 | 27.06 | 24.7<br>2 |
| 21.34 | 18.70 | 23.48 | 22.66 | 21.98 | 27.70 | 25.80 | 19.72 | 29.34 | 27.89 | 26.37 | 24.72 | 27.92 | 24.7      |
| 23.56 | 18.01 | 18.09 | 18.42 | 23.86 | 26.33 | 24.45 | 20.57 | 26.82 | 28.16 | 26.79 | 24.88 | 27.88 | 24.4      |
| 23.84 | 18.62 | 21.36 | 18.37 | 23.75 | 32.08 | 21.80 | 23.20 | 28.89 | 29.05 | 24.64 | 22.85 | 27.09 | 25.6<br>8 |
| 23.78 | 29.38 | 22.55 | 18.47 | 20.19 | 30.05 | 25.88 | 20.93 | 28.22 | 28.18 | 26.81 | 25.03 | 27.07 | 22.5<br>9 |

| 24.61 | 26.60 | 24.64 | 17.85 | 22.25 | 28.18 | 23.52 | 20.19 | 28.08 | 26.65 | 26.81 | 25.67 | 25.28 | 23.9      |
|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-----------|
| 24.43 | 27.35 | 24.50 | 18.46 | 28.17 | 26.75 | 26.83 | 30.58 | 26.96 | 26.85 | 26.04 | 24.18 | 26.43 | 19.9<br>6 |
| 24.17 | 25.62 | 23.62 | 19.98 | 24.66 | 28.16 | 23.29 | 30.42 | 25.06 | 28.19 | 23.88 | 24.73 | 28.14 | 23.8<br>7 |
| 25.59 | 27.79 | 23.61 | 20.76 | 20.29 | 29.89 | 23.20 | 29.99 | 27.17 | 29.17 | 24.94 | 26.37 | 25.59 | 25.6<br>7 |
| 24.24 | 17.03 | 24.86 | 19.84 | 27.33 | 30.70 | 24.03 | 26.32 | 30.90 | 26.86 | 27.18 | 25.30 | 23.76 | 22.5<br>2 |
| 24.80 | 18.47 | 28.15 | 17.52 | 26.22 | 27.30 | 22.84 | 29.89 | 29.40 | 27.64 | 26.81 | 24.49 | 27.41 | 20.0<br>9 |
| 24.22 | 27.46 | 22.87 | 21.62 | 26.29 | 31.82 | 23.52 | 29.16 | 28.27 | 28.14 | 28.09 | 25.03 | 29.05 | 22.8<br>0 |
| 25.07 | 24.41 | 31.84 | 22.58 | 28.20 | 20.06 | 22.68 | 29.18 | 29.66 | 26.93 | 27.23 | 25.27 | 26.13 | 25.3<br>0 |
| 25.53 | 26.54 | 19.78 | 22.99 | 26.37 | 19.52 | 23.43 | 31.21 | 26.81 | 27.03 | 26.17 | 26.15 | 26.30 | 25.3<br>8 |
| 22.12 | 27.31 | 15.95 | 18.32 | 22.16 | 20.61 | 20.88 | 29.58 | 30.93 | 21.96 | 28.98 | 23.39 | 26.28 | 20.0<br>7 |
| 25.14 | 26.28 | 18.64 | 16.88 | 23.59 | 24.97 | 24.95 | 32.02 | 30.59 | 27.74 | 26.02 | 26.37 | 25.63 | 19.4<br>7 |
| 24.55 | 24.47 | 22.69 | 21.06 | 17.99 | 25.51 | 23.72 | 29.29 | 31.18 | 29.14 | 28.14 | 25.12 | 26.25 | 22.9<br>2 |
|       | 28.80 |       | 21.40 |       | 21.83 |       | 31.20 |       | 27.67 |       | 23.81 |       | 26.5<br>4 |
|       | 27.10 |       | 22.40 |       | 25.91 |       | 26.72 |       | 24.95 |       | 25.28 |       | 24.9<br>8 |
|       | 25.62 |       | 22.66 |       | 22.55 |       | 25.35 |       | 26.42 |       | 24.18 |       | 23.8<br>5 |
|       | 26.77 |       | 22.78 |       | 28.39 |       | 31.10 |       | 26.48 |       | 25.32 |       | 25.1<br>9 |

| 25.95 | 22.53   | 26.76 | 29.08 | 27.60 | 24.06 | 23.2 |
|-------|---------|-------|-------|-------|-------|------|
| 26.23 | 18.47   | 25.04 | 27.20 | 24.20 | 21.22 | 19.6 |
| 28.28 | 3 17.85 | 25.60 | 25.77 | 25.13 | 27.01 | 23.3 |
| 26.15 | 5 19.98 | 25.56 | 27.18 | 21.71 | 24.07 | 20.9 |
| 26.94 | 20.36   | 24.00 | 28.92 | 20.16 | 25.66 | 24.1 |
| 18.17 | 21.38   | 24.10 | 29.72 | 24.06 | 24.33 | 24.8 |
| 17.10 | 22.38   | 25.80 | 26.32 | 24.95 | 26.94 | 26.3 |
| 16.96 | 5 18.97 | 24.15 | 30.84 | 27.50 | 26.60 | 24.2 |
| 17.17 | 22.45   | 28.67 | 19.08 | 25.90 | 24.91 | 23.4 |
| 16.98 | 3 19.47 | 28.12 | 18.54 | 22.59 | 25.61 | 25.0 |
| 17.58 | 3 21.68 | 27.74 | 21.27 | 26.42 | 24.18 | 24.2 |
| 18.70 | ) 21.62 | 23.58 | 21.19 | 22.78 | 25.95 | 24.9 |
| 19.07 | 7 19.80 | 23.18 | 28.50 | 24.59 | 29.75 | 23.9 |
| 20.22 | 2 19.05 | 21.90 | 28.33 | 24.62 | 26.11 | 23.1 |
| 26.62 | 21.78   | 24.63 | 28.35 | 20.44 | 28.57 | 22.8 |
| 26.54 | 25.50   | 23.31 | 28.33 | 20.87 | 26.74 | 25.4 |

| 18.63 | 24.72 | 22.90 | 29.86 | 25.99 | 25.41 | 24.0 |
|-------|-------|-------|-------|-------|-------|------|
|       |       |       |       |       |       | 0    |
| 18.34 | 18.42 | 28.07 | 29.74 | 20.77 | 32.32 | 22.3 |
|       |       | 20.97 |       |       |       | 5    |
| 18.34 | 18.3  |       | 30.04 | 24.53 | 26.12 | 21.7 |
|       |       | 28.83 |       |       |       | 5    |
| 18.34 | 24.85 | 20.00 | 28.80 | 23.03 | 27.66 | 23.6 |
|       |       | 29.06 |       |       |       | 6    |
| 20.06 | 21.93 | 20.07 | 27.53 | 24.04 | 26.57 | 23.3 |
|       |       | 29.07 |       |       |       | 4    |
| 25.70 | 19.08 |       | 27.53 | 23.73 | 22.95 | 22.9 |
| 20170 | 19100 | 28.96 | 2,    | 20170 | 22.90 | 3    |
| <br>  |       |       |       |       |       |      |
| 25.59 | 22.20 | 20.18 | 27.51 | 22.36 | 24.15 | 21.0 |
|       |       | 29.10 |       |       |       | 4    |
|       |       |       |       |       |       | т    |

Cq values are the ones presented in Figure 7a-g.

|           | Contr | ol   | CHD  |      |  |  |
|-----------|-------|------|------|------|--|--|
|           | Mean  | SD   | Mean | SD   |  |  |
| Figure 7a | 1.00  | 0.88 | 0.29 | 0.49 |  |  |
| Figure 7b | 1.00  | 1.60 | 4.52 | 5.65 |  |  |
| Figure 7c | 1.00  | 1.24 | 0.40 | 0.95 |  |  |
| Figure 7d | 1.00  | 1.45 | 0.44 | 1.07 |  |  |
| Figure 7e | 1.00  | 1.67 | 3.34 | 8.17 |  |  |
| Figure 7f | 1.00  | 1.35 | 2.55 | 4.55 |  |  |
| Figure 7g | 1.00  | 1.66 | 4.37 | 8.01 |  |  |

Mean and SD values are the ones presented in Figure 7a-g.

|        |                  |           | Control      |              |           | PCB29-pQ     |              |
|--------|------------------|-----------|--------------|--------------|-----------|--------------|--------------|
|        |                  |           | Mean         | SD           | Cavaluas  | Mean         | SD           |
|        |                  | Cq values | (normalized) | (normalized) | Cq values | (normalized) | (normalized) |
|        |                  | 26.22     |              |              | 26.75     |              |              |
|        | <i>LINC01547</i> | 26.28     | 1.00         | 0.04         | 26.86     | 1.00         | 0.07         |
|        |                  | 26.16     |              |              | 26.67     |              |              |
|        |                  | 25.42     |              |              | 24.96     |              |              |
|        | SLCO4A1-AS1      | 25.52     | 1.00         | 0.04         | 24.93     | 2.07         | 0.02         |
|        |                  | 25.43     |              |              | 24.93     |              |              |
|        |                  | 27.58     |              |              | 27.27     |              |              |
|        | NRSN2-ASI        | 27.27     | 1.00         | 0.11         | 27.69     | 1.34         | 0.27         |
|        |                  | 27.52     |              |              | 27.82     |              |              |
|        |                  | 26.39     |              |              | 27.44     |              |              |
|        | FGD5-AS1         | 26.32     | 1.00         | 0.03         | 27.36     | 0.71         | 0.02         |
| Figure |                  | 26.39     |              |              | 27.42     |              |              |
| S3a    |                  | 26.52     |              |              | 27.77     |              |              |
|        | MCM3AP-AS1       | 26.58     | 1.00         | 0.02         | 27.73     | 0.65         | 0.04         |
|        |                  | 26.54     |              |              | 27.60     |              |              |
|        |                  | 25.44     |              |              | 26.86     |              |              |
|        | TUG1             | 25.41     | 1.00         | 0.05         | 27.00     | 0.53         | 0.03         |
|        |                  | 25.55     |              |              | 26.92     |              |              |
|        |                  | 27.29     |              |              | 28.63     |              |              |
|        | RP3-416H24.1     | 27.52     | 1.00         | 0.08         | 28.65     | 0.62         | 0.02         |
|        |                  | 27.41     |              |              | 28.56     |              |              |
|        |                  | 24.52     |              |              | 24.81     |              |              |
|        | AC002550.5       | 25.36     | 1.00         | 0.34         | 24.75     | 1.80         | 0.12         |
|        |                  | 25.24     |              |              | 24.63     |              |              |
| Ī      | RP5-1057120.4    | 24.46     |              |              | 27.49     |              | 0.01         |
|        | (HADC7-AS1)      | 24.52     | 1.00         | 0.08         | 27.57     | 0.18         | 0.01         |

