



**VII CONGRESSO CATARINENSE  
DE OBSTETRÍCIA E GINECOLOGIA**  
II Congresso Catarinense de Perinatologia

25 a 27 de junho de 2015 | Expoville | Joinville | SC

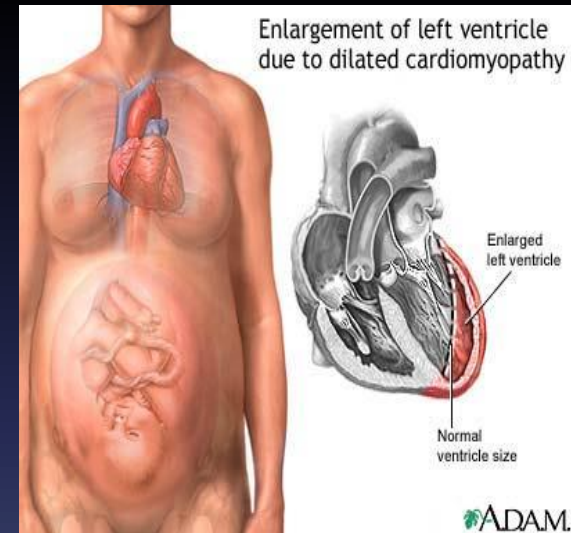
**Conrado Roberto Hoffmann Filho**

**Declaração de conflito de interesse**

Não recebi qualquer forma de pagamento ou auxílio financeiro de entidade pública ou privada para pesquisa ou desenvolvimento de método diagnóstico ou terapêutico ou ainda, tenho qualquer relação comercial com a indústria farmacêutica

# Cardiomiopatia periparto

- diagnóstico, manejo e repercussões maternas e fetais



Declaro não apresentar conflito de interesses



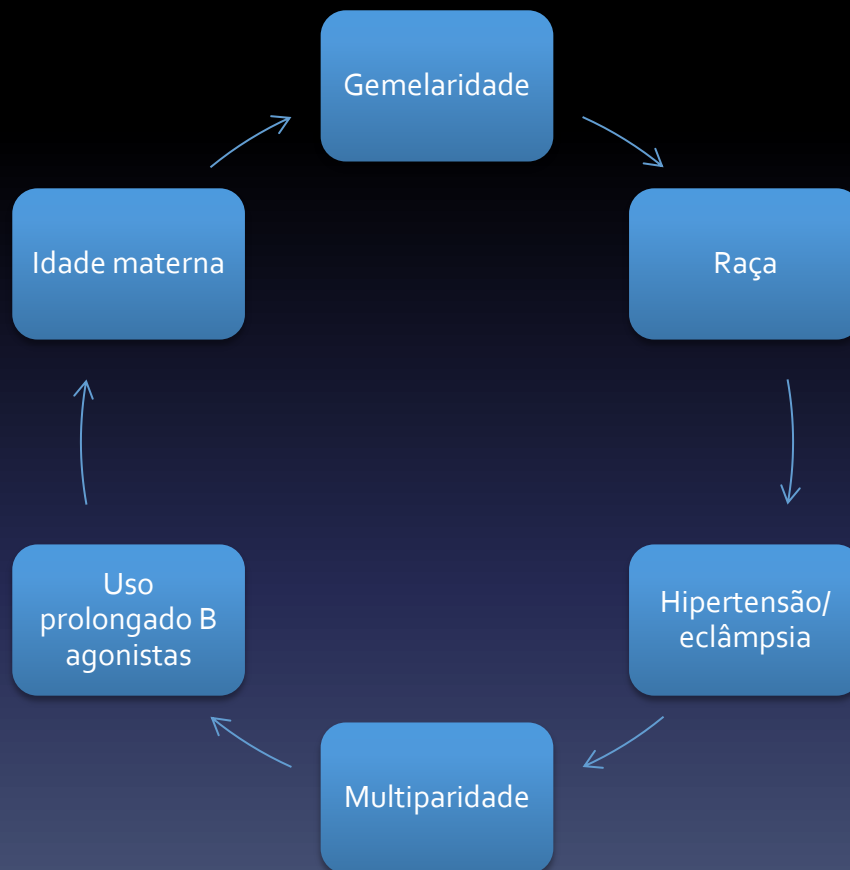
# Definição

Miocardiopatia idiopática, com quadro de insuficiência cardíaca, que se apresenta entre o último mês de gestação e até o quinto mês pós parto, sem outras causas de ICC encontradas, apresentando fração de ejeção reduzida, usualmente  $< 45\%$ . Diagnóstico de exclusão.

Heart Failure Association of the ESC Working Group on PPCM (Sliwa *et al.*, 2010)

