Supplemental Material

Supplemental Methods:

Statistical Analysis

Descriptive statistics of the baseline clinical characteristics are presented as mean ± standard deviation (SD) for continuous variables, median (interquartile range (IQR)) for ordinal variables, and frequency for categorical variables. Baseline characteristics were stratified by presence of PH on right heart catheterization (RHC) (PH by RHC group, mean PAP≥25mmHg), absence of PH on right heart catheterization (No PH by RHC group, mean PAP<25mmHg), and by patients who did not undergo or had a technically inadequate RHC without measurement of mean PAP (Unknown PH status group). The baseline characteristics of the PH by RHC group were compared to the No PH by RHC group, using Kruskall-Wallis testing for continuous variables and chi-square testing for categorical variables. The association between baseline characteristics that were significantly different in the PH by RHC and No PH by RHC groups and PH was further assessed using univariate logistic regression. The RHC hemodynamics were described in the PH by RHC and No PH by RHC groups. Kruskall-Wallis testing or chi-square testing was performed as appropriate to assess differences in hemodynamic values between the two groups, not including the mean PAP, which was different by definition.

Adequacy of PH workup was described as the frequency with which guideline appropriate testing, including echocardiography, assessment of left ventricular function; ANA; liver function tests; HIV serologies; pulmonary function testing; chest imaging; V/Q scan and/or

chest CT angiography; sleep testing or overnight oximetry were performed in patients confirmed to have pulmonary hypertension⁸.

Actual survival from time of first clinic visit was assessed through the final follow up date of 5/31/2016, and time to death for patients who died was calculated; 1-year mortality from time of first clinic visit was calculated. A Kaplan-Meier survival curve for patients with confirmed PH by RHC was drawn.

Changes in treatment and functional status for patients with at least one follow up visit after initial clinic visit were described. Prescription for new or changes in existing supportive medications and other treatments (loop diurectics, including dosage in furosemide equivalents; warfarin; digoxin; PAH-specific therapies; oxygen; CPAP or other noninvasive ventilation), as well as changes in mMRC dyspnea scale, six-minute walk testing, and functional status from first visit to most recent clinic visit (through 5/31/2016) are presented. A follow up visit was defined as a return visit to PH clinic and/or a visit for RHC. Furosemide equivalents were defined as follows: 40mg furosemide=1mg bumetanide=20mg torsemide=50mg ethacrynic acid. Paired t-test for continuous variables, Wilcoxon signed ranks sums test for ordinal variables, or McNemar's chi-square testing for categorical variables was performed as appropriate to assess for changes in these characteristics between the first and most recent follow up visit.

All analysis was performed using Stata/SE version 11.2 software (StataCorp LP). A 2-sided p value of less than 0.05 was considered significant.

Supplemental Results:

The Multispecialty Pulmonary Hypertension Clinic

Prior to institution of the clinic, educational sessions about PH and PH clinic referral were undertaken with various groups within the medical center. The multispeciality PH clinic is staffed by an attending cardiologist and attending pulmonologist with interests in PH clinical care and research. Patients are referred to the clinic by placing a consultation request in the electronic medical record system. Trainees are regularly present in the clinic, including preventive cardiology fellows, pharmacy residents and students. The trainees typically perform the initial assessment of new patients, using a templated note to capture essential data needed for the diagnostic assessment of possible PH. Patients are then jointly discussed with the clinic attending staff, data is reviewed, and a diagnostic and/or care plan established. Patients are then seen and examined by the PH team and, after assessment, the recommendations for further testing or treatment are discussed with the patient. Right heart catheterizations are typically performed in the outpatient setting by the PH cardiologist and cardiology fellow in the catheterization suite, and catheterization results reviewed jointly by the PH attending team. Patients with catheter-confirmed PH return to the PH clinic for follow up; patients are also referred as appropriate for further care in related care settings such as the heart failure clinic or pulmonary rehabilitation program.