# Table S20. The expression of select lncRNAs in HUVECs exposed to PCB29-pQ.

|            | 24.68 |      |      | 27.57 |      |      |
|------------|-------|------|------|-------|------|------|
|            | 24.95 |      |      | 25.06 |      |      |
| SLFNL1-AS1 | 24.95 | 1.00 | 0.11 | 25.20 | 1.37 | 0.07 |
|            | 25.22 |      |      | 25.11 |      |      |

|        |           |           | Control              |                 | PCB29-pQ  |                      |                 |  |  |
|--------|-----------|-----------|----------------------|-----------------|-----------|----------------------|-----------------|--|--|
|        |           | Cq values | Mean<br>(normalized) | SD (normalized) | Cq values | Mean<br>(normalized) | SD (normalized) |  |  |
|        |           | 22.29     |                      |                 | 20.63     |                      |                 |  |  |
|        | MIR-7-5p  | 22.46     | 1.00                 | 0.12            | 20.50     | 6.00                 | 0.35            |  |  |
| Figure |           | 22.11     |                      |                 | 20.47     |                      |                 |  |  |
| S3b    |           | 22.09     |                      |                 | 23.05     |                      |                 |  |  |
|        | MIR-24-3p | 21.85     | 1.00                 | 0.09            | 23.32     | 0.72                 | 0.08            |  |  |
|        |           | 21.88     |                      |                 | 23.35     |                      |                 |  |  |
|        |           | 21.89     |                      |                 | 24.81     |                      |                 |  |  |
|        | MIR-9-5p  | 21.79     | 1.00                 | 0.04            | 25.01     | 0.21                 | 0.02            |  |  |
|        |           | 21.89     |                      |                 | 24.94     |                      |                 |  |  |

Mean and SD values are the ones presented in Figure S3a-b. All data reported relative to control treated cells.

### Table S21. The expression of *HDAC7* level in HUVECs transfected with *HDAC7-AS1* siRNA or pEZ-M61-*HDAC7-AS1*.

|            |            | NC siRNA | ł       | HDAC7-ASI siRNA (25 nm) |         |         | HDAC7-A | 4 <i>S1</i> siRNA | (50 nm) | HDAC7-AS1 siRNA (100 nm) |         |          |  |
|------------|------------|----------|---------|-------------------------|---------|---------|---------|-------------------|---------|--------------------------|---------|----------|--|
|            | Cq Mean SD |          | Cq      | Mean                    | SD      | Cq      | Mean    | SD                | Cq      | Mean                     | SD      |          |  |
|            | values     | (normal  | (normal | values                  | (normal | (normal | values  | (normal           | (normal | values                   | (normal | (normali |  |
|            |            | ized)    | ized)   |                         | ized)   | ized)   |         | ized)             | ized)   |                          | ized)   | zed)     |  |
| Figure S4b | 23.74      | 1.00     | 0.03    | 24.89                   | 0.58    | 0.02    | 25.36   | 0.41              | 0.02    | 25.71                    | 0.33    | 0.02     |  |

| 23.66 | 24.79 |  | 25.46 |  | 25.80 |   |
|-------|-------|--|-------|--|-------|---|
| 23.74 | 24.84 |  | 25.32 |  | 25.67 | l |

|            | p           | EZ-M61-N   | ЧС      | pEZ-M61- <i>HDAC7-AS1</i> (2.5 µg) |         |         | pEZ-M6 | 1 <i>-HDAC7</i><br>ug) | -ASI (5 | pEZ-M61 <i>-HDAC7-AS1</i> (7.5<br>µg) |         |          |  |
|------------|-------------|------------|---------|------------------------------------|---------|---------|--------|------------------------|---------|---------------------------------------|---------|----------|--|
|            | Cq          | Cq Mean SD |         | Cq                                 | Mean    | SD      | Cq     | Mean                   | SD      | Cq                                    | Mean    | SD       |  |
|            | values      | (normal    | (normal | values                             | (normal | (normal | values | (normal                | (normal | values                                | (normal | (normali |  |
|            | ized) ized) |            | ized)   | ized) ized)                        |         |         | ized)  | ized)                  |         | ized)                                 | zed)    |          |  |
|            | 23.72       |            |         | 20.77                              |         |         | 19.77  |                        |         | 18.98                                 |         |          |  |
| Figure S4c | 23.58       | 1.00       | 0.05    | 20.94                              | 8.79    | 0.53    | 19.68  | 20.51                  | 1.07    | 19.01                                 | 32.83   | 1.04     |  |
|            | 23.67       |            |         | 20.88                              |         |         | 19.62  |                        |         | 19.07                                 |         |          |  |

|            | NC siRNA |         | A       | NC siRNA + PCB29-pQ |         |         | HDAC7-AS1 siRNA |         |         | HDAC7-AS1 siRNA +<br>PCB29-pQ |         |          |
|------------|----------|---------|---------|---------------------|---------|---------|-----------------|---------|---------|-------------------------------|---------|----------|
|            | Cq       | Mean    | SD      | Cq                  | Mean    | SD      | Cq              | Mean    | SD      | Cq                            | Mean    | SD       |
|            | values   | (normal | (normal | values              | (normal | (normal | values          | (normal | (normal | values                        | (normal | (normali |
|            |          | ized)   | ized)   |                     | ized)   | ized)   |                 | ized)   | ized)   |                               | ized)   | zed)     |
|            | 23.59    |         |         | 23.89               |         |         | 23.71           |         |         | 24.13                         |         |          |
| Figure S4d | 23.69    | 1.00    | 0.06    | 23.68               | 1.10    | 0.09    | 23.48           | 1.31    | 0.10    | 24.09                         | 0.93    | 0.02     |
|            | 23.51    |         |         | 23.85               |         |         | 23.60           |         |         | 24.09                         |         |          |
|            | 24.90    |         |         | 25.35               |         |         | 25.11           |         |         | 25.51                         |         |          |
| Figure S4e | 24.84    | 1.00    | 0.04    | 25.45               | 0.89    | 0.03    | 25.04           | 1.13    | 0.07    | 25.51                         | 0.81    | 0.09     |
|            | 24.95    |         |         | 25.42               |         |         | 25.20           |         |         | 25.79                         |         |          |

Mean and SD values are the ones presented in Figure S4b-e. All data is HDAC1-AS1 mRNA levels reported relative to levels in NC siRNA or

pEZ-M61-NC.

| Table S22. Protein TGF- | <b>B2 and PPME1 levels in HUVECs</b> | exposed to PCB29-nO with MIk | <i>}-7-5p</i> inhibitor or <i>TGF</i> | <i>F-B2/PPME1</i> siRNA. |
|-------------------------|--------------------------------------|------------------------------|---------------------------------------|--------------------------|
|                         |                                      |                              |                                       |                          |

|                  | NC inhibitor + NC<br>siRNA | NC inhibitor + NC<br>siRNA + PCB29-<br>pQ | NC siRNA +<br><i>MIR-7-5p</i><br>inhibitor | NC siRNA +<br><i>MIR-7-5p</i><br>inhibitor +<br>PCB29-pQ | $MIR-7-5p$ inhibitor + TGF- $\beta$<br>2 siRNA | MIR-7-5p<br>inhibitor + TGF-<br>$\beta 2$ siRNA +<br>PCB29-pQ |
|------------------|----------------------------|---|--|--|--|---|
| Figure S7a       | Grayscale values           | Grayscale values                          | Grayscale values                           | Grayscale values   | Grayscale values                               | Grayscale values  |
| (Lower           | 13486.08                   | 4773.18                                   | 24453.20                                   | 9189.13  | 9331.76  | 6005.81   |
| panel)           | 13486.08                   | 4807.88                                   | 23710.78                                   | 9553.25  | 9037.81  | 6698.18   |
|                  | 13508.49                   | 4833.88                                   | 24159.95                                   | 9516.47  | 9203.93  | 6424.71   |
| Eigung S7h       | Grayscale values           | Grayscale values                          | Grayscale values                           | Grayscale values   | Grayscale values                               | Grayscale values  |
| (Lower           | 12610.66                   | 13479.47                                  | 28440.61                                   | 14589.18   | 5921.18  | 1069.50   |
| (LUWCI<br>nanel) | 12527.00                   | 13590.42                                  | 28704.73                                   | 13716.52   | 6436.83  | 1139.45   |
|                  | 12510.71                   | 13746.54                                  | 27936.61                                   | 14791.59   | 6066.13  | 1023.26   |