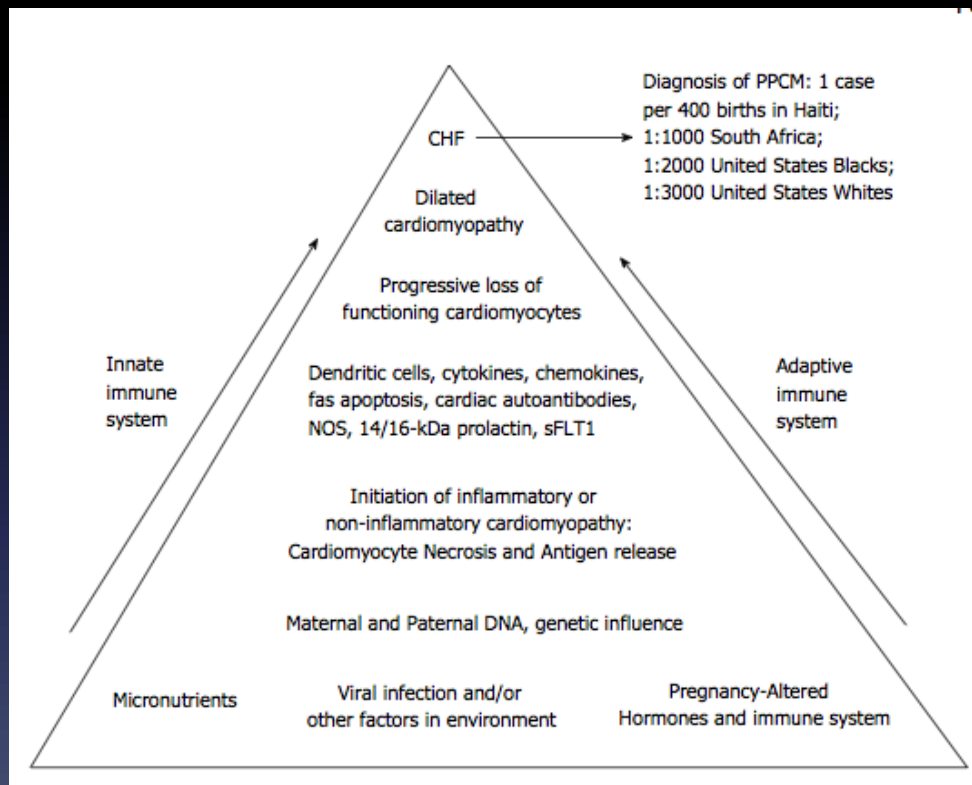
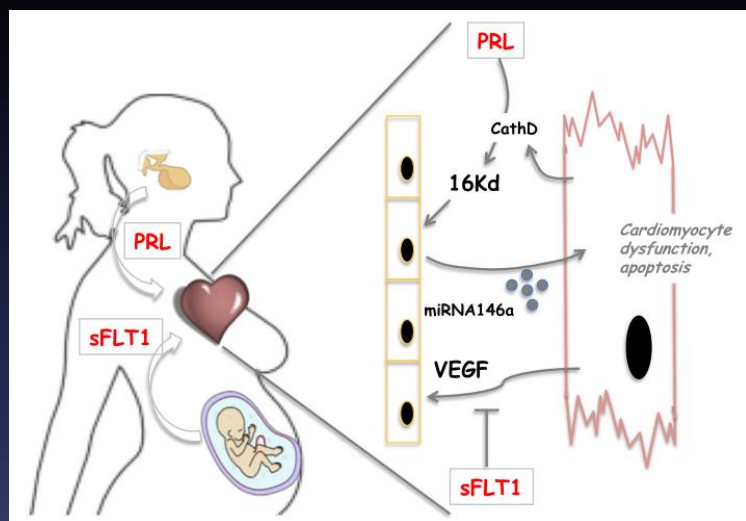


# Epidemiologia: Fatores predisponentes



EURObservational Research Programme: a worldwide registry on peripartum cardiomyopathy (PPCM) European Journal of Heart Failure (2014) 16, 583–591

# Epidemiologia: Etiologia

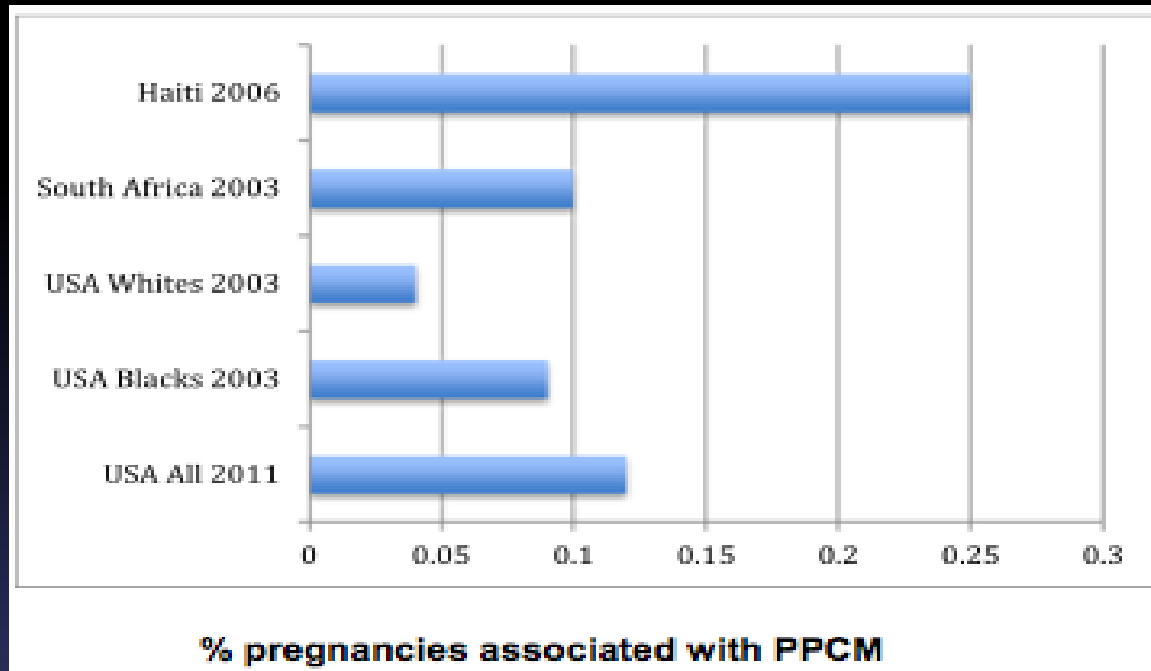


Natalie A. Bello and Zoltan Arany Molecular mechanisms of peripartum cardiomyopathy: A vascular/hormonal hypothesis. Trends in cardiovascular medicine (2015)

Fett JD. Peripartum cardiomyopathy: A puzzle closer to solution. *World J Cardiol* 2014; 6(3): 87-99



# Incidência



60% iniciam 2 meses pós parto  
7% no último trimestre de gravidez

James D. Fett and David W. Markham Discoveries in peripartum cardiomyopathy. Trends in cardiovascular medicine (2014)



# Diagnóstico

Apresentação clínica habitualmente não reconhecida pela confusão dos sintomas da gravidez serem semelhantes ao da IC

Na ausência de sintomas cardíacos observar o crescimento fetal

James D. Fett and David W. Markham Discoveries in peripartum cardiomyopathy. Trends in cardiovascular medicine (2014)