Supplemental Table 1: Baseline characteristics in patients undergoing right heart catheterization, with p values for comparisons (Kruskall-Wallis for continuous and chi-square for categorical variables)

	Patients with	Patients	P value for
	confirmed PH	without PH by	comparison
	by right heart	right heart	
	catheterization	catheterization	
	(n=73)	(n= 21)	
Age	72.8 (9.0)	73 (11.3)	0.679
Male sex	68/73 (93.2%)	21/21 (100%)	0.218
BMI (kg/m2)	34.3 (8.3)	29.3 (5.6)	0.012
AHA ideal health	10/73 (13.7%)	3/21 (14.3%)	0.446
(BMI<25)			
AHA intermediate	18/73 (24.7%)	8/21 (38.1%)	
health (BMI≥25,			
<30)			
AHA poor health	45/73 (61.6%)	10/21 (47.6%)	
(BMI ≥30)			
Heart rate (bpm)	76 (14)	76 (9)	0.952
Systolic blood	126 (18)	113 (13)	0.003
pressure (mmHg)			

Diastolic blood	71 (9)	69 (8)	0.222
pressure (mmHg)			
Pulse pressure	55 (15)	44 (11)	0.001
(mmHg)			
Oxyhemoglobin	94 (4) (n=70)	96 (2)	0.147
saturation (%)			
mMRC dyspnea	3 (2, 3)*	2 (1.5, 3)*	0.034
score, initial visit	(n=72)	(n=20)	
mMRC dyspnea	2/72 (2.8%)	2/20 (10.0%)	0.27
score 0			
mMRC dyspnea	6/72 (8.3%)	3/20 (15.0%)	
score 1			
mMRC dyspnea	17/72 (23.6%)	6/20 (30.0%)	
score 2			
mMRC dyspnea	32/72 (44.4%)	8/20 (40.0%)	
score 3			
mMRC dyspnea	15/72 (20.8%)	1/20 (5.0%)	
score 4			
NYHA functional	3 (2, 3)*	2 (1, 2.5)*	0.003
class, initial visit	(n=51)	(n=12)	
NYHA function	0/51 (0%)	0/12 (0%)	0.015

class 0			
NYHA functional	3/51 (5.9%)	4/12 (33.3%)	
class 1			
NYHA functional	14/51 (27.4%)	5/12 (41.7%)	
class 2			
NYHA functional	27/51 (52.9%)	3/12 (25.0%)	
class 3			
NYHA functional	7/51 (13.7%)	0/12 (0%)	
class 4			
Jugular venous	25/73 (34.2%)	6/21 (28.6%)	0.626
distention, initial			
visit			
Peripheral edema	40/73 (54.8%)	8/21 (38.1%)	0.177
History of anemia	25/73 (34.2%)	3/21 (14.3%)	0.078
History of	2/73 (2.7%)	1/21 (4.8%)	0.642
congenital heart			
disease			
History of valvular	15/73 (20.6%)	7/21 (33.3%)	0.223
heart disease			
History of	30/73 (41.1%)	8/21 (38.1%)	0.805
diabetes			

History of	48/73 (65.8%)	12/21 (57.1%)	0.469
systemic			
hypertension			
History of	5/73 (6.8%)	1/21 (4.8%)	0.730
anorexigen use			
History of	16/73 (21.9%)	3/21 (14.3%)	0.443
bleeding disorder,			
GI bleeding, or			
epistaxis			
History of cancer	25/73 (34.2%)	6/21 (28.6%)	0.626
History of heart	37/73 (51%)	7/21 (33.3%)	0.160
failure			
History of	5/73 (6.8%)	4/21 (19%)	0.094
connective tissue			
disease			
History of	43/73 (58.9%)	7/21 (33.3%)	0.038
coronary artery			
disease			
History of COPD	47/73 (64.4%)	9/21 (43%)	0.076
History of DVT/PE	12/73 (16.4%)	3/21 (14.3%)	0.812
History of heart	16/73 (21.9%)	4/21 (19%)	0.777