| Figure S7a<br>(Lower | NC inhibit<br>siRN | tor + NC<br>NA | NC inhibit<br>siRNA + 1<br>pQ | or + NC<br>PCB29- | NC siF<br><i>MIR-</i><br>inhit | RNA +<br>7-5p<br>bitor | NC siF<br><i>MIR-</i><br>inhibi<br>PCB2 | RNA +<br>7-5p<br>tor +<br>9-pQ | <i>MIR-</i><br>inhibitor<br>2 siR | 7-5p<br>+ TGF-β<br>NA | <i>MIR-</i><br>inhibitor<br>β2 siR<br>PCB2 | 7-5p<br>+ TGF-<br>NA +<br>9-pQ |
|----------------------|--------------------|----------------|-------------------------------|-------------------|--------------------------------|------------------------|---|--------------------------------|-----------------------------------|-----------------------|--|--------------------------------|
| panel)               | Mean               | SD             | Mean                          | SD                | Mean                           | SD                     | Mean                                    | SD                             | Mean                              | SD                    | Mean                                       | SD                             |
|                      | (normaliz          | (norma         | (normaliz                     | (norma            | (normali                       | (norma                 | (normali                                | (norma                         | (normali                          | (norma                | (normali                                   | (norma                         |
|                      | ed)                | lized)         | ed)                           | lized)            | zed)                           | lized)                 | zed)                                    | lized)                         | zed)                              | lized)                | zed)                                       | lized)                         |
|                      | 1.00               | 0.00           | 0.40                          | 0.00              | 2.02                           | 0.03                   | 0.96                                    | 0.02                           | 0.80                              | 0.01                  | 0.65                                       | 0.04                           |

| Figure S7b<br>(Lower | NC inhibitor + NC<br>siRNA |        | NC inhibitor + NC<br>siRNA + PCB29-<br>pQ |        | NC siRNA +<br><i>MIR-7-5p</i><br>inhibitor |        | NC siRNA +<br>MIR-7-5p<br>inhibitor +<br>PCB29-pQ |        | <i>MIR-7-5p</i><br>inhibitor +<br><i>PPME1</i> siRNA |        | <i>MIR-7-5p</i><br>inhibitor +<br><i>PPME1</i> siRNA +<br>PCB29-pQ |        |
|----------------------|----------------------------|--------|---|--------|--|--------|---|--------|--|--------|--|--------|
| panel)               | Mean                       | SD     | Mean                                      | SD     | Mean                                       | SD     | Mean  | SD     | Mean   | SD     | Mean   | SD     |
|                      | (normaliz                  | (norma | (normaliz                                 | (norma | (normali                                   | (norma | (normali  | (norma | (normali   | (norma | (normali   | (norma |
|                      | ed)                        | lized) | ed)                                       | lized) | zed)                                       | lized) | zed)  | lized) | zed)   | lized) | zed)   | lized) |
|                      | 1.00                       | 0.00   | 0.83                                      | 0.04   | 2.80                                       | 0.04   | 1.57  | 0.06   | 0.72   | 0.03   | 0.10   | 0.01   |

Mean and SD values are the ones presented in Figure S7a-b (Lower panel). All data is presented as fold difference relative to NC inhibitor + NC

siRNA.

Table S23. Luciferase analysis of the activity of MIR-7-5p bind to TGF-β2, PPME1, and HDAC7-AS1 in HUVECs transfected with NC

#### mimic or *MIR-7-5p* mimic.

|        |                               | NC m                | nimic     |          | MIR-7-5p mimic      |           |            |  |  |
|--------|-------------------------------|---------------------|-----------|----------|---------------------|-----------|------------|--|--|
|        |                               |                     | Mean      | SD       |                     | Mean      | SD         |  |  |
|        |                               | luciferase activity | (normaliz | (normali | luciferase activity | (normaliz | (normalize |  |  |
|        |                               |                     | ed)       | zed)     |                     | ed)       | d)         |  |  |
|        |                               | 2.95                |           |          | 3.07                |           |            |  |  |
| Figure | NC-3'UTR                      | 2.90                | 1.00      | 0.05     | 3.07                | 1.11      | 0.05       |  |  |
| S8a    |                               | 2.67                |           |          | 3.32                |           |            |  |  |
|        |                               | 0.55                |           |          | 0.48                |           |            |  |  |
|        | <i>TGF-β2</i> (WT)- 3'UTR     | 0.57                | 1.00      | 0.07     | 0.51                | 0.86      | 0.03       |  |  |
|        |                               | 0.63                |           |          | 0.51                |           |            |  |  |
|        | $TCE \rho_2$ (MUT) 2'UTD      | 0.16                | 1.00      | 0.00     | 0.18                | 1.02      | 0.07       |  |  |
|        | $101^{-}p^{2}$ (10101)- 5 01K | 0.16                | 1.00      | 0.08     | 0.16                | 1.05      | 0.07       |  |  |

| 0.10 |  | 0.18 |  |  | 0.18 |  |  |
|------|--|------|--|--|------|--|--|
|------|--|------|--|--|------|--|--|

|        |                    | NC m                | nimic |      | MIR-7-5             | <i>p</i> mimic |      |  |
|--------|--------------------|---------------------|-------|------|---------------------|----------------|------|--|
|        |                    | luciferase activity | Mean  | SD   | luciferase activity | Mean           | SD   |  |
|        |                    | 3.00                |       |      | 3.35                |                |      |  |
|        | NC-3'UTR           | 3.03                | 1.00  | 0.01 | 3.35                | 1.12           | 0.02 |  |
| Figure |                    | 3.02                |       |      | 3.44                |                |      |  |
| S8b    |                    | 0.67                |       |      | 0.50                |                |      |  |
|        | PPME1 (WT)- 3'UTR  | 0.66                | 1.00  | 0.02 | 0.49                | 0.72           | 0.02 |  |
|        |                    | 0.69                |       |      | 0.47                |                |      |  |
|        |                    | 0.19                |       |      | 0.23                |                |      |  |
|        | PPME1 (MUT)- 3'UTR | 0.19                | 1.00  | 0.00 | 0.23                | 1.19           | 0.00 |  |
|        |                    | 0.19                |       |      | 0.23                |                |      |  |

|        |               | NC m                | imic |      | MIR-7-5             | 5 <i>p</i> mimic |      |
|--------|---------------|---------------------|------|------|---------------------|------------------|------|
|        |               | luciferase activity | Mean | SD   | luciferase activity | Mean             | SD   |
|        |               | 3.03                |      |      | 2.56                |                  |      |
|        | NC            | 2.73                | 1.00 | 0.08 | 2.53                | 0.96             | 0.07 |
| Eigung |               | 2.58                |      |      | 2.89                |                  |      |
| Figure |               | 0.65                |      |      | 0.32                |                  |      |
| 300    | HDAC7-AS1-WT  | 0.57                | 1.00 | 0.07 | 0.35                | 0.52             | 0.03 |
|        |               | 0.65                |      |      | 0.31                |                  |      |
|        |               | 0.16                |      |      | 0.16                |                  |      |
|        | HDAC7-AS1-MUT | 0.15                | 1.00 | 0.10 | 0.31                | 1.34             | 0.67 |
|        |               | 0.13                |      |      | 0.12                |                  |      |

Mean and SD values are the ones presented in Figure S8a-c. All data presented as fold difference relative to NC mimic. All data is expressed as

luciferase activity (a.u.).

|           |                           | (                 | Control |       | PCB2              | 9-pQ   |       |
|-----------|---------------------------|-------------------|---------|-------|-------------------|--------|-------|
|           |                           | Individual values | Mean    | SD    | Individual values | Mean   | SD    |
| Eigen CO  |                           | 188.48            |         |       | 103.11            |        |       |
| Figure S9 | NC inhibitor              | 186.73            | 186.50  | 2.12  | 102.49            | 102.00 | 1.41  |
| (Kight    |                           | 184.29            |         |       | 100.41            |        |       |
| paner)    |                           | 208.41            |         |       | 186.33            |        |       |
|           | <i>MIR-7-5p</i> inhibitor | 220.68            | 220.50  | 12.02 | 160.12            | 167.00 | 16.97 |
|           |                           | 232.41            |         |       | 154.55            |        |       |

Table S24. The tube forming ability in HUVECs exposed to PCB29-pQ and transfected with a *MIR-7-5p* inhibitor.

Mean and SD values are the ones presented in Figure S9 (Right panel). Number of branch points is presented.

#### Table S25. HDAC7-AS1, MIR-7-5p, TGF-β2, PPME1 levels and apoptosis or proliferation rates in HUVECs exposed to PCB29-pQ and

transfected with a *MIR-7-5p* inhibitor or *HDAC7-AS1* siRNA.