# Diagnóstico

## Sintomas

Dispneia  
Ortopneia  
DPN  
Tosse  
Dor torácica  
Palpitações  
Dor abdominal

## • Sinais

- Crepitações
- Edema
- Ritmo de galope
- Sopro holossistólico

## Manifestação tromboembólica

James D. Fett and David W. Markham Discoveries in peripartum cardiomyopathy. Trends in cardiovascular medicine (2014)





# Diagnóstico

## Auto teste para diagnóstico de miocardiopatia periparto

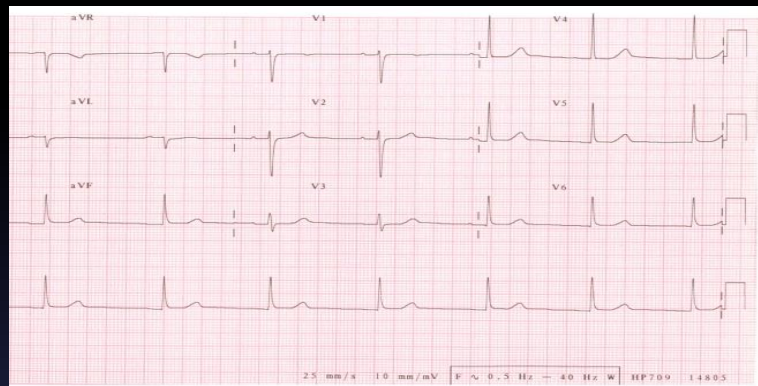
Self-test for early diagnosis of HF in peripartum cardiomyopathy

Symptom	Severity		
Orthopnea (difficulty breathing when lying flat)	None 0 points	Need to elevate head 1 point	Need to elevate $\geq 45^\circ$ 2 points
Dyspnea (shortness of breath on exertion)	None 0 points	Climbing $\geq 8$ steps 1 point	Walking on level 2 points
Unexplained cough	None 0 points	At night 1 point	Day and night 2 points
Swelling (pitting edema) lower extremities	None 0 points	Below knee 1 point	Above and below knee 2 points
Excessive weight gain during last month of pregnancy	<2 lb per week 0 points	2-4 lb per week 1 point	>4 lb per week 2 points
Palpitations (sensation of irregular heartbeats)	None 0 points	When lying down at night 1 point	Day and night, any position 2 points

Total score greater than 4 points suggests a need for further evaluation.

Fett JD. Validation of a self-test for early diagnosis of heart failure in peripartum cardiomyopathy. Crit Pathw Cardiol 2011;10:44-5

