

Pulmonary Angioplasty in Chronic Thromboembolic Pulmonary Hypertension. Are Intravascular Ultrasound and Stents Helpful?



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Abstract

Pulmonary arterial angioplasty (PA) in chronic thromboembolic pulmonary hypertension (CTEPH) is an option for patients rejected for surgical pulmonary endarterectomy. Preliminary results seem encouraging, although pulmonary injury rates are reported. 11 patients (6 women) with CTEPH, mean age 64 ± 16 underwent PA. Four patients were in functional class 3, mean TAPSE was 15 ± 4 mm. Before the procedure, mean pulmonary pressure (mPAP) was 49 ± 13 mmHg, all had normal left ventricular ejection fraction and were under a chronic anticoagulant therapy. 10 out of 11 patients were done under intravascular ultrasound (IVUS) guidance in order to precisely estimate the vessel size. 20 stents were delivered to correct severe initial recoil or arterial occlusive dissection after initial balloon dilatation. There were no procedural complications (no regional pulmonary edema, no pulmonary vessel rupture). On follow-up (16.5 ± 6.4 months), we observed improvement in functional class in 9 out of the 11 patients. No hospital admission and no death were seen on follow-up. Pulmonary angioplasty under IVUS guidance seems to improve the procedure safety in our series, probably avoiding vessel rupture due to a more precise vessel sizing. Stent implantation seems to be safe and has no restenosis.

Keywords: Pulmonary hypertension; Pulmonary angioplasty; Chronic thromboembolic pulmonary hypertension; Intravascular ultrasound; Stents

Abbreviation: PA: Pulmonary Arterial Angioplasty; CTEPH: Chronic Thromboembolic Pulmonary Hypertension; mPAP: Mean Pulmonary Pressure; IVUS: Intravascular Ultrasound; PEA: Pulmonary Endarterectomy; BPA: Balloon Pulmonary Angioplasty; ESC/ERS: European Society of Cardiology/European Respiratory Society; OCT: Optical Coherence Tomography; PH: Pulmonary Hypertension

Introduction

Chronic thromboembolic pulmonary hypertension (CTEPH) is characterized by precapillary pulmonary hypertension (PH) hemodynamically defined as mean pulmonary arterial pressure (mPAP) ≥ 20 mmHg and a normal pulmonary artery wedge pressure ≤ 15 mmHg during resting right heart catheterization, with thromboembolic evidence showing mismatched perfusion defects in pulmonary arteries on computed tomographic pulmonary angiography, lung perfusion scintigraphy or pulmonary angiography despite effective anticoagulation therapy for at least three months. If left untreated, CTEPH will rapidly progress and seriously threaten people's lives [1], and CTEPH patients with

mPAP > 30 mmHg have a relatively poor prognosis [2], with a 5-year survival rate of 30% for those with mPAP > 40 mmHg and 10% for those with mPAP > 50 mmHg [3,4].

Pulmonary endarterectomy (PEA) remains the potentially curative treatment of choice with an in-hospital mortality rate of 2.2% for all eligible CTEPH patients at experienced centers and should be considered whenever possible [5]. Currently, inoperable candidates, either due to comorbidities or to distal arterial disease, have more options to receive medical and/or interventional therapies at specialized centers. Balloon pulmonary angioplasty (BPA) is a catheter-based interventional technique

which uses appropriately sized balloons to dilate and open up stenotic or occluded pulmonary vessels under angiography guidance. Accumulating evidence suggests that it can strikingly improve hemodynamics, cardiac function and exercise tolerance with an acceptable incidence of complications, and the long-term prognosis is encouragingly comparable to PEA surgery [6-10].

After more than 30 years of development and refinements, BPA is emerging as a promising alternative treatment option for CTEPH patients. However, there are still many issues worth exploring for BPA, such as the selection of patients, the establishment of a standardized operation process and the reduction of complications. After years of unremitting efforts, the 2015 European Society of Cardiology/European Respiratory Society (ESC/ERS) guidelines recommended that BPA only be performed in experienced CTEPH expert center [11]. Intravascular imaging techniques such as optical coherence tomography (OCT) [12,13] and intravascular ultrasound (IVUS) [14] offer endovascular information of the vessel wall and lumen that may potentially be useful for a better vessel selection and for a more adequate technique during BPA in order to increase its effectivity and decrease its complications.

