JCOR 2017

Cardiomyopathie du Péripartum

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Réanimation Polyvalente - Monastir

Hammamet, 20 Mai 2017

2016-2017

Défit: Distinction

Modification physiologiques

≠

Pathologie cardio-vasculaire

Confusion → retard et/ou mauvais diagnostic

Histoire en quelques dates

1971

Natural Course of Peripartum Cardiomyopathy

Illinois

By JOHN G. DEMAKIS, M.D., SHAHBUDIN H. RAHIMTOOLA, M.B., M.R.C.P.E.,
GEORGE C. SUTTON, M.D., W. ROBERT MEADOWS, M.D.,
PAUL B. SZANTO, M.D., JOHN R. TOBIN, M.D.,
AND ROLF M. GUNNAR, M.S., M.D.

Figure 4

Histological section of myocardial biopsy specimen taken from the outflow tract of the right ventricle during exploratory thoracotomy from a patient in group B. There is extensive myocardial fibrosis. Mallory stain

- 27 patientes

Terme: **CMPP**

Critères de définition:(disponibilités de l'époque!)

- 1) IC (dernier mois de grossesse ou dans les 5 mois suivant l'accouchement)
- 2) l'absence d'une étiologie d'IC
- 3) Absence d'une path cardiaque connue avant le dernier mois de grossesse.

Circulation. 1971;44:1053-1061

2016

Définition... plus simple

CMPP: LES 4 Critères diagnostiques

- (1) Development of HF in the last month of pregnancy or within 5 months after delivery
- (2) LV systolic dysfunction (LV EF < 45% by echocardiography)
- (3) No identifiable cause for HF
- (4) No recognized heart disease before the last month of pregnancy

Mais pas toujours associée a une dilatation VG

Epidémiologie

Temporal Trends in Incidence and Outcomes of Peripartum Cardiomyopathy in the United States: A Nationwide Population-Based Study

Dhaval Kolte, MD, PhD;* Sahil Khera, MD;* Wilbert S. Aronow, MD; Chandrasekar Palaniswamy, MD; Marjan Mujib, MD, MPH; Chul Ahn, PhD; Diwakar Jain, MD; Alan Gass, MD; Ali Ahmed, MD, MPH; Julio A. Panza, MD; Gregg C. Fonarow, MD

Conclusion—From 2004 to 2011, the incidence of PPCM has increased in the United States. Maternal MAE rates overall have remained unchanged while cardiogenic shock, utilization of mechanical circulatory support, and in-hospital mortality have increased during the study period. Further study of the mechanisms underlying these adverse trends in the incidence and outcomes of PPCM are warranted. (*J Am Heart Assoc.* 2014;3:e001056 doi: 10.1161/JAHA.114.001056)

Critical Care Volume 13 Suppl 1, 2009

29th International Symposium on Intensive Care and Emergency Medicine

Brussels, Belgium, 24–27 March 2009

P163

**Cardiac diseases during pregnancy and peripartum:
13 years in the ICU**

K Baccar, N Baffoun, C Kaddour

National Institute Neurology, Tunis, Tunisia

Critical Care 2009, 13(Suppl 1):P163 (doi: 10.1186/cc7327)

Introduction Pregnancy, labour and the postpartum period constitute major stresses on the cardiovascular system. Patients with heart disease may decompensate due to the physiologic changes that occur during pregnancy and may develop a cardiac event with obstetric events.

Methods A retrospective collection of data for all obstetric patients admitted to our ICU in a 13-year period (September 1995 to September 2008). Data collected include the antepartum or peripartum heart status (abnormal finding on physical examination, ECG and echocardiography), reason for admission, stay on the ICU prognostic scoring (APACHE II, Obstetrical SAPS) and the outcome.

Results During 13 years there were 42 obstetric admissions to our ICU. The reason for admission was due to pulmonary edema 16 (38.1%) cases, hemorrhagic shock 11 (26.2%) cases, cardiogenic shock 8 (19%) cases, pulmonary emboli 4 (9.5%) cases, stroke 2 (4.8%) cases and recent myocardial infarction 1 (2.4%) case.