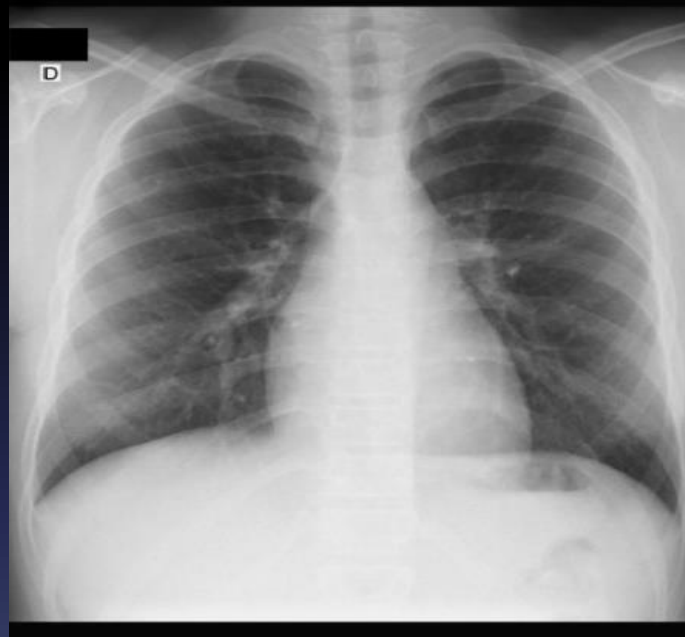
# Diagnóstico



Alterações inspecíficas de ST e T

Arritmias atriais e ventriculares

Distúrbios de condução



Sem cardiomegalia

Congestão venosa e edema pulmonar

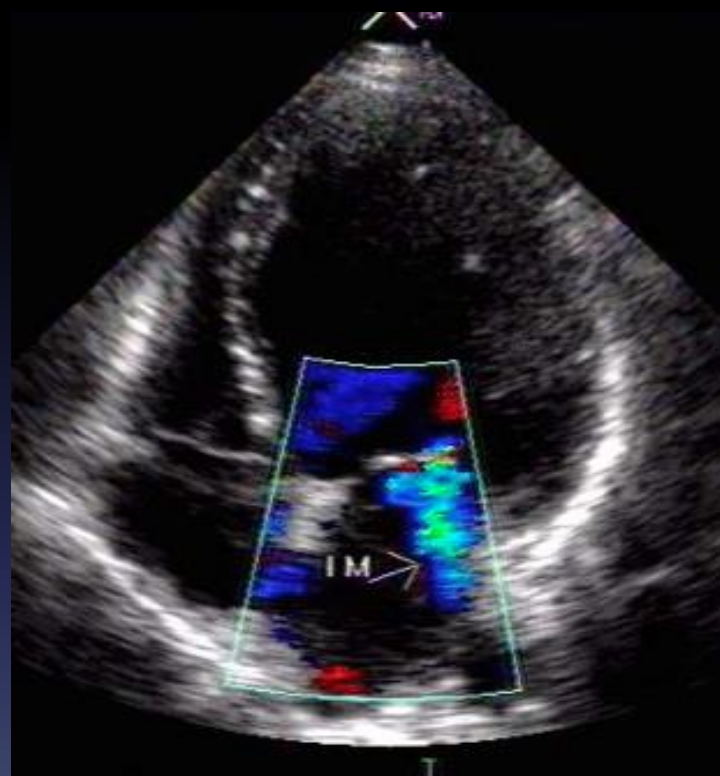


# Miocardiopatia periparto: Diagnóstico

Aumento das 4 câmaras, com diminuição importante da função do VE

Regurgitações mitral, tricúspide e pulmonar

Fração de ejeção, débito cardíaco e movimentação das paredes diminuídos



3. Sliwa K, Fett J, Elkayam U. Peripartum cardiomyopathy. Lancet 2006;368:687 – 693



# Diagnóstico

**Table 1** Overview of biomarkers analysed in peripartum cardiomyopathy patients

Biomarker	Relevance for PPCM
NT-proBNP	Not specific for PPCM, but good sensitivity for heart failure. <sup>23,39</sup>
troponin I and troponin T	Pathophysiological factor of PPCM, high technical effort for measurement, diagnostic accuracy needs to be evaluated. <sup>37,42</sup>
Interferon- $\gamma$	Elevated plasma levels in PPCM patients, diagnostic accuracy needs to be evaluated. <sup>37,49</sup>
Asymmetric Dimethylarginine (ADMA)	Marker for endothelial dysfunction and cardiovascular risk, diagnostic accuracy needs to be evaluated. <sup>23</sup>
Cathepsin D	Activity elevated in plasma of PPCM patients, diagnostic accuracy needs to be further evaluated. <sup>23,37</sup>
Soluble fms-like tyrosine kinase-1 (sFlt-1)	Elevated plasma levels in PPCM patients, diagnostic accuracy needs to be further evaluated. <sup>33</sup>
microRNA-146a	Pathophysiological factor of PPCM, high technical effort for measurement, diagnostic accuracy needs to be further evaluated. <sup>23,38</sup>

BNP



# Miocardiopatia periparto Tratamento

Cardiologista

Obstetra de alto  
risco

Anestesiologista

Neonatalogista

Peripartum cardiomyopathy: current management and future perspectives Denise Hilfiker-Kleiner European Heart Journal (2015) 36, 1090–1097



# Miocardiopatia periparto Tratamento

## Tratamento vigoroso da ICC

Não farmacológico

Restrição de água e sal

Farmacológico

Redução pré carga ( nitratos, diuréticos )

Redução pós carga ( nitratos, amlodipina. Hidralazina)

Inotrópicos ( dopamina. dobutamina. digoxina)

Beta bloqueadores

Anticoagulação

Deambulação precoce

Peripartum cardiomyopathy: current management and future perspectives Denise Hilfiker-Kleiner European Heart Journal (2015) 36, 1090–1097



# Miocardiopatia periparto Tratamento

Drug	Safety during lactation <sup>a</sup>	Absence of complete recovery	Complete and sustained recovery of left-ventricular structure and function (echocardiographic follow-up every 6 months)			
			6 months	6–12 months	>12 months	>18 months
β-Blocker	Bradycardia of newborn rare in rare cases Metoprolol is best-studied β-blocker during lactation.	<b>Complete and sustained recovery of left-ventricular structure and function (echocardiographic follow-up every 6 months)</b>	<b>6 months</b>	<b>6–12 months</b>	<b>&gt;12 months</b>	<b>&gt;18 months</b>
ACE-inhibitor	Low transfer of enalapril and captopril into breast milk.	Continue all drugs for at least 6 months after full recovery to avoid relapse	Continue β-blocker and ACE-inhibitor/ARB for at least 6 months after stopping MRA	Continue β-blocker for at least 6 months after stopping ACE-inhibitor/ARB	Discontinue β-blockade, ensure echocardiographic follow-up	Discontinue β-blockade, ensure echocardiographic follow-up
ARB	Very limited data ARB during lactation and be avoided.	ACE-inhibition. Up-titration to standard or maximally tolerated dosages.		Reduce dosage and then discontinue ACE-inhibitor/ARB		

Peripartum cardiomyopathy: current management and future perspectives Denise Hilfiker-Kleiner European Heart Journal (2015) 36, 1090–1097



# MiocardioPatia periparto Tratamento

Drug	Safety during lactation <sup>a</sup>	Absence of complete recovery	Complete and sustained recovery of left-ventricular structure and function (echocardiographic follow-up every 6 months)			
			6 months	6–12 months	>12 months	>18 months
MRA	Very limited data on MRA during lactation and should be avoided	Recommended for all patients with LVEF < 40%. Eplerenone may be considered due to less hormonal side effects.			Discontinue only if complete and sustained recovery of left-ventricular structure and function	
Ivabradine	No data on ivabradine during lactation available and should be avoided.	For patients with heart rate > 75/min, when β-blocker up-titration is not possible. Should be tapered when β-blocker up-titration is possible and/or heart rate is < 60/min	Continue when heart rate is > 75/min despite β-blocker up-titration		Discontinue only if complete and sustained recovery of left-ventricular structure and function	
Diuretics	Thiazides are the best-studied diuretics during lactation and well tolerated. They may decrease milk production. Very limited data on furosemide and	Only when oedema/ congestion is present. Early tapering of dose according to symptoms, even before full recovery of left-ventricular function	Continue only when symptoms (congestion/oedema) are present without diuretic therapy as part of an antihypertensive drug therapy			

Peripartum cardiomyopathy: current management and future perspectives Denise Hilfiker-Kleiner European Heart Journal (2015) 36, 1090–1097





# Miocardiomatia periparto Tratamento

Ressicronização cardíaca

Cardiodesfibrilador

Assistência ventricular esquerda

Transplante cardíaco

Peripartum cardiomyopathy: current management and future perspectives Denise Hilfiker-Kleiner European Heart Journal (2015) 36, 1090–1097



# Miocardiopatia periparto Tratamento

## Pentoxifilina ??

**Rationale and design of a randomized, controlled multicentre clinical trial to evaluate the effect of bromocriptine on left ventricular function in women with peripartum cardiomyopathy**