OCT produces optically scattered images showing high-resolution internal tissue microstructure. OCT-guided BPA allows us to precisely assess lesion type and vessel diameter and facilitate the selection of appropriate balloon size and length and can also be utilized to evaluate the effectiveness of BPA; however, it may result in potential volume overload and right heart failure [15-18]. IVUS is an endovascular imaging technique that uses a miniaturized ultrasound probe to generate sound waves and produce real-time intravascular images, and due to relatively lower resolution mainly applied to determine vessel diameter so as to select the appropriate balloon. In addition, virtual histology IVUS is capable of identifying vascular lesions vulnerable to balloon compression and helps us decide which organized thrombus can be treated to avoid potential pulmonary artery rupture [17].

The world largest multicenter registry from Japan described pulmonary injury (17.8%) and hemoptysis (14.0%) as the most frequent complications, pulmonary artery perforation, dissection and rupture accounted for 2.9%, 0.4%, and 0.1%, respectively [6]. The recent French BPA experience revealed that lung injury, hemoptysis, pulmonary artery perforation and dissection occurred in 9.1%, 7.1%, 2.8, and 1.9%, respectively, among 1006 BPA sessions, and the incidence of lung injury decreased over time from initial 13.3% to recent 5.9%, clearly indicating that the occurrence of BPA complications depends on center experience [7]. Pulmonary artery stenting may be an option in proximal CTEPH when elastic recoil phenomenon makes balloon angioplasty of a large vessel ineffective [19]. It has been suggested that IVUS-guided pulmonary angioplasty in chronic thromboembolic pulmonary hypertension may increase safety of the procedure [20].

Case Series

We here in describe a case series of IVUS-guided BPA in CTEPH where the IVUS information was useful for the effectiveness and safety of the procedure. Patients submitted to pulmonary angioplasty were either patients with distal disease not candidates for surgery or patients rejected for surgery due to comorbidities despite being angiographically good surgical candidates. No peri nor post procedural complications were observed (no regional pulmonary edema, no pulmonary vessel rupture). All patients were dismissed the day after the procedure. Right heart catheterization was performed through the femoral vein in 7 and through the brachial vein in 13 procedures. 11 patients were submitted to 20 procedures and a total of 28 lesions were treated. IVUS guided PA (Boston Scientific, Opti Cross 40MHz) was performed in 10 out of 11 patients and in 17 out of 20 procedures. Stents were delivered in 10 out of 28 lesions, due to severe immediate elastic recoil or due to angiographic occlusive dissection after initial balloon dilatation.

Basal clinical and hemodynamic data are listed in (Table 1). Procedural variables are listed in (Table 2). Location of dilated lesions is showed in (Figure 1). No immediate or mid-term complications were observed (mean follow up 16.5 ± 6.4 months). Baseline functional class was II in 35% and III in 65% of patients. 9 patients improved at least 1 functional class (6 improved 1 and 3 patients 2) (p<0.01) and 2 remained stable. Only a modest improvement was observed in mean PAP (49.9mmHg vs. 42.8mmHg), and BNP (727 vs. 222pg/dl). No angiographic stent restenosis was seen in angiographic control. No clinical or angiographic complications related to stent delivery were detected on the control angiography at 3-6 months follow-up. Examples of the peri procedural usefulness of IVUS (depicting the mechanisms of arterial occlusion after initial balloon dilatation) are depicted in (Figure 2A, 2B and 2C). Example of stent under expansion not evident in the arterial angiography and showed by the endovascular IVUS image is depicted in (Figure 3).

Table 1: Baseline characteristics.

Patients (n)	11
Lesions treated (n)	28
Age (years)	64.45 ± 16
Sex (%female/male)	55/45
NYHA functional class (I/II/III/IV) (n)	0/4/7/0
TAPSE (mm)	15,1 ± 4
BNP (pg/ml)	727 ± 1186
Right heart catheterisation parameter:	
Mean sPAP (mmHg)	49,9 ± 13
Mean PVR (UW)	11,3 ± 5
RAP (mmHg)	12.7 ± 3

Table 2: Procedural variables.