|             | NC inh | ibitor + N | C siRNA | NC inhib | itor + NC s       | siRNA + | NC inhib | itor + $HD_{A}$ | 1 <i>C7-AS1</i> | MIR-7-5p inhibitor + |          |          |
|-------------|--------|------------|---------|----------|-------------------|---------|----------|-----------------|-----------------|----------------------|----------|----------|
|             |        |            |         | F        | PCB29-pO          |         | siRN     | A + PCB2        | 9-nO            | HDAC/-AST siRNA +    |          |          |
|             |        |            |         | -        | 55 <b>-</b> ) P C |         | birti    | 1002            | Ρ₹              |                      | PCB29-pQ |          |
|             | Cq     | Mean       | SD      | Cq       | Mean              | SD      | Cq       | Mean            | SD              | Cq                   | Mean     | SD       |
|             | values | (normal    | (normal | values   | (normal           | (normal | values   | (normal         | (normal         | values               | (normal  | (normali |
|             |        | ized)      | ized)   |          | ized)             | ized)   |          | ized)           | ized)           |                      | ized)    | zed)     |
|             | 24.00  |            |         | 22.95    |                   |         | 20.55    |                 |                 | 23.43                |          |          |
| Figure S11a | 24.03  | 1.00       | 0.02    | 22.90    | 2.07              | 0.05    | 20.35    | 3.50            | 0.25            | 22.51                | 1.82     | 0.60     |
|             | 23.97  |            |         | 22.88    |                   |         | 20.40    |                 |                 | 23.06                |          |          |
|             | 24.06  |            |         | 24.90    |                   |         | 25.23    |                 |                 | 26.81                |          |          |
| Figure S11b | 24.11  | 1.04       | 0.04    | 24.75    | 0.56              | 0.04    | 25.23    | 0.13            | 0.01            | 26.51                | 0.14     | 0.02     |
|             | 23.99  |            |         | 24.90    |                   |         | 25.12    |                 |                 | 26.81                |          |          |
| Figure S11c | 24.04  | 1.00       | 0.06    | 24.96    | 0.57              | 0.09    | 24.26    | 0.28            | 0.02            | 24.48                | 0.73     | 0.12     |

|             | 24.11 |      |      | 24.56 |      |      | 23.99 |      |      | 24.05 |      |      |
|-------------|-------|------|------|-------|------|------|-------|------|------|-------|------|------|
|             | 23.93 |      |      | 24.93 |      |      | 24.04 |      |      | 24.41 |      |      |
|             | 23.83 |      |      | 24.32 |      |      | 23.28 |      |      | 23.89 |      |      |
| Figure S11d | 23.95 | 1.00 | 0.06 | 24.32 | 0.72 | 0.04 | 23.05 | 0.49 | 0.05 | 23.77 | 0.90 | 0.08 |
|             | 24.00 |      |      | 24.44 |      |      | 23.28 |      |      | 24.02 |      |      |

|             | NC inhibitor + NC siRNA |         |         | NC inhibitor + NC siRNA +<br>PCB29-pQ |         |         | NC inhibitor + <i>HDAC7-AS1</i><br>siRNA + PCB29-pQ |         |         | MIR-7-5p inhibitor +<br>HDAC7-AS1 siRNA +<br>PCB29-pO |         |          |
|-------------|-------------------------|---------|---------|---------------------------------------|---------|---------|---|---------|---------|---|---------|----------|
|             | Graysc                  | Mean    | SD      | Grayscal                              | Mean    | SD      | Grayscal  | Mean    | SD      | Grays   | Mean    | SD       |
|             | ale                     | (normal | (normal | e values                              | (normal | (normal | e values  | (normal | (normal | cale  | (normal | (normali |
|             | values                  | ized)   | ized)   |                                       | ized)   | ized)   |   | ized)   | ized)   | values  | ized)   | zed)     |
| Figure S11f | 26471.                  |         |         |                                       |         | 0.04    |   |         |         | 24470   | 0.61    | 0.04     |
|             | 34                      | 1.00    | 0.03    | 19404.29                              |         |         | 8350.731  |         | 0.02    | .19   |         |          |
|             | 25571.                  |         |         |                                       | 0.75    |         |   | 0.22    |         | 23380   |         |          |
|             | 42                      |         |         | 18907.89                              |         |         | 7680.765  | 0.32    |         | .91   |         |          |
|             | 24789.                  |         |         |                                       |         |         |   |         |         | 21457   |         |          |
|             | 01                      |         |         | 17650.91                              |         |         | 7690.97   |         |         | .98   |         |          |
|             | 35445.                  |         |         |                                       |         |         |   |         |         | 31926   | 0.61    | 0.00     |
|             | 87                      |         |         | 29143.95                              |         |         | 14331.22  |         |         | .00   |         |          |
|             | 35402.                  | 1.00    | 0.00    |                                       | 0.04    | 0.00    |   | 0.42    | 0.00    | 32072   |         |          |
| Figure SIIg | 05                      | 1.00    | 0.00    | 28836.87                              | 0.84    | 0.00    | 14263.22  |         |         | .82   |         |          |
|             | 35530.                  |         |         |                                       |         |         |   |         |         | 31968   |         |          |
|             | 12                      |         |         | 28979.22                              |         |         | 14351.63  |         |         | .29   |         |          |

|             | NC inhibitor + NC siRNA |         |         | NC inhibitor + NC siRNA +<br>PCB29-pQ |         |         | NC inhibitor + <i>HDAC7-AS1</i><br>siRNA + PCB29-pQ |         |         | MIR-7-5p inhibitor +<br>HDAC7-AS1 siRNA +<br>PCB29-pQ |         |          |
|-------------|-------------------------|---------|---------|---------------------------------------|---------|---------|---|---------|---------|---|---------|----------|
|             | Individ                 | Mean    | SD      | Individu                              | Mean    | SD      | Individu  | Mean    | SD      | Indivi  | Mean    | SD       |
|             | ual                     | (normal | (normal | al values                             | (normal | (normal | al values   | (normal | (normal | dual  | (normal | (normali |
|             | values                  | ized)   | ized)   |                                       | ized)   | ized)   |   | ized)   | ized)   | values  | ized)   | zed)     |
| Figure S11h | 100.92                  | 100.00  | 1.91    | 79.83                                 | 81.03   | 3.32    | 76.57   | 71.42   | 4.65    | 87.85   | 88.99   | 1.81     |

| 97.81  | 84.78 |  | 67.54 |  | 91.08 |  |
|--------|-------|--|-------|--|-------|--|
| 101.27 | 78.47 |  | 70.15 |  | 88.04 |  |

Mean and SD values are the ones presented in Figure S11a (mRNA expression of *MIR-7-5p*), b (mRNA expression of *HDAC7-AS1*), c (mRNA expression of *TGFβ-2*), and d (mRNA expression of *PPME1*), and f (protein expression of TGF- $\beta$ 2), g (protein expression of PPME1), and h (% cell viability). All data presented relative to NC inhibitor + NC siRNA.

Table S26. HE staining of aortic root cross-sections and total cholesterol (TC) and low-density lipoprotein (LDL) levels in ApoE<sup>-/-</sup> mice

|               | AAV-NC            |           | AAV-NC            | + PCB29-pQ | AAV-HDA           | 1 <i>C7-AS1</i> | AAV- <i>HDAC7-AS1</i> +<br>PCB29-pQ |       |  |
|---------------|-------------------|-----------|-------------------|------------|-------------------|-----------------|-------------------------------------|-------|--|
|               | Mean              | SD        | Mean              | SD         | Mean              | SD              | Mean                                | SD    |  |
|               | 14.35             | 0.81      | 26.81             | 3.78       | 9.71              | 1.62            | 14.17                               | 2.52  |  |
| Eiguna C12a   | Individu          | al values | Individ           | ual values | Individua         | l values        | Individual values                   |       |  |
| (Dight nonal) | 13.               | .49       | 22                | 2.44       | 11.1              | 8               | 17.05                               |       |  |
| (Right panel) | 14.               | .46       | 29                | 9.09       | 9.9               | 6               | 13.09                               |       |  |
|               | 15.               | .09       | 23                | 8.89       | 7.9               | 8               | 12.38                               |       |  |
|               | 31.60             | 6.03      | 70.40             | 11.89      | 32.20             | 9.83            | 47.60                               | 9.32  |  |
|               | Individual values |           | Individual values |            | Individual values |                 | Individual values                   |       |  |
|               | 36                |           | 68                |            | 26                |                 | 43                                  |       |  |
| Figure S12b   | 3                 | 39        |                   | 69         |                   | 34              |                                     | 41    |  |
|               | 2                 | 8         | 74                |            | 47                |                 | 55                                  |       |  |
|               | 2                 | 24        |                   | 87         |                   |                 | 60                                  |       |  |
|               | 3                 | 31        |                   | 54         |                   | 33              |                                     | 39    |  |
| Figure S12c   | 246.6             | 58.22     | 336.6             | 42.74      | 165.2             | 40.77           | 231.4                               | 46.98 |  |

were *i.v.* injected with AAV-HDAC7-AS1.

| Individual values | Individual values | Individual values | Individual values |
|-------------------|-------------------|-------------------|-------------------|
| 267               | 310               | 211               | 187               |
| 210               | 330               | 103               | 198               |
| 201               | 352               | 190               | 208               |
| 340               | 401               | 165               | 288               |
| 215               | 290               | 157               | 276               |

Mean and SD values are the ones presented in Figure S12a (Right panel) and Figure S12a (plaque area relative to aortic section), b (TC; mmol/l),

and c (LDL; mmol/l).

|             | 0 h    |         |           | 1 h      |             |         | 3 h      |         |         | 6 h    |         |          |      |
|-------------|--------|---------|-----------|----------|-------------|---------|----------|---------|---------|--------|---------|----------|------|
|             | Graysc | Mean    | SD        | Grayscal | Mean        | SD      | Grayscal | Mean    | SD      | Grays  | Mean    | SD       |      |
|             | ale    | (normal | (normal   | e values | (normal     | (normal | e values | (normal | (normal | cale   | (normal | (normali |      |
|             | values | ized)   | ized)     |          | ized)       | ized)   |          | ized)   | ized)   | values | ized)   | zed)     |      |
|             | 3549.9 |         |           |          |             |         |          |         |         | 41060  |         |          |      |
|             | 1      |         |           | 14419.75 |             |         | 21891.17 |         |         | .31    |         |          |      |
| Figure S13a | 3551.9 | 1.00    | 1.00 0.01 | 0.01     |             | 2.62    | 0.01     |         | 6 12    | 0.06   | 40411   | 12 76    | 0.05 |
| (p-CAV1)    | 1      | 1.00    | 0.01      | 14458.87 | 7 5.05<br>0 | 0.01    | 22463.19 | 0.45    | 0.00    | .36    | 12.70   | 0.03     |      |
|             | 3544.5 |         |           |          |             |         |          |         |         | 40784  |         |          |      |
|             | 0      |         |           | 14511.00 |             |         | 22130.58 |         |         | .65    |         |          |      |
|             | 35207. |         |           |          |             |         |          |         |         | 31824  |         |          |      |
|             | 65     |         |           | 39533.43 |             |         | 34135.36 |         |         | .58    |         |          |      |
| Figure S13a | 35245. |         |           |          |             |         |          |         |         | 31567  |         |          |      |
| (CAV1)      | 36     |         |           | 39410.19 |             |         | 34430.60 |         |         | .80    |         |          |      |
|             | 35224. |         |           |          |             |         |          |         |         | 31747  |         |          |      |
|             | 65     |         |           | 39808.55 |             |         | 34124.48 |         |         | .80    |         |          |      |