Arash Haghikia<sup>1</sup> · Edith Podewski<sup>1</sup> · Dominik Berliner<sup>1</sup> · Kristina Sannenschke<sup>1</sup> · Dieter Fischer<sup>2</sup> · Christiane E. Angermann<sup>3</sup> · Michael Böhm<sup>4</sup> · Philipp Röntgen<sup>1</sup> · Johann Bauersachs<sup>1</sup> · Denise Hilfiker-Kleiner<sup>1</sup>

Em andamento

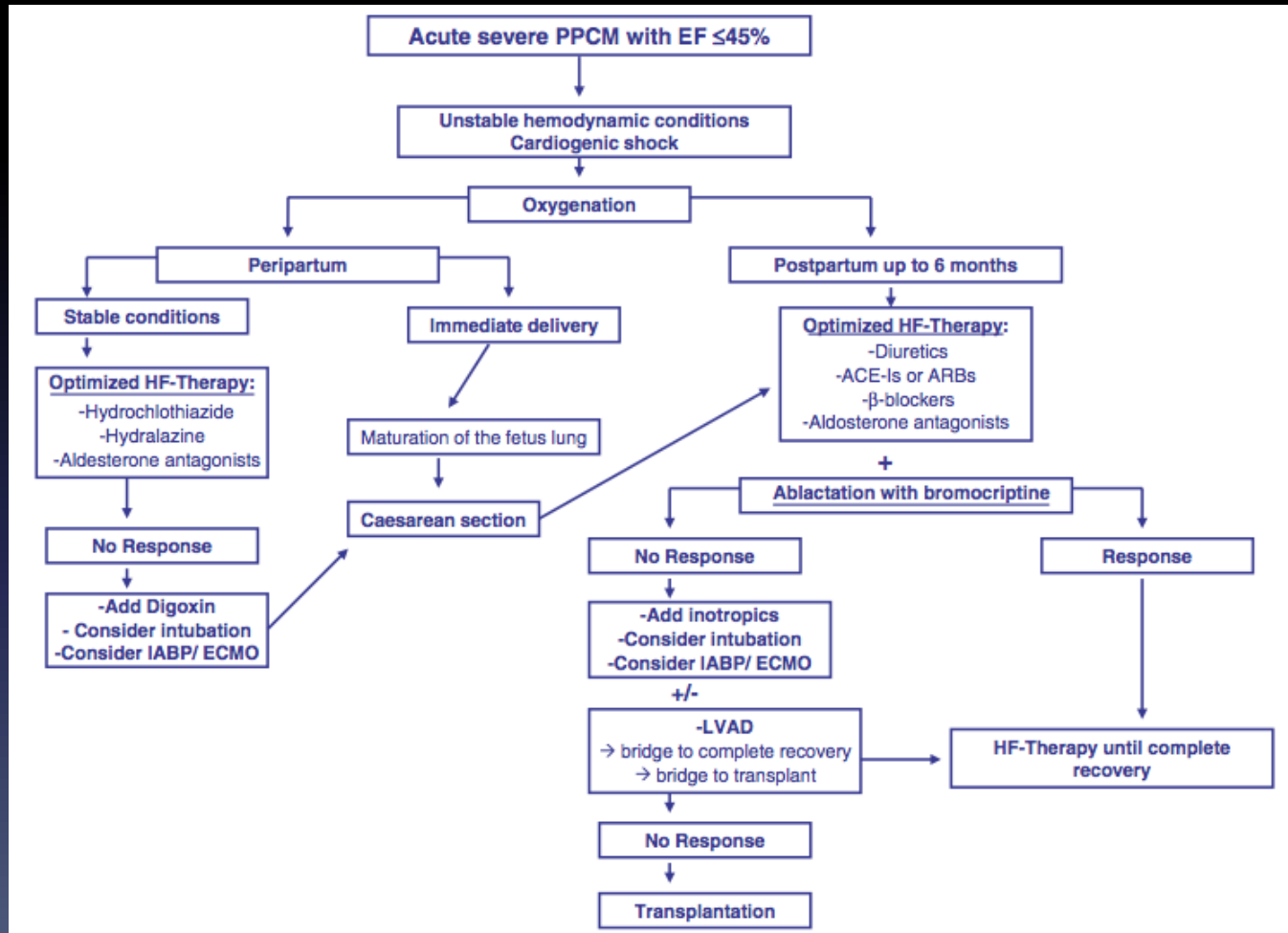
**Effect of levosimendan and predictors of recovery in patients with peripartum cardiomyopathy: a randomized clinical trial**

Murat Bıteker · Nilüfer Eksi Duran · Gökhan K. · Gündüz ·  
Hacı İbrahim Tanboğa · Tayyar ·  
Taylan Akgün · Mustafa Yıldız · Ozkan

Peripartum cardiomyopathy: current management and future perspectives Denise Hilfiker-Kleiner European Heart Journal (2015) 36, 1090–1097



# Algoritmo terapêutico para miocardiopatia periparto aguda severa

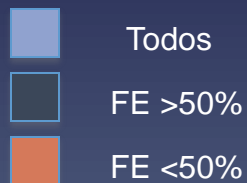
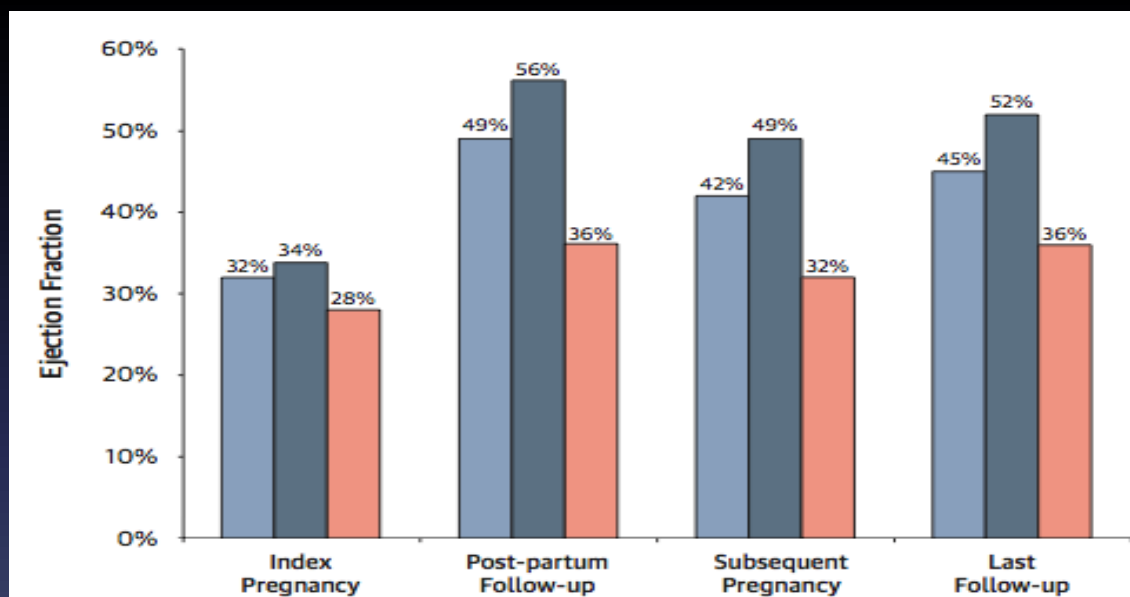


