## Le Cathétérisme Droit dans l'HTAP: Pourqoi est-ce Indispensable?

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## Diagnostic Work-up in Pulmonary Hypertension

- Chest x-ray Pulmonary function tests Echocardiography Laboratory evaluation ♦ ECG Exercise testing ♦ CT Ventilation-perfusion scan Angiography
- Right heart catheterization

## Hemodynamic Data Obtained with Doppler-Echocardiography

#### Volumetric measurements

- Stroke volume and cardiac output
- Regurgitation volume and fraction
- Pulmonary-systemic flow ratio (Qp/Qs)

#### Pressure gradients

- Maximal instantaneous gradient
- Mean gradient

#### Valve area

- Stenotic valve area
- Regurgitant orifice area

#### Intracardiac pressures

- Pulmonary artery pressures
- Left atrial pressure
- Left ventricular end-diastolic pressure

## Problems of Hemodynamic Measurements in Echocardiography

#### No absolute pressure

No direct flow measurement

#### Dependent on quality of echo signal

–PHTN may be underestimated or missed in the presence of a poor signal

- In apical view mitral regurgitation or aortic stenosis signals could be falsly interpreted as tricuspid signals
- Not reliable for PAP measurement in the presence of pulmonary stenosis

### How Good is the Estimation of PA Pressure by Tricuspid Regurgitation Velocity?

#### ? RVSP = Gradient

- RVSP = Gradient + 10 mmHg
- ? RVSP = Gradient + RAP estimated on clinical grounds
- RVSP = Gradient + RAP estimated by cava index

## **Estimation of RA Pressure Based on Diameter of the IVC**

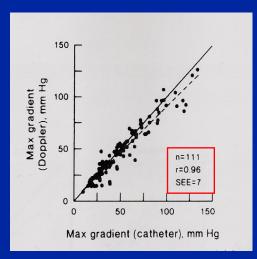
IVC Diameter	Changes IVC Diameter with Inspiration	RA Pressure Estimation (mmHg)	
Small (<1.5 cm)	Collapse	0-5	
Normal (1.5-2.5 cm)	>50% ↓	5-10	
Normal (1.5-2.5 cm)	<50% ↓	10-15	
Dilated (>2.5 cm)	<50% ↓	15-20	
Dilatation also of the hepatic veins	no change	>20	

C. Otto. The Practice of Clinical Echocardiography. 2002

## **Correlation Doppler – Invasive Measurement**

Yock P et al. Circulation 1984;70:657-62 Currie PJ et al. JACC 1985;6.750-6 Stevenson JG JASE 1989;2:157-71

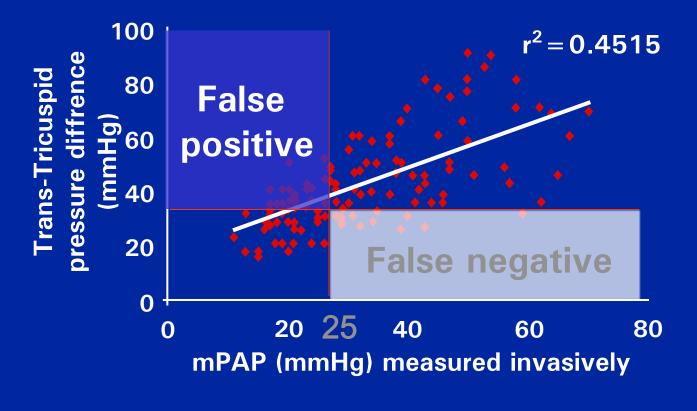
Auteur	n	r	SEE mmHg
Yock 1984	62	0.95	7
Currie 1985	111	0.90	8
Stevenson 1989	50	0.96	6.9



Estimated pressure 50 mmHg  $\rightarrow$  95% confidence limits 34-66 mmHg

Tricuspid regugitant jet estimation
only in 50-60% of patients with no PHTN
only in 80-90% of patients with PHTN

## Doppler Echocardiography vs Invasive Pressure Measurements



Barst RJ, et al. J Am Coll Cardiol 2004; 43:40S-7S.
 Mukerjee D, et al. Rheumatology 2004;43:461-6.