Table S27. TGF-β2, PPME1, and inflammatory factors level in HUVECs exposed to PCB29-pQ with or without CAV1 siRNA.
|             | NC siRNA     |         | A       | NC siRNA + PCB29-pQ |         |         | CAV1 siRNA |         |         | <i>CAV1</i> siRNA + PCB29-pQ |         |          |
|-------------|--------------|---------|---------|---------------------|---------|---------|------------|---------|---------|------------------------------|---------|----------|
|             | Graysc       | Mean    | SD      | Grayscal            | Mean    | SD      | Grayscal   | Mean    | SD      | Grays                        | Mean    | SD       |
|             | ale          | (normal | (normal | e values            | (normal | (normal | e values   | (normal | (normal | cale                         | (normal | (normali |
|             | values       | ized)   | ized)   |                     | ized)   | ized)   |            | ized)   | ized)   | values                       | ized)   | zed)     |
|             | 20472.       |         |         |                     |         |         |            |         |         | 6454.                        |         |          |
|             | 92           |         |         | 29074.26            |         |         | 8883.024   |         |         | 782                          |         |          |
| Figure S13c | 20411.       | 1.05    | 0.00    |                     | 1.65    | 0.01    |            | 0.42    | 0.01    | 6442.                        | 0.32    | 0.01     |
| (p-CAV1)    | 51           | 1.05    | 0.00    | 29425.33            | 1.05    | 0.01    | 8531.782   | 0.42    | 0.01    | 782                          | 0.52    | 0.01     |
|             | 20471.       |         |         |                     |         |         |            |         |         | 6219.                        |         |          |
|             | 34           |         |         | 29228.50            |         |         | 8789.317   |         |         | 468                          |         |          |
|             | 36333.       |         |         |                     |         |         |            |         |         | 10561                        |         |          |
|             | 41           |         |         | 38767.5             |         |         | 16916.46   | -       |         | .56                          |         |          |
| Figure S13c | 36998.       |         |         |                     |         |         |            |         |         | 10408                        |         |          |
| (CAV1)      | 02           |         |         | 38129.48            |         |         | 17327.41   | -       |         | .2                           |         |          |
|             | 36895.       |         |         |                     |         |         |            |         |         | 10447                        |         |          |
|             | 19           |         |         | 38340.43            |         |         | 17230.58   |         |         | .02                          |         |          |
|             | 24553.       |         |         | 10000 54            |         |         | 24677.02   |         |         | 31960                        |         |          |
|             | 22           |         |         | 12233.54            |         |         | 34677.92   |         |         | .07                          |         |          |
| Figure S13d | 24553.       | 1.00    | 0.00    | 10000 54            | 0.55    | 0.01    | 24610.22   | 1.33    | 0.01    | 31996                        | 1.29    | 0.00     |
| U U         | 22           |         |         | 12233.54            |         |         | 34618.22   | -       |         | .0/                          |         |          |
|             | 24478.       |         |         | 10000 54            |         |         | 24202 20   |         |         | 32146                        |         |          |
|             | 9/           |         |         | 12233.34            |         |         | 34383.39   |         |         | .//                          |         |          |
|             | 28510.       |         |         | 20444 51            |         |         | 27711 20   |         |         | 21134                        |         |          |
|             | 29           |         |         | 20444.31            |         |         | 2//11.29   |         |         | .0/                          |         |          |
| Figure S13e | 27909.<br>62 | 1.00    | 0.01    | 20778 41            | 0.80    | 0.01    | 27671 17   | 0.92    | 0.00    | 21252                        | 0.74    | 0.00     |
|             | 28526        |         |         | 20776.41            |         |         | 2/0/1.1/   | -       |         | .24                          |         |          |
|             | 20520.       |         |         | 20288-1             |         |         | 27720 41   |         |         | 21147                        |         |          |
|             | 6365.9       |         |         | 20200.1             |         |         | 27720.71   |         |         | 8396                         |         |          |
|             | 74           |         |         | 29456 14            |         |         | 5738 782   |         |         | 974                          |         |          |
| Figure S13f | 6527.3       | 1.00    | 0.02    |                     | 5.12    | 0.12    | 5750.702   | 0.85    | 0.02    | 8674                         | 1.30    | 0.03     |
|             | 17           |         |         | 30621.72            |         |         | 6009.317   |         |         | 924                          |         |          |

|              | 6636.3 |      |      |          |      |      |          |      |      | 8691. |      |      |
|--------------|--------|------|------|----------|------|------|----------|------|------|-------|------|------|
|              | 88     |      |      | 30772.26 |      |      | 5790.782 |      |      | 338   |      |      |
|              | 12188. |      |      |          |      |      |          |      |      | 2204. |      |      |
|              | 66     |      |      | 32999.77 |      |      | 21290.05 |      |      | 359   |      |      |
| Eigung S12g  | 12851. | 1.00 | 0.02 |          | 2 17 | 0.22 |          | 1.62 | 0.02 | 2515. | 0.10 | 0.01 |
| Figure 515g  | 49     | 1.00 | 0.05 | 37559.54 | 3.17 | 0.25 | 21818.55 | 1.05 | 0.02 | 648   | 0.19 | 0.01 |
|              | 12532. |      |      |          |      |      |          |      |      | 2496. |      |      |
|              | 95     |      |      | 37660.01 |      |      | 21875.55 |      |      | 406   |      |      |
|              | 24172. |      |      |          |      |      |          |      |      | 28610 |      |      |
|              | 53     |      |      | 35359.97 |      |      | 23666.02 |      |      | .48   |      |      |
| Eiguro S12h  | 24116. | 1.00 | 0.00 |          | 1.61 | 0.00 |          | 0.02 | 0.00 | 28470 | 1 16 | 0.00 |
| Figure S1511 | 24     | 1.00 | 0.00 | 35383.38 | 1.01 | 0.00 | 23714.26 | 0.92 | 0.00 | .24   | 1.10 | 0.00 |
|              | 24332. |      |      |          |      |      |          |      |      | 28493 |      |      |
|              | 89     |      |      | 35253.43 |      |      | 23805.09 |      |      | .41   |      |      |

Mean and SD values are the ones presented in Figure S13a (Lower panel; p-CAV1/CAV1 ratio after treatment with PCB29-pQ for 0, 1, 3, or 6h,

no *CAV1* siRNA) and Figure c (p-CAV/CAV ratio), d (TGF-β2), e (PPME1), f (IL-1β), g (IL-6), and h (TNF-α). Data for c-h were normalized to

 $\beta$ -actin expression and presented as fold difference compared to NC siRNA.

|                                | AAV-NC                   |      |      | AAV-NC + PCB29-pQ     |      |      | AAV-HDAC7-AS1         |      |      | AAV- <i>HDAC7-AS1</i> +<br>PCB29-pQ |      |      |
|--------------------------------|--------------------------|------|------|-----------------------|------|------|-----------------------|------|------|-------------------------------------|------|------|
|                                | Individ<br>ual<br>values | Mean | SD   | Individu<br>al values | Mean | SD   | Individu<br>al values | Mean | SD   | Indivi<br>dual<br>values            | Mean | SD   |
| Figure S14<br>(Right<br>panel) | 0.35<br>0.47<br>0.48     | 0.43 | 0.07 | 0.92<br>0.98<br>0.90  | 0.93 | 0.04 | 0.02<br>0.01<br>0.05  | 0.02 | 0.02 | 0.17<br>0.38<br>0.38                | 0.31 | 0.12 |

Table S28. Pearson's correlation coefficient for expression of p-CAV1 and CD31 in ApoE<sup>-/-</sup> mice that were i.v. injected with AAV-HDAC7-AS1.

| 12943. |          |  |          |  | 37683 |  |
|--------|----------|--|----------|--|-------|--|
| 66     | 20477.39 |  | 28475.48 |  | .36   |  |
| 12634. |          |  |          |  | 37516 |  |
| 66     | 20457.39 |  | 28444.24 |  | .89   |  |

Mean and SD values are the ones presented in Figure S14 (Right panel).