EURObservational Research Programme: a worldwide registry on peripartum cardiomyopathy (PPCM) European Journal of Heart Failure (2014) 16, 583–591



# Miocardiopatia periparto evolução

## Fração de ejeção

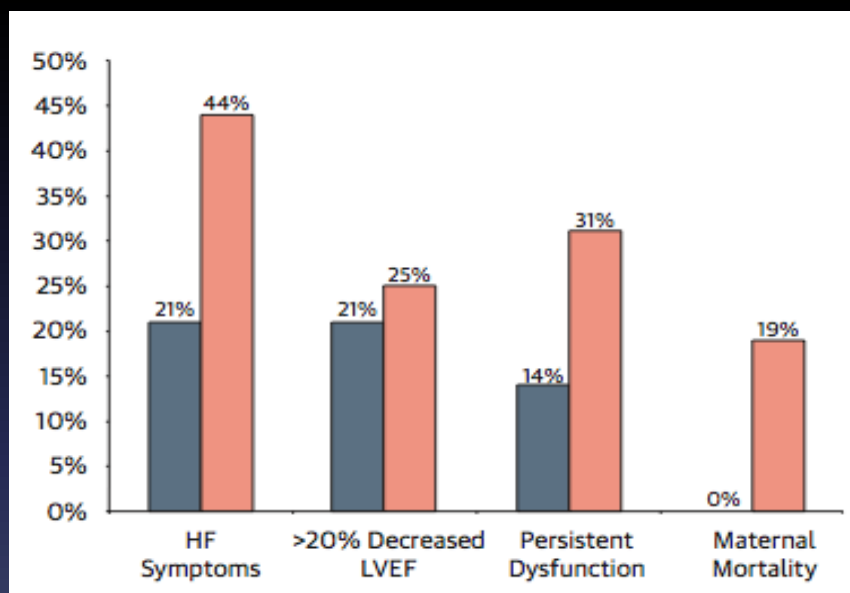


Elkayam Pregnancy Risk After Peripartum Cardiomyopathy (J Am Coll Cardiol 2014;64:1629–36)