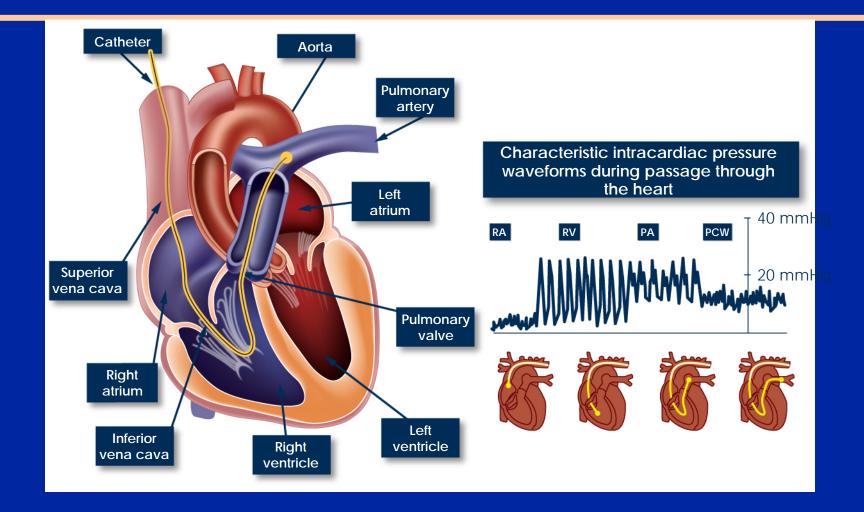
## **Right Heart Catheterization**



#### Cardiac Catheterisation → Essential in the Diagnosis and Management of PHTN

- Diagnostic gold standard
- Confirms the diagnosis of PHTN
- Describes the haemodynamic mechanism (e.g. PAH vs left heart disease)
- Determines severity (CO, RAP, mixed venous oxygen saturation)
- Testing for vasoreactivity
- Overall procedure-related mortality 0.055% (95% CI, 0.01%–0.099%): 4/7218

## **Right Heart Catheterization**

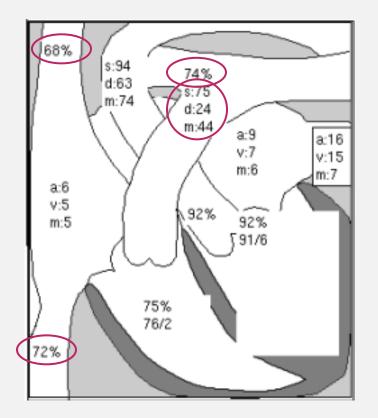


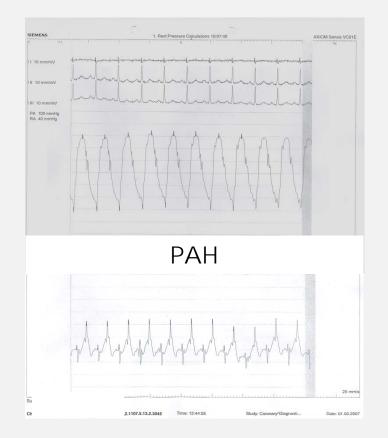
## **Goals of Invasive Assessment**

- Confirm non-invasive estimation of pulmonary pressures
- Measurement of pressures and saturations in all heart chambers
- Find etiology of PHTN (e.g., shunts)
- Test vasoreactivity
- Plan therapy
- Assess prognosis

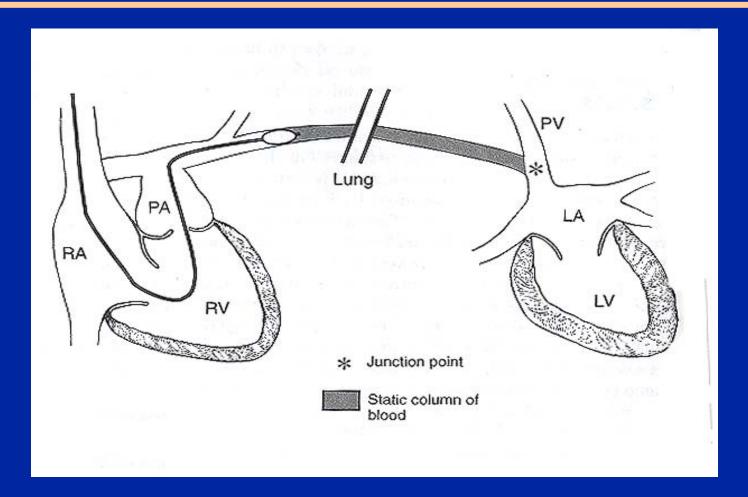
#### Right Heart Catheterization → Insight into Pulmonary Hemodynamics: Pressures, Flow State, Resistances