surgery/CABG			
History of HIV	0/73 (0%)	1/21 (4.8%)	0.061
infection			
History of liver	9/73 (12.3%)	2/21 (9.5%)	0.725
disease			
History of OSA	41/73 (56.2%)	9/21 (42.9%)	0.281
History of	8/73 (11.0%)	3/21 (14.3%)	0.676
pulmonary			
fibrosis			
History of	2/73 (2.7%)	0/21 (0%)	0.443
rheumatic			
fever/heart			
disease			
History of sickle	0/73 (0%)	0/21 (0%)	NA
cell trait or			
disease, or			
thalassemia			
History of thyroid	7/73 (9.6%)	1/21 (4.8%)	0.485
disease			
History of	1/73 (1.4%)	0/21 (0%)	0.590
splenectomy			

History of	1/73 (1.4%)	0/21 (0%)	0.590
myeloproliferative			
disease			
Ever smokers	66/73 (90.4%)	16/21 (76.2%)	0.085
Current smokers	16/73 (21.9%)	1/21 (4.8%)	0.136
Former smokers	50/73 (68.5%)	15/21 (71.4%)	
Never smokers	7/73 (9.6%)	5/21 (23.8%)	
Six minute walk	897 (281)	1220 (458)	0.058
distance (feet)	(n=53)	(n=9)	
Six minute walk	306 (96)	416 (156)	0.058
distance (meters)	(n=53)	(n=9)	
Forced expiratory	65 (21)	76 (20)	0.028
volume in one	(n=69)	(n=19)	
second (FEV1), %			
Forced vital	78 (19)	89 (17)	0.022
capacity (FVC), %	(n=69)	(n=19)	
FEV1/FVC ratio	61 (14)	64 (18)	0.310
	(n=69)	(n=19)	
Total lung	83 (16)	94 (18)	0.032
capacity (TLC), %	(n=66)	(n=18)	
Diffusion capacity	48 (20)	61 (24)	0.039

of the lungs for	(n=66)	(n=19)	
carbon			
monoxide, %			
Airflow	49/69 (71%)	10/19 (52.6%)	0.131
obstruction			
(FEV/FVC<70%)			
Restriction	28/66 (42%)	3/18 (16.7%)	0.045
(TLC<80%)			
Mixed obstruction	14/66 (21.2%)	1/17 (5.6%)	0.124
and restriction			
Right atrial	41/73 (56.2%)	9/20 (45%)	0.375
enlargement			
Right ventricular	6/73 (8.2%)	0/20 (0%)	0.185
hypertrophy			
Systolic septal	19/73 (26%)	1/20 (5%)	0.043
flattening			
Tricuspid plane	1.9 (0.6)	2.1 (0.6)	0.258
annular systolic	(n=63)	(n=16)	
excursion (cm)			
Left atrial size	29.8 (15.2)	26.3 (9.5)	0.483
volume index	(n=61)	(n=17)	

Left ventricular	56 (8)	58 (4)	0.312
ejection fraction	(n=73)	(n=21)	
(%)			
Reduced left	9/73 (12.3%)	0/21 (0%)	0.091
ventricular			
ejection fraction			
(<50%)			
Left ventricular	25/73 (34.2%)	8/20 (40%)	0.634
diastolic			
dysfunction			
present			
Pericardial	3/73 (4.1%)	0/20 (0%)	0.357
effusion present			
Aortic stenosis	12/73 (16.4%)	1/20 (5%)	0.191
Aortic stenosis	1 4/12 (33.3%)	1 0/1 (0%)	0.487
severity*	2 2/12 (16.7%)	2 0/1 (0%)	
	3 3/12 (25%)	3 1/1 (100%)	
	4 0/12 (0%)	4 0/1 (0%)	
	5 3/12 (25%)	5 0/1 (0%)	
Aortic	24/73 (32.9%)	6/14 (30%)	0.807