Admission was unplanned for 36 (85.7%) parturients following emergency caesarean section. Status of the heart was severe aortic or mitral stenosis 21 (50%) cases, mitral or aortic regurgitation with heart failure 5 (11.9%) cases, mechanical prosthetic valve requiring anticoagulation 8 (19%) cases, myocardial infarction 1 (2.4%) case and peripartum cardiomyopathy 7 (16.6%) cases. The median duration of stay was 4 days (range 1 to 11 days), median Obstetrical SAPS was 19.3 (range 10 to 32) and APACHE II score was 13 (range 8 to 32). There were five (11.9%) maternal deaths, due to two cases PPCM, one cerebral death and two prothesis thrombus.

Conclusions The majority of the cases of parturients was unbooked for antenatal care; this leads us to acknowledge the existence of risk factors related to pregnancy. Maternal functional class is an important predictor of outcome; a high index of suspicion for cardiac diseases is essential to identify risk [1], once these patients are referred for a cardiologic opinion there is a needed for cardiologist to develop a systematic approach to their evaluation. Ideally these considerations should be commenced during prepregnancy consultations, but continued throughout pregnancy and the postpartum period [1].

7 CMPP/42 :16%
2deces/5 /CMPP

STATE-OF-THE-ART PAPER

**Clinical Characteristics of
Peripartum Cardiomyopathy in the United States**

Diagnosis, Prognosis, and Management

Uri Elkayam, MD

**Surtout postpartum immédiat
60 %**

Figure 1: Time of onset of symptoms of peripartum cardiomyopathy according to country.

Facteurs de risque

Esther Beus
Walther N. K. A. Mook
Graham Ramsay
Jan L. M. Stappers
Hans W. H. M. Putten

Peripartum cardiomyopathy: a condition intensivists should be aware of

Reported associated condition

Age

Multiparité

Africaine

Tocolyse

Esther Beus
Walther N. K. A. Mook
Graham Ramsay
Jan L. M. Stappers
Hans W. H. M. Putten

Peripartum cardiomyopathy: a condition intensivists should be aware of

Article

Serum Selenium and Ceruloplasmin in Nigerians with Peripartum Cardiomyopathy

Kamilu M. Karaye^{1,2,*}, **Isah A. Yahaya**³, **Krister Lindmark**^{2,4} and **Michael Y. Henein**^{2,4}

Int. J. Mol. Sci. **2015**, *16*, 7644–7654

Sélénium

30% sans aucun facteur de risque...

Phenotyping and outcome on contemporary management in a German cohort of patients with peripartum cardiomyopathy

**A. Haghikia · E. Podewski · E. Libhaber · S. Labidi · D. Fischer ·
P. Roentgen · D. Tsikas · J. Jordan · R. Lichtinghagen · C. S. von Kaisenberg ·
I. Struman · N. Bovy · K. Sliwa · J. Bauersachs · Denise Hilfiker-Kleiner**

Cas familiaux

Review

Pregnancy and the cardiovascular system

Amr E. Abbas^{a,*}, Steven J. Lester^a, Heidi Connolly^b

Table 1

Cardiovascular adaptations during pregnancy

Parameter	Average change during pregnancy
Blood volume	↑ 35%
Cardiac output	↑ 40–43%
Stroke volume	↑ 30%
Heart rate	↑ 15–17%
Systemic vascular resistance	↓ 15–21%
Mean arterial pressure	No significant change
Systolic blood pressure	↓ 3–5 mm Hg
Diastolic blood pressure	↓ 5–10 mm Hg
Central venous pressure	No significant change
Serum colloid osmotic pressure	↓ 14%
Hemoglobin	↓ 2.1 g/dl

Donc au cours de la grossesse

Reprogrammation

- Hormonale
 - Métabolique
 - Hémodynamique

La question

THE LANCET

Seminar
2006

Peripartum cardiomyopathy

Karen Sliwa, James Fett, Uri Elkavam

Physiopathologie

- **Hypothèse inflammatoire :**

- Biopsies myocardique : lésion « myocardite aigue »
- Perturbation réponse inflammatoire au cours de la grossesse
- Facteur infectieux initiant/aggravant cette réaction

