

Figure 5.2 Monocytes (MO) from PE patients co-cultured with HUVECs exhibit an amplified inflammatory response of IL-6

Top panels: Basal levels of IL-6 production by HUVEC monoculture, NP/PE MO monocultures, HUVEC-NP MO co-culture or HUVEC-PE MO co-culture (*left panel*-pg/ml or *right panel*-MFI). *Middle panels:* Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures, and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-pg/ml or *right panel*-MFI). *Bottom panels:* Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-fold change (pg/ml) or *right panel*-fold change (MFI)). Fold change was calculated by dividing pg/ml or MFI values over untreated control pg/ml or MFI values and illustrated as mean ± SEM. Comparisons were made between NP/PE MO monocultures versus NP/PE co-cultures; NP versus PE mono- or co-cultures; and non-stimulated cultures. Statisticance was determined by the Mann-Whitney U test to compare NP/PE monocultures versus NP/PE co-cultures versus PE mono- or co-cultures; and One Way ANOVA (Bonferroni multiple comparison test) to compare differences between the unstimulated cells to different stimulators (bacterial and endogenous ligands of TLRs) (n=7-9). \*\*P < 0.01 NP/PE MO vs HUVEC-NP/PE MO, †p < 0.05 PE mono- or co-cultures as compared with NP mono- or co-cultures, ^p < 0.05 non-stimulated control compared with stimulated cultures; and §\$p < 0.01 LPS-treated HUVECs as compared to untreated HUVECs.

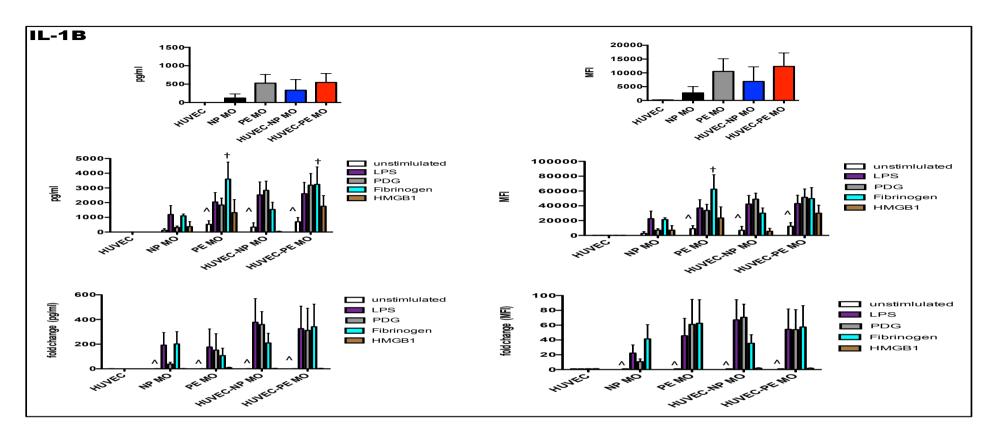


Figure 5.3 Monocytes (MO) from PE patients co-cultured with HUVECs did not exhibit a significant change of IL-1β

Top panels: Basal levels of IL-1β production by HUVEC monoculture, NP/PE MO monocultures, HUVEC-NP MO co-culture or HUVEC-PE MO co-culture (*left panel*-pg/ml or *right panel*-MFI). *Middle panels*: Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures, and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-pg/ml or *right panel*-MFI). *Bottom panels*: Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-fold change (pg/ml) or *right panel*-fold change (MFI)). Fold change was calculated by dividing pg/ml or MFI values over untreated control pg/ml or MFI values and illustrated as mean ± SEM. Comparisons were made between NP/PE MO monocultures versus NP/PE co-cultures; NP versus PE mono- or co-cultures; and non-stimulated cultures. Statistical significance was determined by the Mann-Whitney U test to compare NP/PE monocultures versus NP/PE co-cultures and NP mono- or co-cultures versus PE mono- or co-cultures; and One Way ANOVA (Bonferroni multiple comparison test) to compare differences between the unstimulated cells to different stimulators (bacterial and endogenous ligands of TLRs) (n=7-9).  $^{\dagger}$ P < 0.05 PE mono- or co-cultures as compared with NP mono- or co-cultures,  $^{\dagger}$ P < 0.05 PE mono- or co-cultures.