regurgitation			
Aortic	1 21/24 (87.5%)	1 6/7 (85.7%)	0.782
regurgitation	2 2/24 (8.3%)	2 1/7 (14.3%)	
severity*	3 1/24 (4.2%)	3 0/7 (0%)	
Mitral stenosis	0/73 (0%)	0/20 (0%)	NA
Mitral stenosis	NA	NA	NA
severity			
Mitral	35/73 (48%)	10/20 (50%)	0.871
regurgitation			
Mitral	1 30/37 (81.1%)	1 9/10 (90%)	0.793
regurgitation	2 3/37 (8.1%)	2 0/10 (0%)	
severity*	3 2/37 (5.4%)	3 1/10 (10%)	
	4 1/37 (2.7%)	4 0/10 (0%)	
	5 1/37 (2.7%)	5 0/10 (0%)	
Tricuspid	52/73 (71.2%)	18/20 (90%)	0.085
regurgitation			
Tricuspid	1 31/53 (58.5%)	1 12/18 (66.7%)	0.412
regurgitation	2 3/53 (5.7%)	2 0/18 (0%)	
severity*	3 9/53 (17.0%)	3 2/18 (11.1%)	
	4 4/53 (7.6%)	4 0/18 (22.2%)	
	5 6/53 (11.3%)	5 4/18 (22.2%)	

Estimated right	49 (13) (n=56)	43 (10) (n=15)	0.033
ventricular			
systolic pressure			
(mmHg)			
Estimated right	49/56 (87.5%)	9/15 (60.0%)	0.014
ventricular			
systolic pressure			
≥40mmHg			
Estimated right	16/56 (28.6%)	0/15 (0%)	0.019
ventricular			
systolic pressure			
≥60mmHg			

^{*1=}mild; 2=mild to moderate; 3=moderate; 4=moderate to severe; 5=severe

Supplemental Table 2: Changes in treatments and functional factors over time in patients with confirmed PH

	At baseline PH	At most recent	p value
	clinic visit	follow up visit	
Loop diuretic	42/68	57/68	0.0001**
prescription			
Loop diurectic	41 (53)	66 (59)	<0.001
dosage			
(furosemide			
equivalents):			
mean (SD)			
Change in	Mean increase of	24 (32)	
furosemide			
dose: mean (SD)			
Warfarin	20/68	22/68	0.41**
prescription			
Digoxin	3/68	4/68	0.56**
prescription			
Pulmonary	5/68	8/68*	0.41**
arterial			

hypertension-			
specific therapy			
prescription			
Supplemental	35/68	41/68	0.014**
oxygen			
prescription			
СРАР	28/68	38/68	0.004**
prescription			
Weight in lbs:	224 (56) (n=68)	218 (58) (n=64)	0.26
mean (SD)	221 (56) (n=64		
	with follow up		
	weight too)		
mMRC dyspnea	3 (2, 3) (n=67)	3 (2, 3) (n=58)	0.037***
score: median	3 (2, 3) (n=57	3 (2, 3) (n=57	
(IQR)	with mMRC	with mMRC	
	dyspnea score at	dyspnea score at	
	recent visit)	first visit)	
Functional	3 (2, 3) (n=48)	3 (2, 3) (n=50)	1.00 (NS)***
classification:	3 (2, 3((n=39	3 (2, 3) (n=39	
median (IQR)	with NYHA class	with NYHA class	
	at recent visit)	at first visit)	

Six minute walk	906 (278) (n=51)	887 (355) (n=26)	0.65
distance, feet	908 (262) (n=26		
	with follow up		
	6MWD)		
Six minute walk	274 (86) (n=53)	270 (108) (n=26)	
distance,	277 (80) (n=26)		
meters: mean			
(SD)			

^{*}An additional 4/68 were actively enrolled in clinical trial of a PAH-specific therapy vs. placebo

Paired t-test was performed to compare continuous variables.

^{**}McNemar test

^{***}Wilcoxson signed rank sum test