Inflammation inappropriée : virus+++

Physiopathologie

- Hypothèse inflammatoire
- Hypothèse Auto-immune
- **Hypothèse Hormonale**
 - Voie métabolique de la prolactine**

Peripartum Cardiomyopathy

Zolt Arany, MD, PhD; Uri Elkayam, MD

Diagnostic

Présentations cliniques

• Symptomatologie cardiaque

- oedème aigu du poumon
- troubles du rythme
- insuffisance C globale
- embolie artérielle

- **Oxygéno-dependance**
- **Toux persistante**

• Symptômes « classiques » de la grossesse

- asthénie
- douleurs abdominales
- dyspnée (anémie)
- oedèmes

- **Arrêt cardio-respiratoire**

Phenotyping and outcome on contemporary management in a German cohort of patients with peripartum cardiomyopathy

A. Haghikia · E. Podewski · E. Libhaber · S. Labidi · D. Fischer ·
P. Roentgen · D. Tsikas · J. Jordan · R. Lichtinghagen · C. S. von Kaisenberg ·
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A % of Patients with Pathologic Values of Serum Markers

B Log NT-proBNP (pmol/l)

C ADMA (mmol/l)

D Cathepsin D (U/l)

Echo graphie

Hypokinésie - Dilatation

IM

IT

↑ PRG

T. compliance

Thrombus VG

Echocardiographie 2D strain

- L'imagerie de *strain* et de *strain rate* est un outil de quantification précise de la fonction et de la contractilité myocardiques.
- Elle analyse le déplacement myocardique par le *tracking* des *speckles* (marqueurs acoustiques naturels) au sein de l'image ultrasonique bidimensionnelle

Echocardiographie 2D-*strain*

Mouvements du cœur
dans les 3 plans
-hauteur
-largeur
-rotation

Echocardiographie 2D strain

Longitudinal systolic strain profiles and outcomes in peripartum cardiomyopathy

Alexandros Briasoulis M.D. | Marian Mocanu M.D. | Karolina Marinescu M.D. |
Osama Qaqi M.D. | Mohan Palla M.D. | Tesfaye Telila M.D. | Luis Afonso M.D.

**Le strain est modifié dans la MCPP même si la FE est normale
(détection des formes frustrées ou précoces ?)**

IRM

Thrombus

Suivi

Biopsie endomyocardique

International Journal of Cardiology 104 (2005) 92 – 100

International Journal of
Cardiology

“Endomyocardial biopsy may be considered in view of a strong clinical suspicion of myocarditis or no improvement **after 2 weeks** of optimal medical **therapy** for heart failure, especially if malignant ventricular **arrhythmias** are present and the diagnosis of giant cell myocarditis is being considered. Cardiac **MRI** can be a useful technique to **guide site of biopsy** to improve the diagnostic yield”

International Journal of Cardiology 112 (2006) 125 – 126

International Journal of
Cardiology

Selon le moment pendant lequel on réalise la biopsie dans l’histoire de la maladie ou suivant l’endroit où il est effectué, les résultats histologiques sont très variables!!

Diagnostics différentiels

oui

non

*peut-
être*

Diagnostics différentiels

Table 1 Peripartal acute dyspnoea: differential diagnosis of acute peripartum cardiomyopathy

	PPCM	Pre-existing CMP, valve disease or congenital heart disease	Pregnancy-associated myocardial infarction	Pulmonary embolism/ amniotic liquid embolism	Myocarditis
History	Most commonly post-partal onset of dyspnoea	Earlier onset (during second trimester) Sometimes family history	Retrosternal chest pain, abdominal discomfort, nausea	Pleuritic chest pain	Infection
Biomarkers	Elevated natriuretic peptides	Elevated natriuretic peptides	Elevated troponin	Elevated D-dimer, troponin, natriuretic peptides	Elevated troponin Possibly elevated natriuretic peptides
Echocardiography	Left and/or right ventricular dysfunction	Evidence of pre-existing valve disease or congenital defect	Regional hypokinesis/akinesis	RV dysfunction, elevated RV pressure, McConnell's sign	Regional or general hypokinesis
Additional tests	Consider MRI	Consider MRI Consider genetic test	Coronary angiography	CT-scan or V/Q scintigraphy; consider angiography	MRI Consider myocardial biopsy

CMP, cardiomyopathy; MRI, magnetic resonance imaging; PPCM, peripartum cardiomyopathy; RV, right ventricular.