#### To rule out shunts-droit



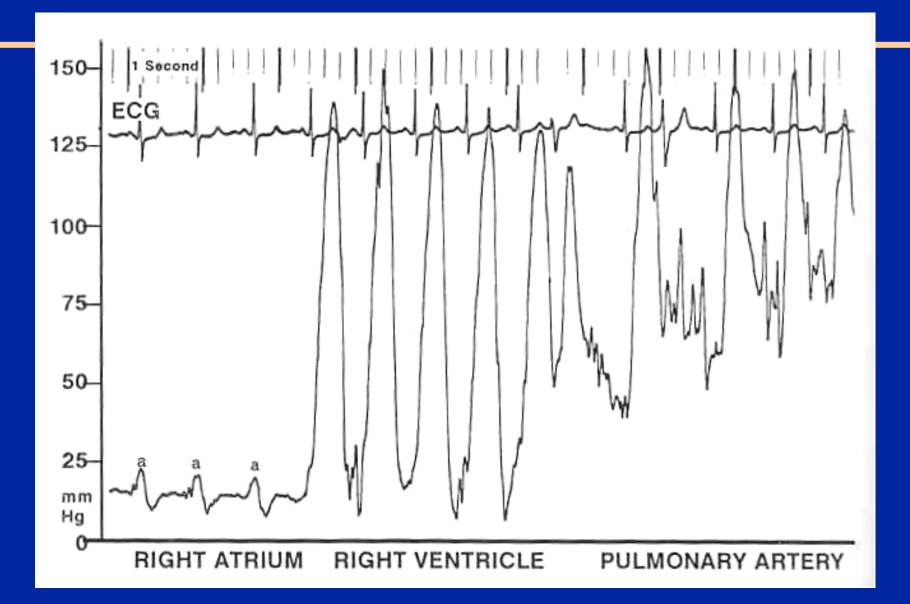


## Pulmonary Artery Wedge Pressure Measurement

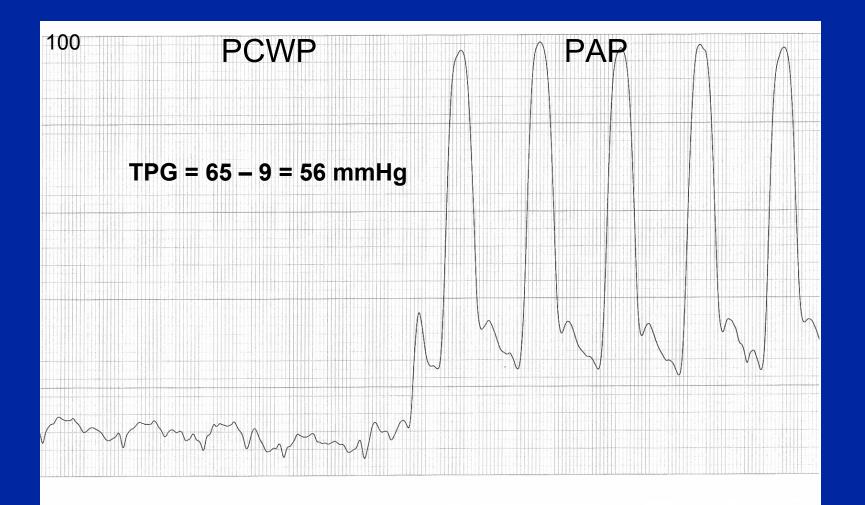


#### Transpulmonal gradient = mean PAP – mean PCWP

## **Right Heart Catheter**



## Transpulmonal Gradient (TPG) = mean PA pressure - PCWP



## **Right Heart: Normal Hemodynamics**

- Syst. PA pressure 18 25 mmHg
- Diast PA pressure 6 10 mmHg
- Mean PAP 12 16 mmHg

PCWP 6 – 10 mmHg

 $\frac{\text{Mean PAP} - \text{PCWP}}{\text{Cardiac Output}} \times 80 = 60-120 \text{ dyn.sec.cm}^{-5}$ 

## **Right Heart Catheterization in PAH**

#### Increased mPAP

- normal mPAP < 20 mmHg; PAH defined as</p>
- mPAP > 25 mmHg
- Normal PCWP
  - normal range <15 mmHg</p>
- PVR↑, > 3 Wood units (250 dyn/sec/cm<sup>-5</sup>)\*
   J
- Right atrial pressure↑
  - normal right atrial pressure 2–7 mmHg
- ◆ Cardiac output↓
  - normal cardiac output 4–8 liters per minute
- ◆ Cardiac index↓
  - normal cardiac index 2.5–4.0 liters/min/m<sup>2</sup>