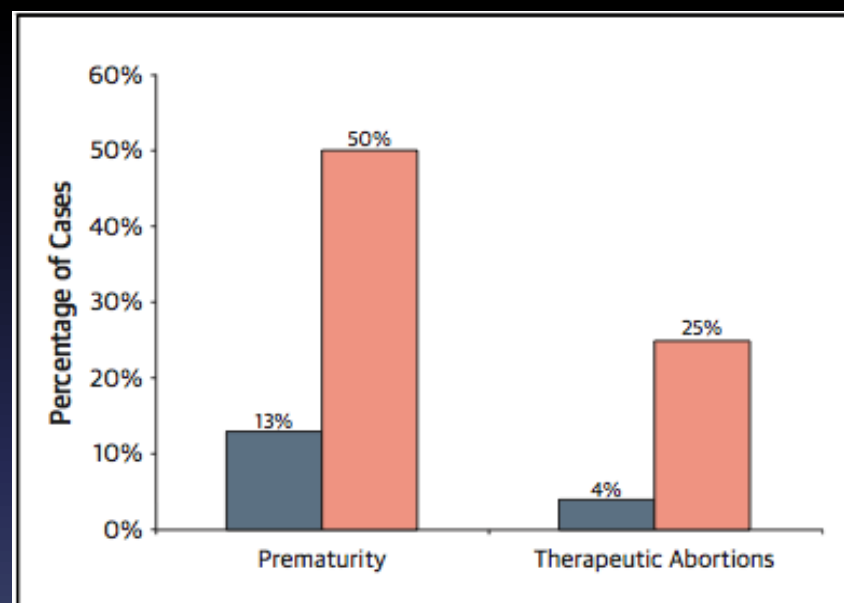


# Miocardiopatia periparto evolução

## Complicações maternas



## Complicações fetais



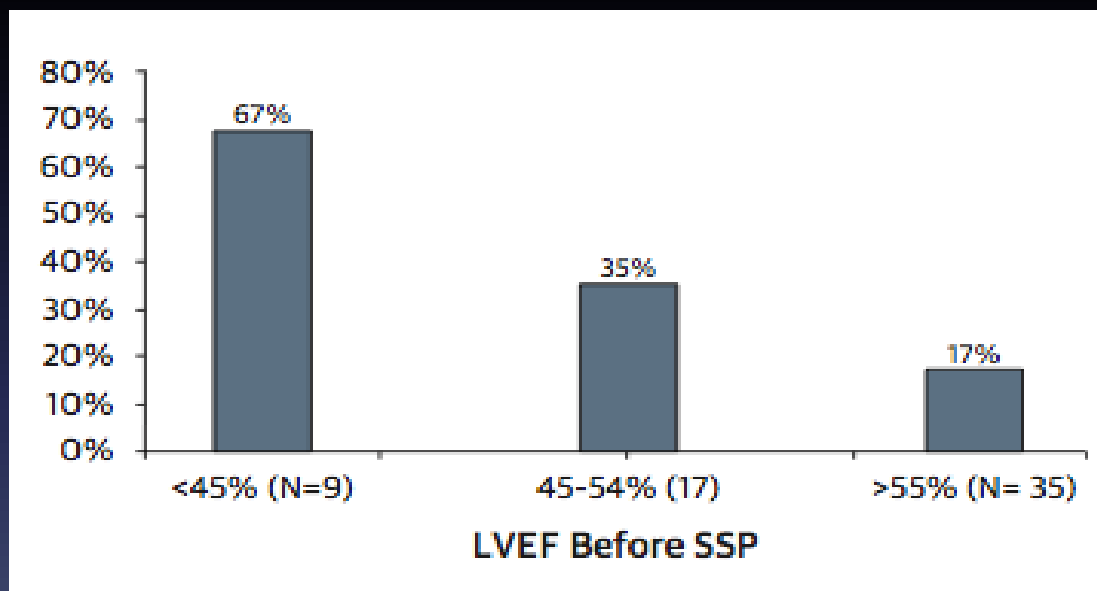
■ FE >50%  
■ FE <50%

Elkayam Pregnancy Risk After Peripartum Cardiomyopathy (J Am Coll Cardiol 2014;64:1629–36)



# Miocardiopatia periparto evolução

## Recorrência



Elkayam Pregnancy Risk After Peripartum Cardiomyopathy (J Am Coll Cardiol 2014;64:1629–36)



# Miocardiopatia Gravidez subsequente

**TABLE 1** Outcome of Subsequent Pregnancy in Women With a History of Peripartum Cardiomyopathy: Results of a Survey

Group	Maternal Outcome			Fetal Outcome		
	No Relapse	Relapse	Death	Live Birth	Abortions	Stillbirth
Group A	74.4%	23.3%	2.3%	93%	4.7%	2.3%
Group B	37.5%	54.2%	8.3%	83.3%	16.7%	0%

Values are %. Modified with permission from Ostrzega and Elkayam (9).

Group A = women with recovered left ventricular function; Group B = women with persistent left ventricular dysfunction.

Elkayam Pregnancy Risk After Peripartum Cardiomyopathy (J Am Coll Cardiol 2014;64:1629–36)



# Miocardiopatia Gravidez subsequente

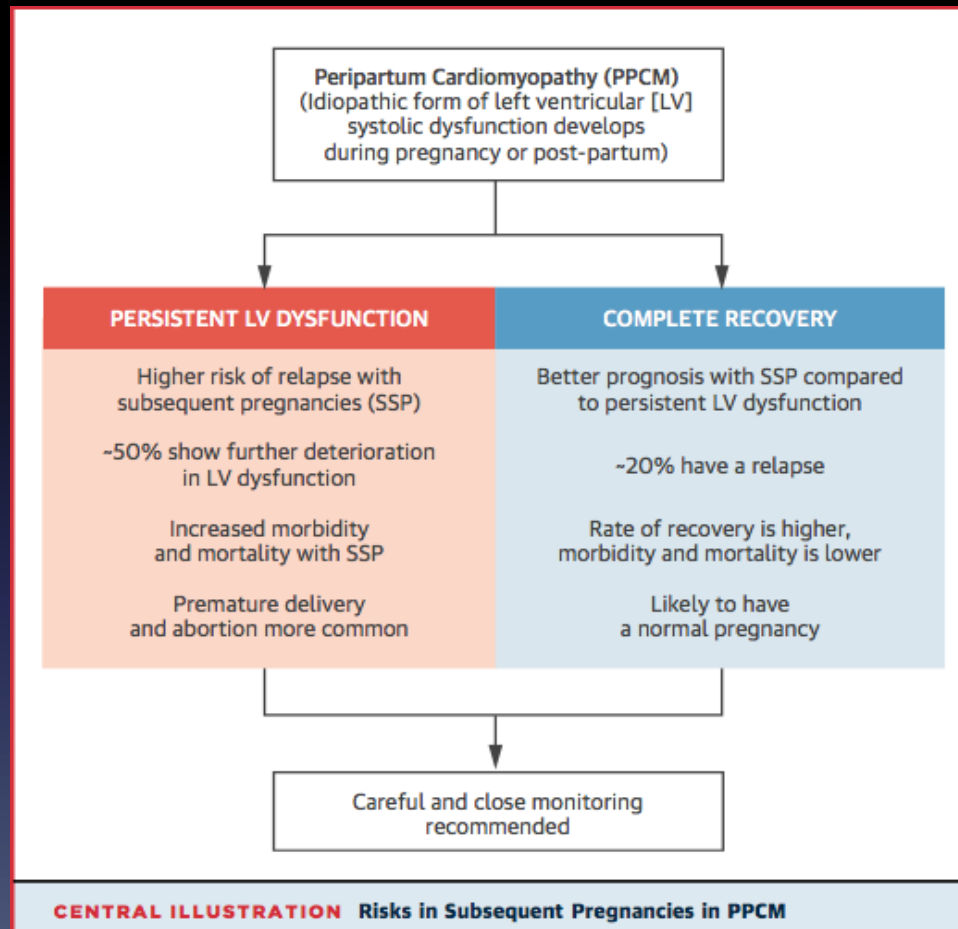
Baseline LVEF	(n)	Percentage with relapse heart failure	Country	Outcome comments
<0.45	10	66.7	Haiti <sup>a</sup>	1 death, 7/10 left with worse cardiomyopathy
0.45–0.49	8	37.5	USA <sup>b</sup>	4/8 returned to LVEF pre-subsequent pregnancy
0.50–0.54	6	33.3	USA <sup>b</sup>	5/6 LVEF $\geq$ 0.50 last echocardiography
$\geq$ 0.55	26	23.1	USA <sup>b</sup>	22/26 LVEF $\geq$ 0.50 last echocardiography
$\geq$ 0.55+CR	12	0 <sup>c</sup>	USA <sup>b</sup>	All LVEF $\geq$ 0.50 last echocardiography