## Table S29. Immunohistochemistry staining of CD68, total cholesterol (TC), and low-density lipoprotein level (LDL) in ApoE<sup>-/-</sup> and ApoE<sup>-</sup>

/-/*CAV1*-/- mice.

|               |          |                   | ApoE-/- |      | ApoE <sup>-/-</sup> /CAV1 <sup>-/-</sup> |       |      |  |  |
|---------------|----------|-------------------|---------|------|--|-------|------|--|--|
|               |          | Individual values | Mean    | SD   | Individual values                        | Mean  | SD   |  |  |
|               |          | 19.85             |         |      | 6.33                                     |       |      |  |  |
| Figure S16a   | Control  | 14.40             | 16.00   | 3.35 | 6.94                                     | 7.88  | 2.17 |  |  |
| (Right panel) |          | 13.75             |         |      | 10.35                                    |       |      |  |  |
|               |          | 21.76             |         |      | 15.48                                    |       |      |  |  |
|               | PCB29-pQ | 22.45             | 20.86   | 2.19 | 11.70                                    | 12.38 | 2.83 |  |  |
|               |          | 18.36             |         |      | 9.95                                     |       |      |  |  |

|             |          |                   | ApoE-/- |       | A                 | poE <sup>-/-</sup> /CAV1 <sup>-/-</sup> |      |
|-------------|----------|-------------------|---------|-------|-------------------|---|------|
|             |          | Individual values | Mean    | SD    | Individual values | Mean                                    | SD   |
|             |          | 39.00             |         |       | 25.00             |   |      |
|             |          | 32.00             |         |       | 39.00             |   |      |
| Eigung S16h | Control  | 29.00             | 31.80   | 6.22  | 32.00             | 33.80                                   | 5.63 |
| rigule 5100 |          | 23.00             |         |       | 38.00             |   |      |
|             |          | 36.00             |         |       | 35.00             |   |      |
|             |          | 69.00             |         |       | 42.00             |   |      |
|             | PCB29-pQ | 74.00             | 71.80   | 10.57 | 39.00             | 46.20                                   | 6.14 |
|             |          | 85.00             |         |       | 45.00             |   |      |

| 56.00 |  | 52.00 |  |
|-------|--|-------|--|
| 75.00 |  | 53.00 |  |

|             |          |                   | ApoE-/- |       | A                 | poE <sup>-/-</sup> /CAV1 <sup>-/-</sup> |        |
|-------------|----------|-------------------|---------|-------|-------------------|---|--------|
|             |          | Individual values | Mean    | SD    | Individual values | Mean                                    | SD     |
|             |          | 268.00            |         |       | 301.00            |   |        |
|             |          | 126.00            |         |       | 301.00            |   |        |
|             | Control  | 206.00            | 207.40  | 77.72 | 206.00            | 275.60                                  | 40.937 |
| Figure S16a |          | 136.00            |         |       | 299.00            |   |        |
| Figure STOC |          | 301.00            |         |       | 271.00            |   |        |
|             |          | 543.00            |         |       | 405.00            |   |        |
|             |          | 571.00            |         |       | 321.00            |   |        |
|             | PCB29-pQ | 432.00            | 510.20  | 52.47 | 401.00            | 396.80                                  | 44.59  |
|             |          | 501.00            |         |       | 436.00            |   |        |
|             |          | 504.00            |         |       | 421.00            |   |        |

Mean and SD values are the ones presented in Figure S16a (Right panel;), b (TC, mmol/l), and c (LDL, mmol/l).

| Table <b>S</b> | <b>S30</b> . | Adhesion | molecules, | pro-inflan | nmatory | cytokines | and p65 | ' protein ex | xpression le | evel in H | UVEC ex | posed to | PCB29- | pQ. |
|----------------|--------------|----------|------------|------------|---------|-----------|---------|--------------|--------------|-----------|---------|----------|--------|-----|
|                |              |          |            |            |         |           |         |              |              |           |         |          |        |     |

|             |        | 0 h     |         | 1 h      |         |         | 3 h      |         |         | 6 h    |         |          |
|-------------|--------|---------|---------|----------|---------|---------|----------|---------|---------|--------|---------|----------|
|             | Graysc | Mean    | SD      | Grayscal | Mean    | SD      | Grayscal | Mean    | SD      | Grays  | Mean    | SD       |
|             | ale    | (normal | (normal | e values | (normal | (normal | e values | (normal | (normal | cale   | (normal | (normali |
|             | values | ized)   | ized)   |          | ized)   | ized)   |          | ized)   | ized)   | values | ized)   | zed)     |
|             | 21710. |         |         |          |         |         |          |         |         | 34571  |         |          |
|             | 31     |         |         | 32831.89 |         |         | 35262.31 |         |         | .68    |         |          |
| Figure S17h | 22002. | 1.00    | 0.02    |          | 1 27    | 0.00    |          | 1.52    | 0.01    | 34389  | 1 50    | 0.02     |
| Figure 5170 | 72     | 1.00    | 0.02    | 33634.02 | 1.57    | 0.00    | 35212.19 | 1.55    | 0.01    | .10    | 1.30    | 0.02     |
|             | 22448. |         |         |          |         |         |          |         |         | 34622  |         |          |
|             | 72     |         |         | 33584.48 |         |         | 34742.26 |         |         | .68    |         |          |

|             | 5039.5<br>4  |      |      | 18047.31 |      |      | 26252.14 |      |      | 37682<br>.02 |      |      |
|-------------|--------------|------|------|----------|------|------|----------|------|------|--------------|------|------|
| Figure S17c | 5142.3<br>7  | 1.00 | 0.01 | 17668.19 | 3.63 | 0.04 | 26381.60 | 5.05 | 0.16 | 38094<br>.14 | 6.42 | 0.13 |
|             | 5138.3<br>7  |      |      | 17703.19 |      |      | 27830.45 |      |      | 36646<br>.00 |      |      |
|             | 13157.<br>73 |      |      | 29704.43 |      |      | 31683.31 |      |      | 41349<br>.53 |      |      |
| Figure S17d | 13400.<br>10 | 1.00 | 0.03 | 29558.07 | 2.31 | 0.02 | 31253.95 | 2.39 | 0.04 | 41482<br>.58 | 2.76 | 0.02 |
|             | 12635.<br>20 |      |      | 30513.92 |      |      | 31385.89 |      |      | 40789<br>.29 |      |      |
|             | 16167.<br>24 |      |      | 17999.00 |      |      | 26253.14 |      |      | 32660<br>.29 |      |      |
| Figure S17e | 16070.<br>17 | 1.00 | 0.02 | 18002.41 | 1.19 | 0.01 | 26429.67 | 1.59 | 0.01 | 32522<br>.17 | 1.79 | 0.00 |
|             | 15514.<br>68 |      |      | 18385.89 |      |      | 26264.26 |      |      | 32662<br>.29 |      |      |
|             | 9014.2<br>2  |      |      | 20417.84 |      |      | 24193.53 |      |      | 35338        |      |      |
| Figure S17f | 8564.2<br>0  | 1.00 | 0.03 | 19950.19 | 2.39 | 0.04 | 24735.14 | 2.63 | 0.04 | 35652        | 3.48 | 0.03 |
|             | 9080.6       |      |      | 20655.31 |      |      | 24087.82 |      |      | 34981<br>.51 |      |      |
|             | 12851.<br>66 |      |      | 20475.39 |      |      | 28455.95 |      |      | 3/633        |      |      |
| Figure S17i | 12943.<br>66 | 1.00 | 0.01 | 20477.39 | 1.69 | 0.06 | 28475.48 | 2.47 | 0.06 | 3/683<br>.36 | 3.40 | 0.01 |
|             | 12634.<br>66 |      |      | 20457.39 |      |      | 28444.24 |      |      | 3/516<br>.89 |      |      |
| Figure S17j | 31806.<br>92 | 1.00 | 0.00 | 25913.87 | 0.84 | 0.00 | 11533.10 | 0.40 | 0.02 | 11508<br>.73 | 0.39 | 0.00 |

| 31920. |          |          | 11025 |  |
|--------|----------|----------|-------|--|
| 92     | 25681.92 | 11297.73 | .27   |  |
| 31736. |          |          | 10419 |  |
| 92     | 25889.75 | 11409.44 | .42   |  |

|                    |            | Control      |              | PCB29-pQ          |              |              |  |
|--------------------|------------|--------------|--------------|-------------------|--------------|--------------|--|
|                    | Individual | Mean         | SD           | Individual values | Mean         | SD           |  |
| Figure S17g (Right | values     | (normalized) | (normalized) |                   | (normalized) | (normalized) |  |
| panel)             | 6.15       |              |              | 17.99             |              |              |  |
|                    | 4.89       | 4.94         | 1.18         | 15.26             | 15.27        | 2.71         |  |
|                    | 3.79       |              |              | 12.56             |              |              |  |

|             |        | Control |         | P        | PCB29-pQ |         |          | PDTC    |         | PD     | TC + PCB | 29-pQ    |
|-------------|--------|---------|---------|----------|----------|---------|----------|---------|---------|--------|----------|----------|
|             | Graysc | Mean    | SD      | Grayscal | Mean     | SD      | Grayscal | Mean    | SD      | Grays  | Mean     | SD       |
|             | ale    | (normal | (normal | e values | (normal  | (normal | e values | (normal | (normal | cale   | (normal  | (normali |
|             | values | ized)   | ized)   |          | ized)    | ized)   |          | ized)   | ized)   | values | ized)    | zed)     |
|             | 24902. |         |         |          |          |         |          |         |         | 33129  |          |          |
|             | 37     |         |         | 37563.55 |          |         | 25411.1  |         |         | .51    |          |          |
| Eiguro S171 | 24895. | 1.00    | 0.00    |          | 1.57     | 0.00    |          | 1 20    | 0.00    | 33175  | 1 20     | 0.00     |
| Figure S1/I | 37     | 1.00    |         | 37531.84 |          | 0.00    | 25381.39 | 1.29    | 0.00    | .92    | 1.30     | 0.00     |
|             | 24917. |         |         |          |          |         |          |         |         | 33020  |          |          |
|             | 2      |         |         | 37384.19 |          |         | 25371.63 |         |         | .92    |          |          |
|             | 7863.1 |         |         |          |          |         |          |         |         | 10033  |          |          |
|             | 96     |         |         | 28066.72 |          |         | 12419.56 |         |         | .95    |          |          |
| Figure S17m | 7853.7 | 1.00    | 0.00    |          | 2 72     | 0.00    |          | 2.00    | 0.00    | 9969.  | 1.22     | 0.00     |
|             | 82     | 1.00    | 0.00    | 28090.14 | 3.72     | 0.00    | 12406.85 | 2.00    | 0.00    | 246    | 1.32     | 0.00     |
|             | 7863.1 |         |         |          |          |         |          |         | 10031   |        |          |          |
|             | 96     |         |         | 28115.67 |          |         | 12380.32 |         |         | .95    |          |          |

| Figure S17n | 18254.<br>39<br>18289.<br>8<br>18234. | 1.00 | 0.00 | 30108.24<br>30100.24 | 1.70 | 0.08 | 20426.8<br>20369.1 | 1.41 | 0.00 | 15814<br>.39<br>15725<br>.15<br>15602 | 0.89 | 0.01 |
|-------------|---------------------------------------|------|------|----------------------|------|------|--------------------|------|------|---------------------------------------|------|------|
|             | 56                                    |      |      | 30089.24             |      |      | 20425.51           |      |      | .02                                   |      |      |

Mean and SD values are the ones presented in Figure S17b (IL-1 $\beta$ ), c (IL-6), d (TNF $\alpha$ ), e (ICAM-1), f (VCAM-1), g (The fluorescence intensity of Calcein-AM) (a.u.), i (p-p65/p65 ratio), j (I $\kappa$ B $\alpha$ ), l (IL-1 $\beta$ ), m (IL-6), and n (TNF- $\alpha$ ). Data in b-e normalized to  $\beta$ -actin and presented as fold difference compared to 0 h control. Data in i presented as fold difference compared to 0 h. Data in 1-n normalized to  $\beta$ -actin and presented as fold difference compared to control.

