## Supplementary data

## Supplementary Methods

The sensitivities for the MMP Luminex assays were MMP-1 $1.1 \mathrm{pg} / \mathrm{ml}, \mathrm{MMP}-212.6 \mathrm{pg} / \mathrm{ml}, \mathrm{MMP}-3$ $7.3 \mathrm{pg} / \mathrm{ml}$, MMP-7 $6.6 \mathrm{pg} / \mathrm{ml}, ~ M M P-816.6 \mathrm{pg} / \mathrm{ml}$, MMP-9 $13.7 \mathrm{pg} / \mathrm{ml}, \mathrm{MMP}-103.2 \mathrm{pg} / \mathrm{ml}, \mathrm{MMP}-12$ $0.7 \mathrm{pg} / \mathrm{ml}, \mathrm{MMP}-1363.5 \mathrm{pg} / \mathrm{ml}$, EMPRIN $5.6 \mathrm{pg} / \mathrm{ml}$.

The sensitivities for the TIMP Luminex assays were TIMP-1 $3.43 \mathrm{pg} / \mathrm{ml}$, TIMP-2 $40.1 \mathrm{pg} / \mathrm{ml}$, TIMP-3, 20.0pg/ml, TIMP-4 0.28pg/ml.

The sensitivities for the cytokine Luminex assays were $\mathrm{IL} 1 \beta 0.18 \mathrm{pg} / \mathrm{ml}, \mathrm{IL}-20.28 \mathrm{pg} / \mathrm{ml}, \mathrm{IL}-60.31 \mathrm{pg} / \mathrm{ml}$, $\mathrm{IL}-80.07 \mathrm{pg} / \mathrm{ml}$, IL-10 0.24, IFN-g $0.08 \mathrm{pg} / \mathrm{ml}$ and TNF- $\alpha 0.54 \mathrm{pg} / \mathrm{ml}$.

Supplementary Table 1 Partial Spearman correlation coefficients between CT markers and the MMP with the strongest bivariate correlation, controlling for the effect of each other MMP in turn

|  | Controlling for: |  |  |  |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | MMP1 | MMP2 | MMP3 | MMP7 | MMP8 | MMP9 | MMP10 | MMP12 | MMP13 |
| FEV1\% | $\begin{gathered} -0.54 \\ (p=.008) \end{gathered}$ | $\begin{gathered} -0.56 \\ (p=.005) \end{gathered}$ | $\begin{gathered} -0.58 \\ (p=.004) \end{gathered}$ | $\begin{gathered} -0.53 \\ (p=.01) \end{gathered}$ |  | $\begin{gathered} -0.20 \\ (p=.370) \end{gathered}$ | $\begin{gathered} -0.58 \\ (p=.003) \end{gathered}$ | $\begin{gathered} -0.42 \\ (p=.048) \end{gathered}$ | $\begin{gathered} -0.60 \\ (p-.002) \end{gathered}$ |
| FEF25/75 | $\begin{gathered} -0.53 \\ (p=.010) \end{gathered}$ | $\begin{gathered} -0.56 \\ (p=.006) \end{gathered}$ | $\begin{gathered} -0.62 \\ (p=.002) \end{gathered}$ | $\begin{gathered} -0.65 \\ (p=.001) \end{gathered}$ |  | $\begin{gathered} -0.25 \\ (p=.247) \end{gathered}$ | $\begin{gathered} -0.61 \\ (p=.002) \end{gathered}$ | $\begin{gathered} -0.48 \\ (p .021) \end{gathered}$ | $\begin{gathered} -0.61 \\ (p=.002) \end{gathered}$ |
| LAA\% | $\begin{gathered} 0.53 \\ (p=.009) \end{gathered}$ | $\begin{gathered} 0.57 \\ (p=.005) \end{gathered}$ | $\begin{gathered} 0.32 \\ (p=.139) \end{gathered}$ | $\begin{gathered} 0.33 \\ (p=.124) \end{gathered}$ | $\begin{gathered} 0.46 \\ (p=.026) \end{gathered}$ | $\begin{gathered} 0.46 \\ (p=.027) \end{gathered}$ |  | $\begin{gathered} 0.51 \\ (p=.013) \end{gathered}$ | $\begin{gathered} 0.56 \\ (p=.006) \end{gathered}$ |
| EI/MLD | $\begin{gathered} 0.66 \\ (p=.007) \\ \hline \end{gathered}$ | $\begin{gathered} 0.60 \\ (p=.019) \end{gathered}$ | $\begin{gathered} 0.46 \\ (p=.083) \\ \hline \end{gathered}$ | $\begin{gathered} 0.37 \\ (p=.172) \end{gathered}$ |  | $\begin{gathered} 0.28 \\ (p=.318) \end{gathered}$ | $\begin{gathered} 0.46 \\ (p=.084) \end{gathered}$ | $\begin{gathered} 0.38 \\ (p=.161) \end{gathered}$ | $\begin{gathered} 0.60 \\ (p=.019) \end{gathered}$ |

$\mathrm{N}=24$ except for $\mathrm{EI} / \mathrm{MLD}$ ( $\mathrm{N}=16$ ).
MMP8 had the strongest bivariate correlation with all CT markers apart from LAA\%, which had the strongest bivariate correlation with MMP10.

Supplementary Table 2 Spearman's correlation analysis between cytokines, neutrophils and lung function and CT measures of disease in COPD subjects

|  | IL-1 $\boldsymbol{\beta}$ | IL-6 | IL-8 | GM-CSF | Neutrophils |
| :--- | :--- | :--- | :--- | :--- | :--- |
| FEV1\% | -0.16 | -0.39 | $-\mathbf{0 . 4 1 ^ { * }}$ | -0.05 | -0.40 |
| FEF75-25\% | $-\mathbf{0 . 4 4 ^ { * }}$ | -0.37 | $-\mathbf{0 . 5 7 * *}$ | -0.18 | -0.18 |
| Emphysema \% <br> (LAA\%) | -0.30 | 0.28 | 0.22 | -0.13 | 0.36 |
| Small airways <br> Disease <br> (E/I MLD) | 0.12 | 0.29 | $\mathbf{0 . 5 3 *}$ | -0.19 | 0.43 |
| Bronchial wall <br> area (Pi10) | 0.05 | -0.10 | 0.10 | 0.09 | 0.10 |
| Spearman's rho values given. <br> N=24, apart from associations with E/I MLD (n=16) <br> $*$ <br> p<0.05 $* * p<0.01 ~ * * * p<0.001 ~$ |  |  |  |  |  |