Practical management of peripartum cardiomyopathy

Mi-Jeong Kim¹ and Mi-Seung Shin²

Flow chart for suspected peripartum cardiomyopathy

Prise en charge 2017

Pas de traitement spécifique

Prise en charge

**Défaillance
cardiaque**

Traitements de l'insuffisance cardiaque durant la grossesse

Traitements autorisés

Traitements contre-indiqués

Digoxine

Dérivés nitrés

Héparine

Diurétiques

Bêtabloquants

Dobutamine

Dopamine

IEC/ARA II

Amiodarone

Antivitamines K

Severe PPCM with cardiopulmonary distress

Consider mechanical circulatory support if refractory cardiopulmonary distress

Recovery?

Transplantation

Weaning

PPCM without cardiopulmonary distress

Contemporary Reviews in Cardiovascular Medicine

Peripartum Cardiomyopathy

Zolt Arany, MD, PhD; Uri Elkayam, MD

Evaluation of Bromocriptine in the Treatment of Acute Severe Peripartum Cardiomyopathy: A Proof-of-Concept Pilot Study

Karen Sliwa, Lori Blauwet, Kemi Tibazarwa, Elena Libhaber, Jan-Peter Smedema, Anthony Becker, John McMurray, Hatice Yamac, Saida Labidi, Ingrid Struman and Denise Hilfiker-Kleiner

- 20 patientes avec CMPP récente et FEVG < 35%
- Randomisées bromocriptine (5 mg/24h 2 semaines puis 2,5 mg/24h 6 semaines) vs pas de ttt
- End-point: Décès / NYHA III-IV / FEVG < 35% à 6 mois

Limites Bromocriptine

- Pas d'allaitement → dénutrition nourrisson ?

Figure 5. Growth and survival of children of PPCM study mothers from birth to 3 months plotted on World Health Organization growth charts.

Sliwa, Circulation, 2010

- Risque nécrose myocardique ?

James, JACC, 2010

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

Murat Biteker · Nilüfer Ekşi Duran · Hasan Kaya · Sabahattin Gündüz · Halil İbrahim Tanboğa · Tayyar Gökdeniz · Gökhan Kahveci · Taylan Akgün · Mustafa Yıldız · Mehmet Özkan

Conclusions Addition of levosimendan to conventional therapy did not improve outcome in patients with PPCM. In patients with PLVD or patients who died, LVEF, LVESD and left atrial diameter were worse than those with complete resolution.

PPCM Participant enrolment in ESC & NON-ESC Countries as of 26 July 2016.

148 registered centres in 61 countries of which, 43 are active countries.

ENROLLMENT ≥ 1	
	ESC countries
	NON-ESC Countries
•	1 – 5 participants
●	6 – 15 participants
●	> 15 participants

Clinical characteristics of patients from the worldwide registry on peripartum cardiomyopathy (PPCM)

EURObservational Research Programme in conjunction with the Heart Failure Association of the European Society of Cardiology Study Group on PPCM

« Thérapeutiques d'exception »

RESEARCH

Open Access

Acute and critically ill peripartum cardiomyopathy and 'bridge to' therapeutic options: a single center experience with intra-aortic balloon pump, extra corporeal membrane oxygenation and continuous-flow left ventricular assist devices

Sofie Gevaert^{1*}, Yves Van Belleghem², Stefaan Bouchez³, Ingrid Herck⁴, Filip De Somer², Yasmina De Block^{1,2}, Fiona Tromp¹, Els Vandecasteele¹, Floor Martens², Michel De Pauw¹