• "25–15–3" rule

#### \*Gradient DPAP-Wedge < 6mmHg

## Acute vasodilator responsiveness in different forms of PAH

	N° of patients tested*	Acute responders# (n, %)	Long-term responders to CCB
Sporadic PPH	430	57 (13 %)	27 (6.3 %)
Appetite suppressant	127	13 (10 %)	10 (7.9 %)
Connective Tissue D.	166	15 (9 %)	2
PVOD / PCH	34	3 (9 %)	0
HIV-associated PAH	123	2 (1.5 %)	1
Portopulmonary Ht.	153	1 (0.6 %)	0
Congenital Heart D.	41	0	NA
Familial PPH	34	. 0	NA
* With NO and/or $PgI_2$	# fall in mPAP and PVR > 20%		Personal unpublished data

Sitbon Venice 2003

## **Acute Vasodilation Testing**

<u>DRUG</u>	<u>ROUTE</u>	<u>DOSE</u>	<u>HALF LIFE</u>
NO	inhaled	10-20ppm	15-30s
Adenosine (USA)	iv	50-200µg/kg/min	5-10s
Nifedipine	ро	10-40mg	2 h
Epoprostenol	iv	2-20ng/kg/min	3-5min
Iloprost	inhaled	10-20µg	45 min
Bosentan	ро	62.5-125mg	5 h
Sildenafil	ро	25-100mg	60 min

## PHTN: Positive Vasodilator Response

Decrease of mean pulmonary artery pressure by  $\geq$ 10 mmHg to reach  $\leq$ 40 mmHg with an increased or unchanged cardiac output.

= new definition (Dana Point 2008)

## Importance of Vasoreactivity Testing

 Initiation of vasodilator therapy
 Surgical closure of shunts in congenital disease
 Detection of right ventricular dysfunction European Heart Journal (2009) **30**, 2493–2537 doi:10.1093/eurheartj/ehp297

#### **ESC/ERS GUIDELINES**

#### Guidelines for the diagnosis and treatment of pulmonary hypertension

The Task Force for the Diagnosis and Treatment of Pulmonary Hypertension of the European Society of Cardiology (ESC) and the European Respiratory Society (ERS), endorsed by the International Society of Heart and Lung Transplantation (ISHLT)

Authors/Task Force Members: Nazzareno Galiè (Chairperson) (Italy)<sup>\*</sup>; Marius M. Hoeper (Germany); <u>Marc Humbert (France);</u> Adam Torbicki (Poland); Jean-Luc Vachiery (France); Joan Albert Barbera (Spain); Maurice Beghetti (Switzerland); Paul Corris (UK); Sean Gaine (Ireland); J. Simon Gibbs (UK); Miguel Angel Gomez-Sanchez (Spain); Guillaume Jondeau (France); Walter Klepetko (Austria) Christian Opitz (Germany); Andrew Peacock (UK); Lewis Rubin (USA); Michael Zellweger (Switzerland); Gerald Simonneau (France)

# Table I IRecommendations for right heartcatheterization (A) and vasoreactivity testing (B)

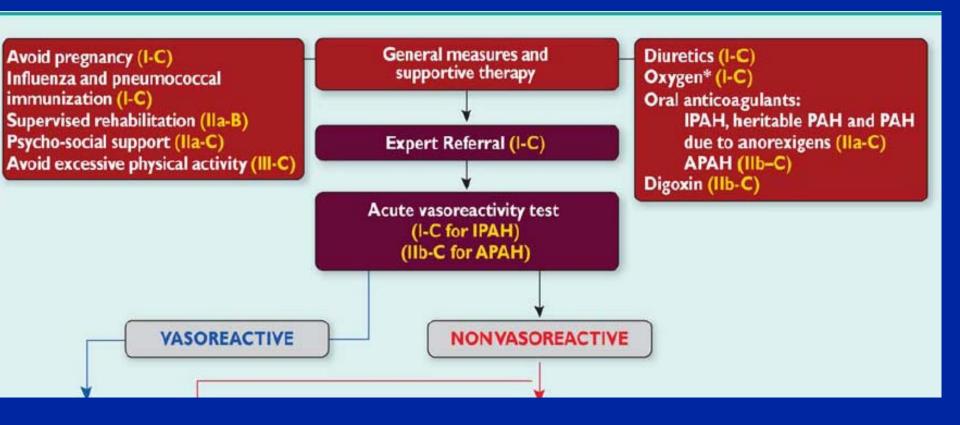
	<b>C</b> lass <sup>a</sup>	Level <sup>b</sup>
A		
RHC is indicated in all patients with PAH to confirm the diagnosis, to evaluate the severity, and when PAH specific drug therapy is considered	I	С
RHC should be performed for confirmation of efficacy of PAH-specific drug therapy	lla	С
RHC should be performed for confirmation of clinical deterioration and as baseline for the evaluation of the effect of treatment escalation and/or combination therapy	lla	С