Table S31. Adhesion molecules, pro-inflammatory cytokines and p65 protein expression levels in HUVEC exposed to PCB29-pQ with *CAV1* siRNA.

|             |        | NC siRNA |         |          | NC siRNA + PCB29-pQ |         |          | CAV1 siRNA |         |        | <i>CAV1</i> siRNA + PCB29-pQ |          |  |
|-------------|--------|----------|---------|----------|---------------------|---------|----------|------------|---------|--------|------------------------------|----------|--|
|             | Graysc | Mean     | SD      | Grayscal | Mean                | SD      | Grayscal | Mean       | SD      | Grays  | Mean                         | SD       |  |
|             | ale    | (normal  | (normal | e values | (normal             | (normal | e values | (normal    | (normal | cale   | (normal                      | (normali |  |
|             | values | ized)    | ized)   |          | ized)               | ized)   |          | ized)      | ized)   | values | ized)                        | zed)     |  |
|             | 30214. |          |         |          |                     |         |          |            |         | 23260  |                              |          |  |
|             | 87     |          |         | 37211.05 |                     |         | 23534.58 |            |         | .63    |                              |          |  |
| Eiguro C19h | 30368. | 1.00     | 0.02    |          | 1.24                | 0.02    |          | 0.91       | 0.02    | 24800  | 0.86                         | 0.02     |  |
| Figure 5160 | 41     | 1.00     | 0.02    | 38160.58 | 1.24                | 0.02    | 24974.92 | 0.01       | 0.05    | .87    | 0.80                         | 0.05     |  |
|             | 31474. |          |         |          |                     |         |          |            |         | 24913  |                              |          |  |
|             | 41     |          |         | 38190.24 |                     |         | 24898.73 |            |         | .41    |                              |          |  |

|                        | 27480.       |      |      |          |      |      |          |      |      | 13017        |       |      |
|------------------------|--------------|------|------|----------|------|------|----------|------|------|--------------|-------|------|
|                        | 92           |      |      | 33946.92 |      |      | 25009.53 |      |      | .08          |       |      |
| <b>F</b> ' <b>G</b> 10 | 27800.       | 1.00 | 0.01 |          | 1.04 | 0.01 |          | 0.01 | 0.01 | 13238        | 0 51  | 0.00 |
| Figure S18c            | 12           | 1.00 | 0.01 | 33688.38 | 1.24 | 0.01 | 24770.82 | 0.91 | 0.01 | .20          | 0.51  | 0.00 |
|                        | 27472.       |      |      |          |      |      |          |      |      | 13060        |       |      |
|                        | 05           |      |      | 34324.87 |      |      | 24471.58 |      |      | .37          |       |      |
|                        | 16321.       |      |      |          |      |      |          |      |      | 13659        |       |      |
|                        | 92           |      |      | 30156.51 |      |      | 14698.73 |      |      | .00          |       |      |
| Eiguro C19d            | 16318.       | 1.00 | 0.01 |          | 1 97 | 0.00 |          | 0.02 | 0.00 | 13695        | 0.01  | 0.01 |
| Figure 5180            | 75           | 1.00 | 0.01 | 30156.51 | 1.0/ | 0.00 | 14738.73 | 0.92 | 0.00 | .71          | 0.91  | 0.01 |
|                        | 16022.       |      |      |          |      |      |          |      |      | 13699        |       |      |
|                        | 02           |      |      | 30156.51 |      |      | 14686.73 |      |      | .00          |       |      |
|                        | 24843.       |      |      |          |      |      |          |      |      | 19715        |       |      |
|                        | 80           |      |      | 34943.65 |      |      | 33345.68 |      |      | .92          |       |      |
| Figure S18e            | 24493.       | 1.00 | 0.00 |          | 1.43 | 0.00 |          | 1.36 | 0.01 | 19912        | 0.87  | 0.00 |
| i iguie sitee          | 92           | 1100 | 0.00 | 35010.12 | 1110 | 0.00 | 33010.68 | 1100 | 0101 | .63          | 0.07  | 0.00 |
|                        | 24677.       |      |      |          |      |      |          |      |      | 19799        |       |      |
|                        | 87           |      |      | 35044.12 |      |      | 32993.15 |      |      | .51          |       |      |
|                        | 16432.       |      |      |          |      |      |          |      |      | 14917        |       |      |
|                        | 29           |      |      | 29701.36 |      |      | 16065.10 |      |      | .08          |       |      |
| Figure S18f            | 16188.       | 1.00 | 0.01 | 2068665  | 1.83 | 0.00 | 16100 62 | 0.79 | 0.00 | 15005        | 0.99  | 0.01 |
| _                      | 16429        |      |      | 29080.03 |      |      | 10100.03 |      |      | .20          |       |      |
|                        | 10438.<br>20 |      |      | 20721 77 |      |      | 16100.63 |      |      | 14972        |       |      |
|                        | 2062.2       |      |      | 7220.66  |      |      | 25073 10 |      |      | .70          |       |      |
|                        | 68           |      |      | 7550.00  |      |      | 23073.10 |      |      | 20131<br>15  |       |      |
|                        | 1008 1       |      |      | 7178 832 |      |      | 24649 32 |      |      | .+J<br>28210 |       |      |
| Figure S18h            | 30           | 1.00 | 0.01 | /1/0.052 | 1.42 | 0.10 | 27077.32 | 0.17 | 0.00 | 50           | 0.29  | 0.00 |
|                        | 4134.1       |      |      | 7261 953 |      |      | 24475 80 |      |      | 27942        |       |      |
|                        | 96           |      |      | ,201.755 |      |      | 211/3.00 |      |      | .84          |       |      |
|                        | 21031.       | 4.00 | 0.05 | 12072.82 | 0.55 | 0.00 | 30701.04 |      | 0.00 | 18197        | 0.6.1 | 0.01 |
| Figure S18j            | 05           | 1.00 | 0.02 | 120,2002 | 0.53 | 0.02 | 20,01101 | 1.44 | 0.00 | .58          | 0.84  | 0.01 |

| 21909. | 11440.39 | 30541.67 | 18714 |
|--------|----------|----------|-------|
| 72     |          |          | .48   |
| 21875. | 11506.80 | 30697.50 | 18468 |
| 14     |          |          | .00   |

|             |        | 0 h     |         | 1 h      |         |         | 3 h      |         |         | 6 h    |         |          |
|-------------|--------|---------|---------|----------|---------|---------|----------|---------|---------|--------|---------|----------|
|             | Graysc | Mean    | SD      | Grayscal | Mean    | SD      | Grayscal | Mean    | SD      | Grays  | Mean    | SD       |
|             | ale    | (normal | (normal | e values | (normal | (normal | e values | (normal | (normal | cale   | (normal | (normali |
|             | values | ized)   | ized)   |          | ized)   | ized)   |          | ized)   | ized)   | values | ized)   | zed)     |
| Figure S18j | 0.07   |         |         | 0.72     |         |         | 0.01     |         |         | 0.17   |         |          |
| (Right      | 0.08   | 0.08    | 0.01    | 0.88     | 0.83    | 0.09    | 0.07     | 0.05    | 0.03    | 0.28   | 0.23    | 0.06     |
| panel)      | 0.10   |         |         | 0.90     |         |         | 0.08     |         |         | 0.24   |         |          |

Mean and SD values are the ones presented in Figure S18b (IL-1β), c (IL-6), d (TNF-α), e (ICAM-1), f (VCAM-1), h (p-p65/p65 ratio), I (IκBα),

and j (Pearson's coefficient). Data in c-f and i normalized to  $\beta$ -actin and presented as fold difference compared to NC siRNA. Data in h presented as fold difference compared to NC siRNA.