Table 1 Patient characteristics

Patient	1	2	3	4	5	6
Year	2001	2008	2010	2007	2008	2009
Age (y)	34	35	36	37	38	28
BSA (m ²)	1.90	1.88	1.58	1.83	1.60	1.79
Race	C	A	C	C	C	C
Obstetrical Hx	G4A0P4	G2A0P2	G4A0P4	G1A0P0	G2A0P1	G3A1P1
Symptom onset	3wPP	5mPP	5mPP	38wPr	35wPr	38wPr
Diagnosis	3w PP	10mPP	18mPP	38wPr	36wPr	38wPr
Clinical picture	APE	ADHF	ADHF	ADHF	ADHF	CS
LVEDD (mm)	55	62	79	53	61	68
Serology	Negative	Mycoplasma IgM	Negative	Negative	Negative	Mycoplasma IgM
Coronary angio	Normal	-	Normal	-	Normal	-
Biopsy	-	Negative	Negative	-	Negative	Negative
IABP (d)	7	13	5	4	6	1
ECMO (d)	-	-	-	-	-	7
LVAD (d)	-	126	Since 26/04/2010	-	360	78
Complications	-	Perop. rupture aorta Tamponade 2x Pocket Infection	Rectus hematoma Occlusion AFC	-	-	Bleeding anast. aorta
Outcome	Recovery	Tx, SD 535 days postTx	Alive, on Tx list	Recovery	Tx	Tx

A, African; APE, acute pulmonary edema; ADHF, acute decompensated heart failure; AFC, Arteria Femoralis Communis; anast, anastomosis; BSA, body surface area; C, Caucasian; CS, cardiogenic shock; d, days; Hx, history; LVEDD, left ventricular end diastolic diameter at presentation; Perop, peroperative; PP, postpartum; Pr, pregnancy; SD, sudden death; Tx, cardiac transplantation; Y, year.

Peripartum Cardiomyopathy

Definition, Incidence, Etiopathogenesis, Diagnosis, and Management

Jalaj Garg, MD, Chandrasekar Palaniswamy, MD, and Gregg M. Lanier, MD

- Pentoxifylline (antiTNF-alpha) (30 patientes, Sliwa K; Eur J Heart Fail. 2002)
- Immunoglobulines (6 patientes-Bozkurt B; J Am Coll Cardiol. 1999)
- Défibrillateur AI (EF < 30% malgré traitement ou arythmie ventriculaire ou QRS > 120)
- Anticoagulation (FEVG<25%)
- Mechanical Assist Device –ECMO -BCPIA
- Heart Transplantation

Pronostic

Mortalité

- Mortalité $\approx$ **10-16%** à 6 mois

Sliwa et al. Eur J Heart Fail 2010;12:767-78

- Meilleur pronostic que les autres cardiomyopathies

Felker et al. N Engl J Med 2000;342:1077-84

STATE-OF-THE-ART PAPER

Clinical Characteristics of Peripartum Cardiomyopathy in the United States

Diagnosis, Prognosis, and Management

Uri Elkayam, MD

Clinical Profile and Predictors of Complications in Peripartum Cardiomyopathy

SOREL GOLAND, MD,¹ KALGI MODI, MD,² FAHED BITAR, MD,³ MUNIR JANMOHAMED, MD,³ JAMES M. MIROCHA, MS,¹ LAWRENCE SC. CZER, MD,¹ SANDRA ILLUM, MD,² PARTA HATAMIZADEH, MD,³ AND URI ELKAYAM, MD³

MCP

182 patientes

**50-60%
Guérison
totale
(3-6 mois)**

**20-30%
Guérison
partielle**

Evènements indésirables majeurs :

Choc irréversible, greffe, décès, ACR, assistance, ECMO, Tr du rythme (pacemaker, defib), Insuff cardiaque chronique