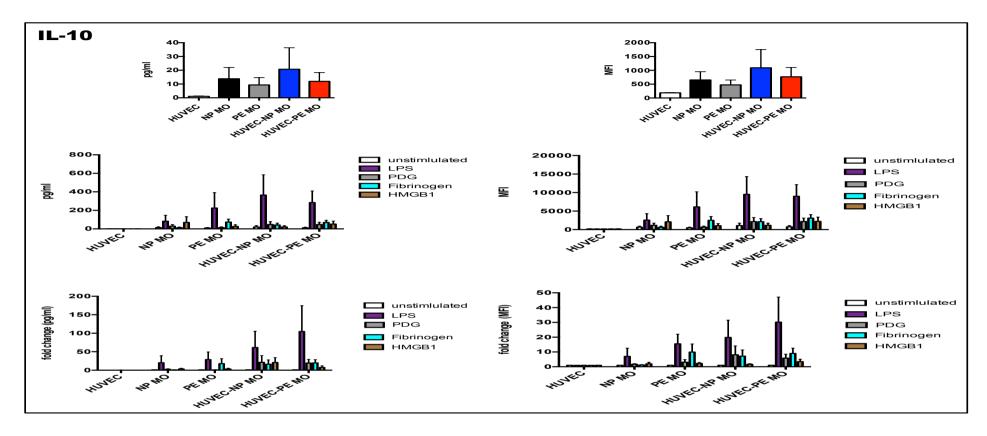


Figure 5.4 Monocytes (MO) from PE patients co-cultured with HUVECs exhibit a declined anti- inflammatory response of IL-10

Top panels: Basal levels of IL-10 production by HUVEC monoculture, NP/PE MO monocultures, HUVEC-NP MO co-culture or HUVEC-PE MO co-culture (*left panel*-pg/ml or *right panel*-MFI). *Middle panels*: Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures, and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-pg/ml or *right panel*-MFI). *Bottom panels*: Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-fold change (pg/ml) or *right panel*-fold change (MFI)). Fold change was calculated by dividing pg/ml or MFI values over untreated control pg/ml or MFI values and illustrated as mean ± SEM. Comparisons were made between NP/PE MO monocultures versus NP/PE co-cultures; NP versus PE mono- or co-cultures; and non-stimulated cultures. Statistical significance was determined by the Mann-Whitney U test to compare NP/PE monocultures versus NP/PE co-cultures and NP mono- or co-cultures versus PE mono- or co-cultures; and One Way ANOVA (Bonferroni multiple comparison test) to compare differences between the unstimulated cells to different stimulators (bacterial and endogenous ligands of TLRs) (n=7-9).

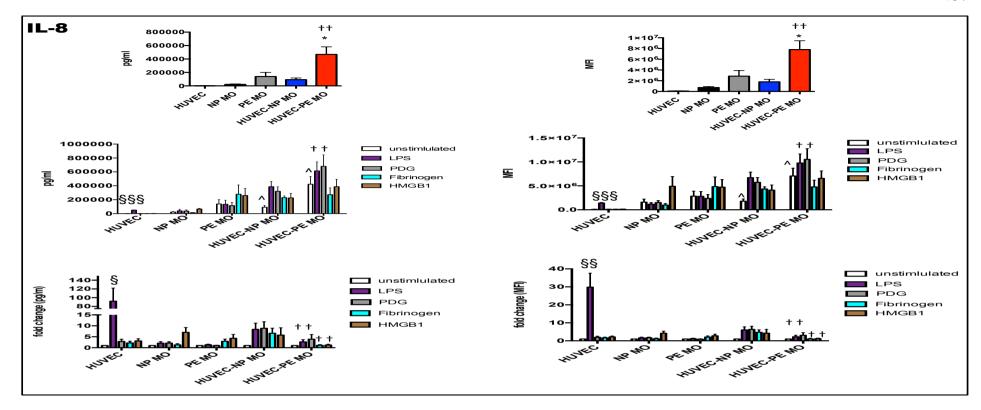


Figure 5.5 Monocytes (MO) from PE patients co-cultured with HUVECs exhibit an amplified inflammatoryresponse of IL-8