Fett J.D. Why Do Some Recovered Peripartum Cardiomyopathy Mothers Experience Heart Failure With a Subsequent Pregnancy? Curr Treat Options Cardio Med (2015) 17:354





# Miocardiopatia periparto



Elkayam Pregnancy Risk After Peripartum Cardiomyopathy (J Am Coll Cardiol 2014;64:1629–36)



# Miocardio patia periparto e nova gestação

<b>IV</b>	Extremely high risk of maternal mortality or severe morbidity; pregnancy contraindicated. If pregnancy occurs termination should be discussed. If pregnancy continues, care as for class III.
• Pulmonary arterial hypertension of any cause	
• Severe systemic ventricular dysfunction (LVEF <30%, NYHA III–IV)	
• Previous peripartum cardiomyopathy with any residual impairment of left ventricular function	
• Severe mitral stenosis, severe symptomatic aortic stenosis	
• Marfan syndrome with aorta dilated >45 mm	
• Aortic dilatation >50 mm in aortic disease associated with bicuspid aortic valve	
• Native severe coarctation	

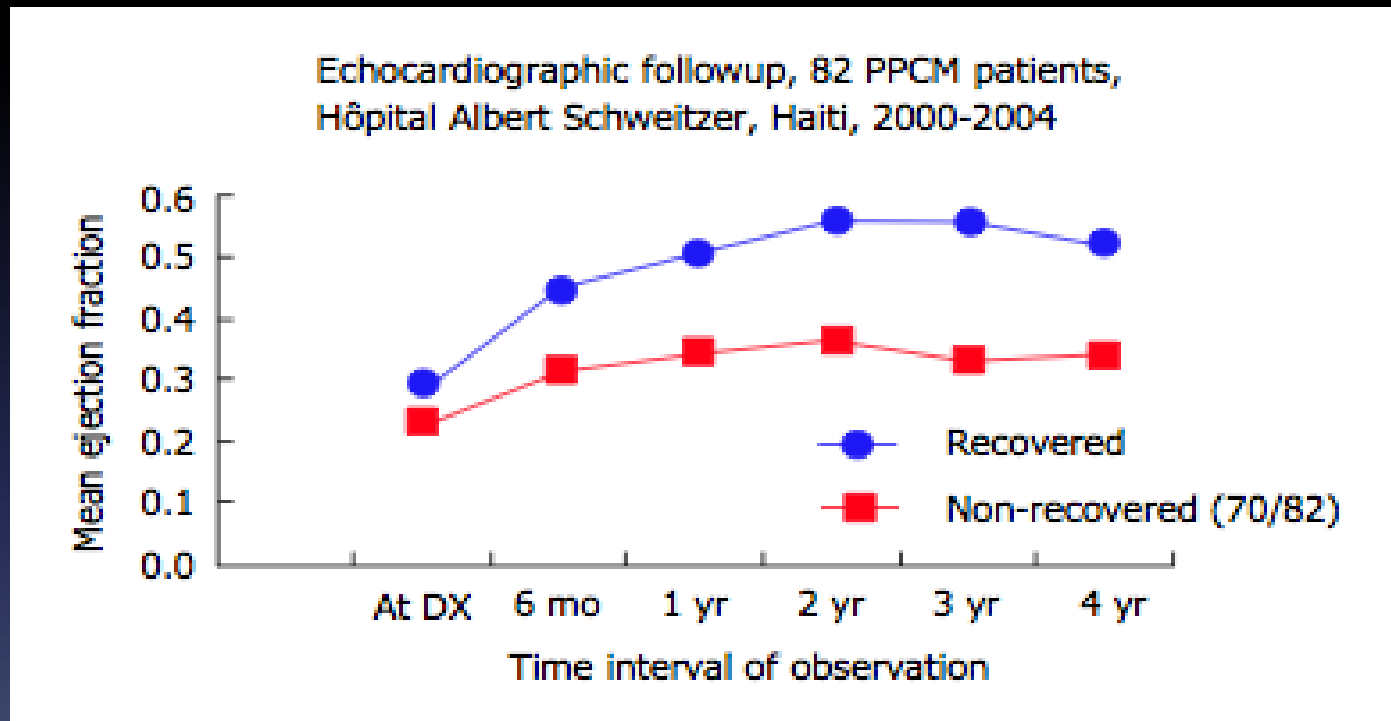
Adapted from Thorne et al.<sup>73</sup>  
LVEF = left ventricular ejection fraction; NYHA = New York Heart Association; WHO = World Health Organization.

ESC Guidelines on the management of cardiovascular diseases during pregnancy European Heart Journal (2011) 32, 3147–3197

Tedoldi CL, Freire CMV, Bub TF et al Arq Bras Cardiol.2009;93(6 supl.1):e110-e178



# Prognóstico



Fett JD. Peripartum cardiomyopathy: A puzzle closer to solution. *World J Cardiol* 2014; 6(3): 87-99



# Prognóstico

- Dependente da recuperação da FE.
- Recuperação varia entre 23 e 54%.
- Dependente da raça.

EURObservational Research Programme: a worldwide registry on peripartum cardiomyopathy (PPCM) European Journal of Heart Failure (2014) 16, 583–591



## Prognóstico

Pacientes que persistem com FE baixa 50% terão recorrência da miocardiopatia

Em pacientes com recuperação da FE 20% terão recidiva da miocardiopatia

Elkayam Pregnancy Risk After Peripartum Cardiomyopathy (J Am Coll Cardiol 2014;64:1629–36)



## Mensagens para guardar

Sintomas semelhantes a gestações normais

Retardo de crescimento fetal pode auxiliar no diagnóstico

Manutenção do tratamento é essencial para o prognóstico

Desaconselhamento em relação a novas gestações

Deteção precoce é essencial para o prognóstico

Peripartum cardiomyopathy: current management and future perspectives Denise Hilfiker-Kleiner European Heart Journal (2015) 36, 1090–1097



# Muito Obrigado

