

1 **SUPPLEMENTARY MATERIAL**

2 **Table S1.** Assessment of disease progression—proportions of patients experiencing ≥10%
 3 decline in FVC or death, ≥50-m decline in 6MWD or death and ≥20-point worsening of UCSD
 4 SOBQ score or death in the pooled placebo population stratified by baseline FVC (≥80% and
 5 <80% predicted) or baseline GAP index stage (GAP I and GAP II–III) at 12 months

Outcome at 12 Months	Baseline FVC ≥80%	Baseline FVC <80%	Baseline GAP Stage I	Baseline GAP Stage II-III
Pooled placebo population, n	170	454	235	387
FVC, % of patients				
≥10% decline or death	21.8	28.0	21.7	28.9
No decline	21.2	16.3	20.4	15.8
6MWD, % of patients				
≥50-m decline or death	24.1	38.8	26.6	39.7
No decline	37.3	33.4	37.3	32.9
UCSD SOBQ, % of patients				
≥20-point change in UCSD SOBQ score or death	18.0	36.4	21.4	37.5
No decline	37.1	24.9	36.8	23.1

6 6MWD: 6-minute walk distance; FVC: forced vital capacity; UCSD SOBQ: University of
 7 California—San Diego Shortness of Breath Questionnaire.

8 **Table S2.** Proportions of patients experiencing $\geq 10\%$ decline in FVC or death, $\geq 50\text{-m}$ decline in
9 6MWD or death and ≥ 20 -point worsening of UCSD SOBQ score or death in the pooled
10 pirfenidone population stratified by baseline FVC ($\geq 80\%$ and $< 80\%$ predicted) or baseline GAP
11 index stage (GAP I and GAP II–III) at 12 months

Outcome at 12 Months	Baseline FVC $\geq 80\%$	Baseline FVC $< 80\%$	Baseline GAP Stage I	Baseline GAP Stage II-III
Pooled pirfenidone population, n	146	477	247	376
FVC, % of patients				
$\geq 10\%$ decline or death	14.4	14.9	13.4	15.7
No decline	37.7	25.2	31.2	26.1
6MWD, % of patients				
$\geq 50\text{-m}$ decline or death	16.8	27.2	19.7	28.2
No decline	46.2	39.0	47.5	36.2
UCSD SOBQ, % of patients				
≥ 20 -point change in UCSD SOBQ score or death	18.1	25.8	17.4	28.3
No decline	40.3	27.9	35.5	27.7

12 6MWD, 6-minute walk distance; FVC, forced vital capacity; UCSD SOBQ, University of
13 California—San Diego Shortness of Breath Questionnaire.

14 **Table S3.** Linear change in FVC (percent predicted and L) without imputation through 12
 15 months by treatment group and baseline FVC and GAP index stage

Slope (Year)*	Pirfenidone 2403 mg/day (N=477)	Placebo (N=454)	Mean Difference[†]	Relative Difference, %[‡]	P Value[§]
FVC, % predicted					
Baseline FVC ≥80%	-1.92	-5.19	3.269	63.0	<0.0001
Baseline FVC <80%	-3.66	-5.95	2.290	38.5	<0.0001
Baseline GAP Stage I	-2.93	-5.78	2.847	49.3	<0.0001
Baseline GAP Stage II-III	-3.41	-5.67	2.262	39.9	<0.0001
FVC, L					
Baseline FVC ≥80%	-0.083	-0.187	0.1047	55.9	0.0010
Baseline FVC <80%	-0.149	-0.238	0.0891	37.5	<0.0001
Baseline GAP Stage I	-0.120	-0.215	0.0945	44.0	0.0002
Baseline GAP Stage II-III	-0.140	-0.226	0.0853	37.8	<0.0001

16 FVC, forced vital capacity.

17 *Change from baseline was calculated as the 12-month value minus the baseline value.

18 [†]Difference in mean: (pirfenidone – placebo).

19 [‡]Relative difference in mean: 100*(pirfenidone – placebo)/(|placebo|).

20 [§]Calculated from the mixed linear model comparing pirfenidone 2403 mg/day to placebo, with
 21 change as the outcome variable; study (PIPF-004, PIPF-006 and PIPF-016), geographic region
 22 (United States and the rest of the world) within study and assessment time by treatment as fixed
 23 effects; baseline as a covariate; subject and subject by assessment time as random effects
 24 with variance components covariance structure.