| Table S32. Protein ex | pression of inflammator | v factors and ROS levels in | HUVEC exi | posed to PCB29-pO. |
|-----------------------|-------------------------|-----------------------------|-----------|--------------------|
|                       |                         |                             |           |                    |

|        | Control          | PCB29-pQ         | PCB29-pQ + VC    | PCB29-pQ + VE    | PCB29-pQ + NAC   |  |
|--------|------------------|------------------|------------------|------------------|------------------|--|
|        | Grayscale values |  |
| Figure | 13974.88         | 14314.47         | 6892.91          | 4899.10          | 6949.18          |  |
| S19b   | 12348.40         | 14314.47         | 6892.91          | 4899.10          | 6954.59          |  |
|        | 13750.64         | 14314.47         | 6892.91          | 4899.10          | 6949.18          |  |
|        | Grayscale values |  |
| Figure | 15912.10         | 21827.13         | 11045.47         | 12279.83         | 16616.30         |  |
| S19c   | 15863.28         | 22036.83         | 11003.47         | 12345.83         | 16551.59         |  |
|        | 15839.98         | 21941.25         | 11018.93         | 12351.83         | 16497.59         |  |

|        | Grayscale values |
|--------|------------------|------------------|------------------|------------------|------------------|
| Figure | 16573.56         | 23407.17         | 16591.27         | 20362.92         | 15709.90         |
| S19d   | 16479.90         | 23292.87         | 16712.22         | 20046.56         | 16281.73         |
|        | 16693.39         | 23259.34         | 16577.97         | 20474.51         | 15738.20         |
|        | Grayscale values |
| Figure | 24377.20         | 27848.27         | 9483.76          | 19209.90         | 16895.78         |
| S19e   | 24422.78         | 27853.27         | 9527.47          | 19147.08         | 16898.95         |
|        | 24457.78         | 27858.27         | 9443.76          | 19198.78         | 16931.78         |
|        | Grayscale values |
| Figure | 20508.34         | 3491.42          | 24095.95         | 22248.27         | 506.34           |
| S19f   | 20143.15         | 3515.83          | 24081.41         | 22567.15         | 494.34           |
|        | 20916.39         | 3403.59          | 23976.77         | 22010.61         | 506.34           |
|        | Grayscale values |
| Figure | 8957.13          | 11376.25         | 7745.71          | 4248.71          | 9647.69          |
| S19g   | 9396.02          | 10545.05         | 7818.54          | 4345.42          | 9577.69          |
|        | 9970.21          | 10050.10         | 7784.83          | 4246.42          | 9650.40          |

|        | Control             | PCB29-pQ            | PCB29-pQ + VC       | PCB29-pQ + VE       | PCB29-pQ + NAC      |
|--------|---------------------|---------------------|---------------------|---------------------|---------------------|
|        | Fluorescence values |
| Figure | 1001.23             | 1391.82             | 1121.46             | 961.25              | 1041.36             |
| S19h   | 1001.37             | 1361.78             | 1171.53             | 901.18              | 1081.41             |
|        | 1001.31             | 1351.76             | 1031.35             | 901.18              | 1091.42             |

|       | Control    |            | PCB29-pQ   |            | PCB29-pQ + VC |            | PCB29-pQ + VE |            | PCB29-pQ + NAC |            |
|-------|------------|------------|------------|------------|---------------|------------|---------------|------------|----------------|------------|
|       | Mean       | SD         | Mean       | SD         | Mean          | SD         | Mean          | SD         | Mean           | SD         |
|       | (normalize | (normalize | (normalize | (normalize | (normalize    | (normalize | (normalize    | (normalize | (normalize     | (normalize |
|       | d)         | d)         | d)         | d)         | d)            | d)         | d)            | d)         | d)             | d)         |
| Figur |            |            |            |            |               |            |               |            |                |            |
| e     | 1.00       | 0.07       | 1.36       | 0.06       | 0.58          | 0.02       | 0.39          | 0.02       | 0.60           | 0.01       |
| S19b  |            |            |            |            |               |            |               |            |                |            |

| Figur | 1.00 | 0.00 | 1 71 | 0.01 | 0.74 | 0.00 | 0.82 | 0.00 | 1 20 | 0.00 |
|-------|------|------|------|------|------|------|------|------|------|------|
| S19c  | 1.00 | 0.00 | 1.71 | 0.01 | 0.71 | 0.00 | 0.02 | 0.00 | 1.20 | 0.00 |
| Figur |      |      |      |      |      |      |      |      |      |      |
| e     | 1.00 | 0.01 | 1.74 | 0.01 | 1.07 | 0.00 | 1.29 | 0.01 | 1.10 | 0.02 |
| S19d  |      |      |      |      |      |      |      |      |      |      |
| Figur |      |      |      |      |      |      |      |      |      |      |
| e     | 1.00 | 0.00 | 1.41 | 0.10 | 0.41 | 0.00 | 0.83 | 0.00 | 0.79 | 0.00 |
| S19e  |      |      |      |      |      |      |      |      |      |      |
| Figur |      |      |      |      |      |      |      |      |      |      |
| e     | 1.00 | 0.02 | 1.25 | 0.00 | 0.21 | 0.00 | 0.14 | 0.01 | 0.03 | 0.00 |
| S19f  |      |      |      |      |      |      |      |      |      |      |
| Figur |      |      |      |      |      |      |      |      |      |      |
| e     | 1.00 | 0.05 | 1.40 | 0.09 | 0.88 | 0.00 | 0.48 | 0.01 | 1.17 | 0.01 |
| S19g  |      |      |      |      |      |      |      |      |      |      |
| Figur |      |      |      |      |      |      |      |      |      |      |
| e     | 1.00 | 0.07 | 1.37 | 0.02 | 1.11 | 0.07 | 0.92 | 0.03 | 1.07 | 0.03 |
| S19h  |      |      |      |      |      |      |      |      |      |      |

Mean and SD values are the ones presented in Figure S19b (p-65/p65 ratio), c (IL-1β), d (IL-6), e (TNF-α), f (ICAM-1), g (VCAM-1), and h (ROS

level). Data in c-g are normalized to  $\beta$ -actin. All data presented as fold difference relative to control.

| Table S33. p-CAV1 pro | otein expression rela | ative to CAV1 total <b>p</b> | protein in HUVEC ex | posed to PCB29-p | Q and antioxidants. |
|-----------------------|-----------------------|------------------------------|---------------------|------------------|---------------------|
|-----------------------|-----------------------|------------------------------|---------------------|------------------|---------------------|

|            | Control    |            | PCB29-pQ   |            | PCB29-pQ + VC |            | PCB29-pQ + VE |            | PCB29-pQ + NAC |            |
|------------|------------|------------|------------|------------|---------------|------------|---------------|------------|----------------|------------|
|            | Mean       | SD         | Mean       | SD         | Mean          | SD         | Mean          | SD         | Mean           | SD         |
|            | (normalize | (normalize | (normalize | (normalize | (normalize    | (normalize | (normalize    | (normalize | (normalize     | (normalize |
|            | d)         | d)         | d)         | d)         | d)            | d)         | d)            | d)         | d)             | d)         |
| Figur<br>e | 1.00       | 0.03       | 2.02       | 0.03       | 0.82          | 0.06       | 0.42          | 0.02       | 0.95           | 0.03       |

| S20a  |  |  |  |  |  |
|-------|--|--|--|--|--|
| (Righ |  |  |  |  |  |
| t     |  |  |  |  |  |
| panel |  |  |  |  |  |
| )     |  |  |  |  |  |

|  | Control    |            | PCB29-pQ   |            | PCB29-pQ + PEG-CAT |            | PCB29-pQ + PEG-SOD |            | PCB29-pQ + GSH-MEE |            |
|--|------------|------------|------------|------------|--------------------|------------|--------------------|------------|--------------------|------------|
|  | Mean       | SD         | Mean       | SD         | Mean               | SD         | Mean               | SD         | Mean               | SD         |
|  | (normalize | (normalize | (normalize | (normalize | (normalize         | (normalize | (normalize         | (normalize | (normalize         | (normalize |
|  | d)         | d)         | d)         | d)         | d)                 | d)         | d)                 | d)         | d)                 | d)         |
| Figur<br>e<br>S20b<br>(Righ<br>t<br>panel<br>) | 1.00       | 0.17       | 20.01      | 0.32       | 8.35               | 0.16       | 9.23               | 0.16       | 4.64               | 0.11       |

|                         | Control          | PCB29-pQ         | PCB29-pQ + VC    | PCB29-pQ + VE    | PCB29-pQ + NAC   |
|-------------------------|------------------|------------------|------------------|------------------|------------------|
|                         | Grayscale values |
| Figure S20a<br>(p-CAV1) | 13410.85         | 27941.75         | 11037.39         | 4832.47          | 10080.42         |
|                         | 13993.92         | 27406.58         | 10979.39         | 5269.59          | 9641.52          |
|                         | 13638.61         | 28534.65         | 9943.78          | 5056.23          | 9682.05          |
|                         | Grayscale values |
| Figure S20a<br>(CAV1)   | 30972.85         | 31152.49         | 28496.59         | 27020.02         | 22916.02         |
|                         | 30766.20         | 30912.78         | 29202.61         | 26508.83         | 23050.15         |
|                         | 30282.13         | 31115.78         | 29372.49         | 26523.42         | 23198.27         |

|             | Control          | PCB29-pQ         | PCB29-pQ + PEG-  | PCB29-pQ + PEG-  | PCB29-pQ + GSH-  |  |
|-------------|------------------|------------------|------------------|------------------|------------------|--|
|             |                  |                  | CAT              | SOD              | MEE              |  |
| Figure S20b | Grayscale values |  |
| (p-CAV1)    | 1181.16          | 30552.43         | 11266.49         | 12159.02         | 6742.50          |  |

|             | 981.63           | 31232.09         | 11089.49         | 12099.44         | 6548.62          |
|-------------|------------------|------------------|------------------|------------------|------------------|
|             | 1374.41          | 30331.07         | 10787.42         | 12484.56         | 6891.33          |
|             | Grayscale values |
| Figure S20b | 22054.73         | 28945.39         | 24954.80         | 24981.15         | 27437.22         |
| (CAV1)      | 22386.22         | 28874.73         | 24947.92         | 24977.61         | 27180.97         |
|             | 22204.80         | 28874.44         | 24811.39         | 24980.73         | 27335.80         |

Mean and SD values are the ones presented in Figure S20a and b. Data presented as fold difference relative to control.