	Nº procedures	IVUS	Lobes	Total Lesions	PCI	Highest Diameter Device (mm)
PATIENT 1	1	IVUS guided	Inferior left (2)	2	BA	3,0
PATIENT 2	1	IVUS guided	Inferior left (2)	2	BA + 2 DES	6,0
PATIENT 3	3	IVUS guided	Med. right lobe (1) Inferior left (2)	3	BA + BMS	4,5
PATIENT 4	1	IVUS guided	Inferior left (2)	2	BA + BMS	7,0
PATIENT 5	2	IVUS guided	Inferior left (2)	2	BA	2,5
PATIENT 6	2	IVUS guided	Inferior left (1)	1	BA + DES	4,5
PATIENT 7	2	IVUS guided	Inferior left (2) Right superior (1)	3	BA + 2 DES	5,0
PATIENT 8	3	IVUS guided	Inferior left (2) Right superior (4) Right Inferior (1)	7	Simple BA (x3)	4,0
PATIENT 9	1	IVUS guided	Inferior left (1)	1	BA + 1 DES	5,0
PATIENT 10	2	IVUS guided	Right Inferior (3)	3	BA + 2 DES	8,0
PATIENT 11	2	No	Inferior left (2)	2	BA	5,0

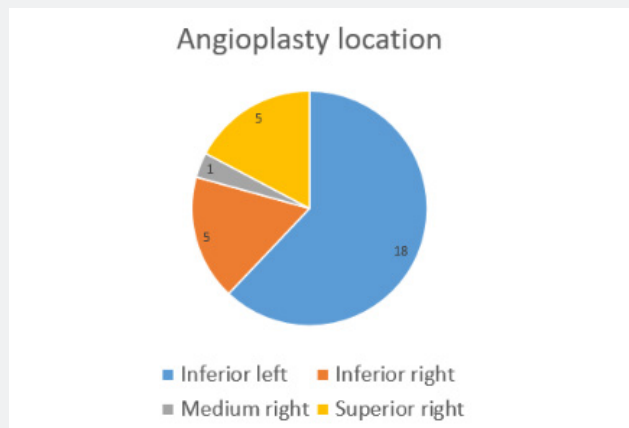


Figure 1: Location of the treated lesions.

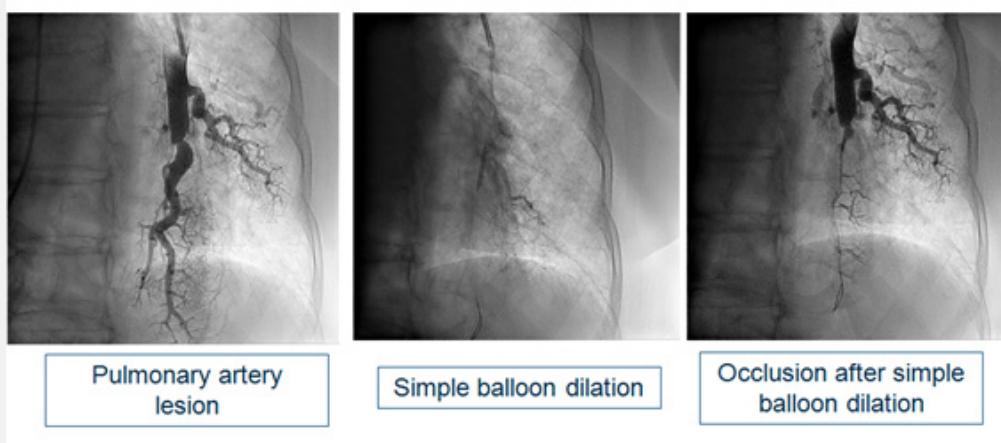


Figure 2A: Example of initial total occlusion after initial balloon dilatation.