Top panels: Basal levels of IL-8 production by HUVEC monoculture, NP/PE MO monocultures, HUVEC-NP MO co-culture or HUVEC-PE MO co-culture (*left panel*-pg/ml or *right panel*-MFI). *Middle panels*: Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures, and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-pg/ml or *right panel*-MFI). *Bottom panels*: Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-fold change (pg/ml) or *right panel*-fold change (MFI)). Fold change was calculated by dividing pg/ml or MFI values over untreated control pg/ml or MFI values and illustrated as mean  $\pm$  SEM. Comparisons were made between NP/PE MO monocultures versus NP/PE co-cultures; NP versus PE mono- or co-cultures; and non-stimulated cultures. Statistical significance was determined by the Mann-Whitney U test to compare NP/PE monocultures versus NP/PE co-cultures and NP mono- or co-cultures versus PE mono- or co-cultures; and One Way ANOVA (Bonferroni multiple comparison test) to compare differences between the unstimulated cells to different stimulators (bacterial and endogenous ligands of TLRs) (n=7-9). \*P < 0.05 NP/PE MO vs HUVEC-NP/PE MO, \*p < 0.05, \*f\*p < 0.01 PE mono- or co-cultures as compared with NP mono- or co-cultures,  $\Delta$  0.05 non-stimulated control compared with stimulated cultures; and \$p < 0.05, \$\frac{8}{9}\$p < 0.01, \$\frac{8}{9}\$p < 0.01 LPS-treated HUVECs.

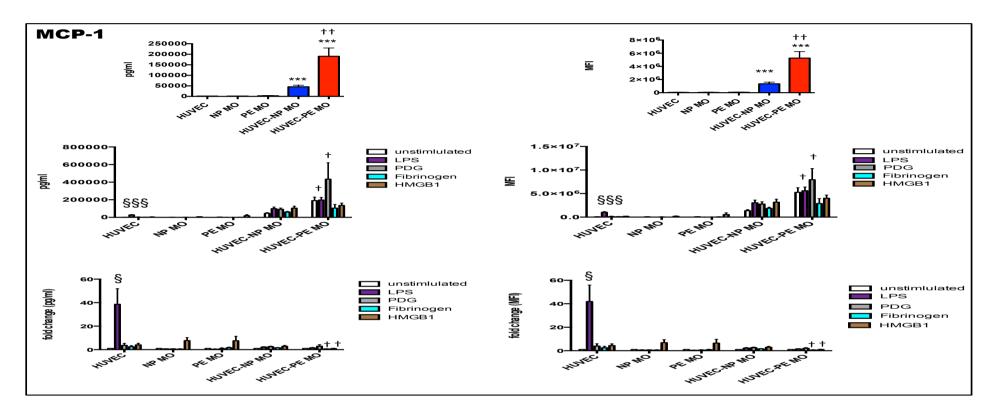


Figure 5.6 Monocytes (MO) from PE patients co-cultured with HUVECs exhibit an amplified inflammatory response of MCP-1

Top panels: Basal levels of MCP-1 production by HUVEC monoculture, NP/PE MO monocultures, HUVEC-NP MO co-culture or HUVEC-PE MO co-culture (*left panel*-pg/ml or *right panel*-MFI). *Middle panels*: Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures, and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-pg/ml or *right panel*-MFI). *Bottom panels*: Graphic representation of the response of HUVEC monoculture, NP/PE MO monocultures and HUVEC-NP or PE MO co-cultures to LPS, PDG, fibrinogen and HMGB1 (*left panel*-fold change (pg/ml) or *right panel*-fold change (MFI)). Fold change was calculated by dividing pg/ml or MFI values over untreated control pg/ml or MFI values and illustrated as mean ± SEM. Comparisons were made between NP/PE MO monocultures versus NP/PE co-cultures; NP versus PE mono- or co-cultures; and non-stimulated cultures. Statistical significance was determined by the Mann-Whitney U test to compare NP/PE monocultures versus NP/PE co-cultures and NP mono- or co-cultures versus PE mono- or co-cultures; and One Way ANOVA (Bonferroni multiple comparison test) to compare differences between the unstimulated cells to different stimulators (bacterial and endogenous ligands of TLRs) (n=7-9). \*\*\*P < 0.001 NP/PE MO vs HUVEC-NP/PE MO, †p < 0.05, ††p < 0.01 PE mono- or co- cultures as compared with NP mono- or co- cultures; and \$p < 0.05, §\$p < 0.01, §\$\$p < 0.01 LPS-treated HUVECs as compared to untreated HUVECs.