A positive response to vasoreactivity testing is I C
 defined as a reduction of mean PAP
 ≥ 10 mmHg to reach an absolute value of mean
 PAP ≤40 mmHg with an increased or
 unchanged CO
 Vasoreactivity testing should be performed only in IIa C
 referral centres
 Vasoreactivity testing should be performed using IIa C
 nitric oxide as vasodilator

#### **Dedicated interventionalists (HUG → Dr. Keller)**

- Indication for the cath discussed in the multidisciplinary PHT team
- Knows about the patients
- Knows the specific question that the invasive test is suppose to answer
- Is able to integrate the results in the clinical contaxt

## Central Role of Cardiac Catheterization and Vasoreactivity Test



## **Goals of Invasive Assessment**

 Confirm non-invasive estimation of pulmonary pressures

- Measurement of pressures and saturations in all heart chambers
- Find etiology of PHTN
- Test vasoreactivity
- Plan therapy
- Assess prognosis

## **Hemodynamic Classification**

Class	Symptoms	Echocardio-	RV
		graphy	Catheterization
Mild	NYHA I	Syst. PAP 35- 55 mmHg	Mean PAP 21- 40 mmHg
Moderate	NYHA II	Syst PAP > 55 mmHg	Mean PAP > 40 mmHg
Severe	NYHA III	RV function impaired	SVO <sub>2</sub> < 60 %
Very severe	NYHA IV	RV function severely impaired	SVO <sub>2</sub> < 50 %

## Hemodynamic Adverse Prognostic Indicators in Primary Pulmonary Hypertension

#### Pulmonary arterial oxygen saturation < 63%</li>

- >63%: 55% survival at 3 years
- < 63%: 17% survival at 3 years</p>

#### Cardiac index < 2.1 l/min/m<sup>2</sup>

- < 2.1: 17 months median survival

#### Right atrial pressure > 10 mmHg

- < 10 mmHg: 4 years mean survival</p>
- > 20 mmHg: 1 month mean survival

 Lack of pulmonary vasodilator response to acute challenge

## **Prognostic Implications**

Better prognosis	Determinants of prognosis	Worse prognosis
No	Clinical evidence of RV failure	Yes
Slow	Rate of progression of symptoms	Rapid
No	Syncope	Yes
I, II	WHO-FC	IV
Longer (>500 m) <sup>a</sup>	6MWT	Shorter (<300 m)
Peak O <sub>2</sub> consumption >15 mL/min/kg	Cardio-pulmonary exercise testing	Peak O <sub>2</sub> consumption <12 mL/min/kg
Normal or near-normal	BNP/NT-proBNP plasma levels	Very elevated and rising
No pericardial effusion TAPSE <sup>b</sup> >2.0 cm	Echocardiographic findings <sup>b</sup>	Pericardial effusion TAPSE <sup>b</sup> <1.5 cm
RAP <8 mmHg and CI ≥2.5 L/min/m <sup>2</sup>	Haemodynamics	RAP >15 mmHg or CI ≤2.0 L/min/m <sup>2</sup>

### Conclusions: Why is Right Heart Catheterization Necessary

RHC is indicated in all patients with PAH to confirm the diagnosis, to evaluate the severity, and when PAH specific drug therapy is considered

### Conclusions: Why is Right Heart Catheterization Necessary

- Pressure measurement not estimation
  - On Echo PHTN cannot be estimated in in 50-60% of patients with no PHTN and in 80-90% of patients with PHTN
- Allows to exclude "treatable" causes of PHTN (shunts)
- Can differentiate PHTN related or not related to LV dysfunction
- Insight into pulmonary hemodynamics: pressures, flow state, resistances
- Invasive, but low complication rates
- Vasoreactivity testing by non-invasive measurements not reliable
   Planning of therapy without vasoreactivity test questionable
- Has prognostic implications at the time of diagnosis
- To follow the patient response to vasodilator therapy if the clinical evolution and the echocardiographic parameters are discordant:if PHTN stable or decreasing but also the cardiac output is decreasing the prognosis is poorue.
- ◆ Prognosis of the patient with severe PAHT unfavorable → the highest degree of accuracy for diagnosis and assessment of vasoreativity is indicated

## Last but not least

- Indication, interpretation, and therapeutic consequences of right heart catheterization and vasoreactivity nedd to be discussed nin multidicliplinary fashion
- Right heart catheterization should be done by a « dedicated » interventionalcardiologist
- Cardiac catheterization should be performed in a dedicated pulmonary hypertension center