	No MAE n=136	MAE n=46	p-value
Age (years)	30±6	27±8	0.03
Age > 30 years	53%	42%	0.3
Non-Caucasian	37%	61%	0.005
<u>Parity</u>	2.0±1.6	2.0±2.1	0.3
Multipara	53%	41%	0.3
Twin or triplet pregnancy			0.02
Gestation hypertensive			0.2
Tocolytic therapy			1.0
Index pregnancy			0.3
1 st			
2 nd			
3 rd and more			
Caesarian delivery	21%	15%	0.7
Occurrence of symptoms antepartum	30%	41%	0.2
Occurrence of symptoms postpartum	64%	54%	0.3
Diagnosis delay (weeks)	1.7±3.0	3.8±6.1	0.02
<u>Diagnosis delay (≥ 1 week)</u>	48%	60%	0.3
LVEF (%) baseline	31±11	24±10	<0.001
<u>LVEF < 25%</u>	31%	63%	0.001
LVEF (%) at ≥ 6month	47±13	32±14	<0.0001
LV Recovery (LVEF ≥ 50%)	45%	18%	<0.001

**facteurs de mauvais pronostic:
Echo+++**

Evaluation of the Clinical Relevance of Baseline Left Ventricular Ejection Fraction as a Predictor of Recovery or Persistence of Severe Dysfunction in Women in the United States With Peripartum Cardiomyopathy

SOREL GOLAND, MD,¹ FAHED BITAR, MD,¹ KALGI MODI, MD,² JORDAN SAFIRSTEIN, MD,³ ANGELA RO, MD,³ JAMES MIROCHA, MS,¹ NUDRAT KHATRI, MD,¹ AND URI ELKAYAM, MD¹

Los Angeles, California; Shreveport, Louisiana; and New York, New York

CMPP: Facteurs de mauvais pronostic

- Gravité de la défaillance HD **initiale** (FEVG) et sa réponse au ttt
- Complications **emboliques**
- Existence d'une cardiopathie **séquellaire** au delà **du 6 ème** mois après diagnostic

Une nouvelle grossesse

Outcome of Subsequent Pregnancy in Patients With Documented Peripartum Cardiomyopathy

Karen Sliwa, MD, PhD, Olaf Forster, MD, Fitzgerald Zhanje, MD, Geoff Candy, PhD, John Kachope, MD, and Rafique Essop, MD

FIGURE 1. Time course pattern of left ventricular function in patients with PC and subsequent pregnancy. † $p \leq 0.05$ comparing 8 months of pregnancy versus 3 months postpartum.

FIGURE 2. Time course pattern of TNF- α in patients with PC and subsequent pregnancy. † $p \leq 0.05$ comparing 8 months of pregnancy versus 3 months postpartum.

Risk of Subsequent Pregnancy in Women With a History of Peripartum Cardiomyopathy

Uri Elkayam, MD

Recommandations avant nouvelle grossesse

- CID si FE < 50%
- Déconseillée même si FE normale (20% récursive)
- Arrêt des IEC 3 mois avant la grossesse (test !)
- Echographie de référence sans IEC *(car sont CID si grossesse)*
- Epreuve d'effort conseillée
- Surveillance par BNP + écho
- Information (ITG)
- Traitement bromocryptine après naissance ?

Take home messages

Ce qu'on sait

- Incidence variable
- Caractéristiques semblables
- HTA- Femme africaine
- Les complications les plus graves peuvent être diminuées ou évitées
- La récupération complète se produit plus fréquemment qu'avec toute autre cardiomyopathie
- Auto-immunité-inflammation
- Des guidelines efficaces fondées sur des preuves disponibles
- Nouvelle grossesse : à risque

Des questions

- Les «déclencheurs» réels qui initient le processus
- Rôle du virus dans la pathogenèse
- Incidence plus élevée/ plus sévère chez les africaines : prédisposition génétique
- Prolactine / inhibition de la prolactine
- Pourquoi certaines patientes ayant récupérés ont-ils une rechute d'insuffisance cardiaque lors des grossesses ultérieures?
- Rôle des micronutriments et des métaux traces dans la pathogenèse
- Biomarqueurs