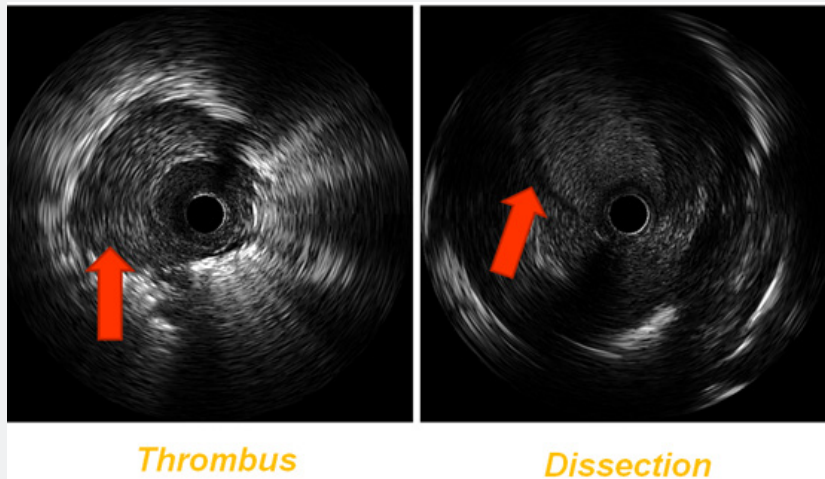


Figure 2B: IVUS shows clearly arterial dissection with thrombus formation.

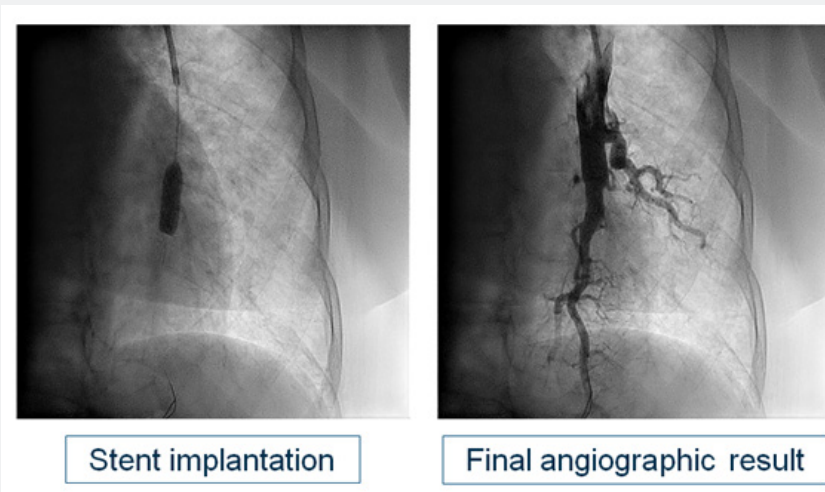


Figure 2C: Stent implantation with good final result.

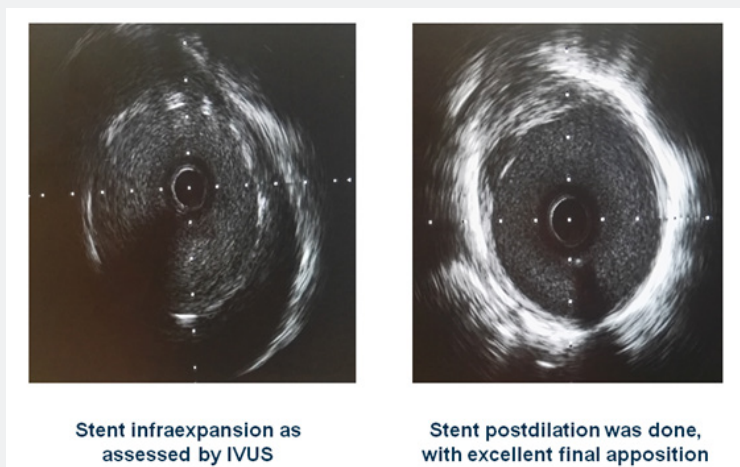


Figure 3: Example of stent infra expansion showed by IVUS and solved by stent post dilatation.

Conclusions

IVUS allows a better vessel sizing of pulmonary artery and thus a more accurate selection of balloon and stent diameters. IVUS is better than angiography alone for detecting balloon post dilatation recoil and stent under expansion. IVUS guided PA may also detect potential complications, such as arterial dissection, differentiating true vs false lumen, thrombus formation, and may increase the safety of the procedure. No angiographic stent restenosis was seen in angiographic control.

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