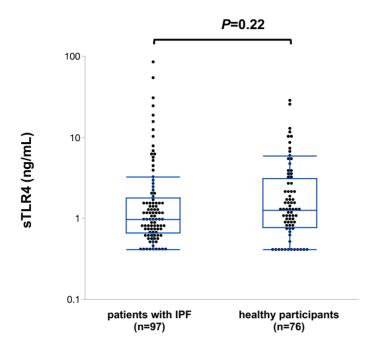
# **Supplementary Information (SI)**

### S1. Serum sTLR4 levels in patients with IPF and healthy participants



There were no significant differences in serum sTLR4 levels between patients with IPF and healthy participants (1.0 ng/mL [0.7-1.8] and 1.2 ng/mL [0.8-3.1], P=0.22, respectively). sTLR4, soluble toll-like receptor 4; IPF, idiopathic pulmonary fibrosis

# S2: Clinical characteristics of the study participants

|                             | All participants (n=173) | Patients with IPF (n=97) | Healthy participants (n=76) | P-value* |
|-----------------------------|--------------------------|--------------------------|-----------------------------|----------|
| Age, years                  | 66.0 (59.0–73.0)         | 68.0 (62.0-74.0)         | 61.5 (58.0-72.0)            | 0.024    |
| Sex, male/female            | 149/24                   | 86/11                    | 63/13                       | 0.28     |
| BMI, kg/m <sup>2</sup>      | 23.7 (21.4–25.7)         | 23.7 (21.6–26.2)         | 23.7 (21.2–25.4)            | 0.66     |
| Smoking history, pack years | 31.8 (8.1–50.0)          | 32.8 (15.9–49.5)         | 30.0 (0-57.5)               | 0.40     |
| FVC, %predicted             | 82.8 (67.4–95.0)         | 74.3 (64.1–88.1)         | 95.0 (86.0–111.3) **        | <.00010  |
| sTLR4, ng/mL                | 1.1 (0.7–2.2)            | 1.0 (0.7–1.8)            | 1.2 (0.8–3.1)               | 0.22     |

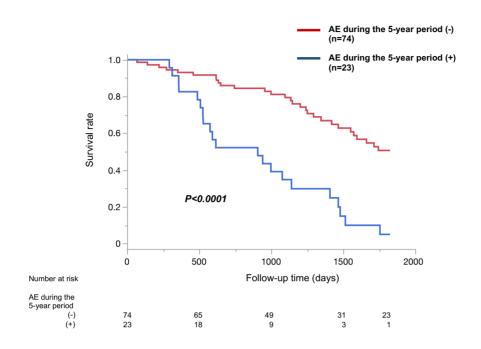
Data are presented as the median (interquartile range)

BMI, body mass index; FVC, forced vital capacity; IPF, idiopathic pulmonary fibrosis; sTLR4, soluble toll-loke receptor 4

<sup>\*</sup> Comparison of patients with IPF and healthy participants

<sup>\*\*</sup> There were 30 missing data in healthy participants.

# S3: The 5-year survival rate based on acute exacerbation



\* P< 0.05, Spearman correlation coefficients

AE, acute exacerbation

S4: Association between baseline characteristics and serum sTLR4 levels

|                                 | sT     | LR4     |
|---------------------------------|--------|---------|
|                                 | $r_s$  | P-value |
| Age, years                      | 0.160  | 0.12    |
| BMI, kg/m <sup>2</sup>          | -0.083 | 0.50    |
| Smoking history, pack years     | 0.072  | 0.49    |
| FVC, %                          | -0.073 | 0.48    |
| WBC, $/\mu L$                   | 0.130  | 0.21    |
| Neutrophil, /μL                 | 0.081  | 0.43    |
| Lymphocyte, /µL                 | 0.058  | 0.58    |
| Monocyte, $/\mu L$              | 0.23   | 0.020*  |
| Platelet, $\times 10^4 / \mu L$ | 0.044  | 0.67    |
| CRP, mg/dL                      | 0.17   | 0.11    |

<sup>\*</sup> P< 0.05, Spearman correlation coefficients

BMI, body mass index; CRP, C-reactive protein; FVC, forced vital capacity; sTLR4, soluble toll-like receptor 4; WBC, white blood cells

# S5: Clinical characteristics of the study participants

|                             | All patients with IPF (n=97) | Combination of sTLR4 and monocyte |                |                |
|-----------------------------|------------------------------|-----------------------------------|----------------|----------------|
|                             |                              | Group A (n=17)                    | Group B (n=36) | Group C (n=44) |
| Age, years                  | 68.0                         | 68.0                              | 69.0           | 67.0           |
|                             | (62.0-74.0)                  | (61.5-74.5)                       | (62.3-73.8)    | (59.8-74.0)    |
| Sex, male/female            | 86/11                        | 17/0                              | 33/3           | 36/8           |
| BMI, kg/m <sup>2</sup>      | 23.7                         | 23.7                              | 23.8           | 23.8           |
|                             | (21.6–26.2)                  | (22.8–25.2)                       | (21.4-25.6)    | (21.5–27.0)    |
| Smoking history, pack years | 32.8                         | 40.0                              | 30.0           | 33.3           |
|                             | (15.9-49.5)                  | (22.0-60.0)                       | (18.8-40.0)    | (11.3-49.5)    |
| FVC, %predicted             | 74.3                         | 76.4                              | 74.3           | 73.6           |
|                             | (64.1-88.1)                  | (65.6-89.8)                       | (61.8-86.8)    | (63.7-86.7)    |

Group A: Patients with soluble toll-like receptor 4 (sTLR4) level higher than 2.2 ng/mL and monocyte level higher than 381.0/μL

Group B: Patients with higher sTLR4 or higher monocyte levels

Group C: Patients with lower sTLR4 and lower monocyte levels

Data are presented as the median (interquartile range).

BMI, body mass index; FVC, forced vital capacity; IPF, idiopathic pulmonary fibrosis; sTLR4, soluble toll-like receptor 4