

# L' ENFANT MACROSOME PRISE EN CHARGE PÉDIATRIQUE

Pratiques reposant sur les faits scientifiques prouvés



44<sup>e</sup> journées de la SFMP  
Lyon 2014



24/10/2014

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AUDIPOG

" Estimation du poids fœtal ou néonatal et pratiques "

# Incidence of and risk factors for nodding off at scientific sessions



Kenneth Rockwood, David B. Hogan, Christopher J. Patterson; for the Nodding at Presentations (NAP) Investigators CMAJ • DEC. 7, 2004; 171 (12)

**Table 1: Risk factors for nodding off at lectures**

Factor	Odds ratio (and 95% CI)
<b>Environmental</b>	
Dim lighting	1.6 (0.8–2.5)
Warm room temperature	1.4 (0.9–1.6)
Comfortable seating	1.0 (0.7–1.3)
<b>Audiovisual</b>	
Poor slides	1.8 (1.3–2.0)
Failure to speak into microphone	1.7 (1.3–2.1)
<b>Circadian</b>	
Early morning	1.3 (0.9–1.8)
Post prandial	1.7 (0.9–2.3)
<b>Speaker-related</b>	
Monotonous tone	6.8 (5.4–8.0)
Tweed jacket	2.1 (1.7–3.0)
Losing place in lecture	2.0 (1.5–2.6)

Note: CI = confidence interval.

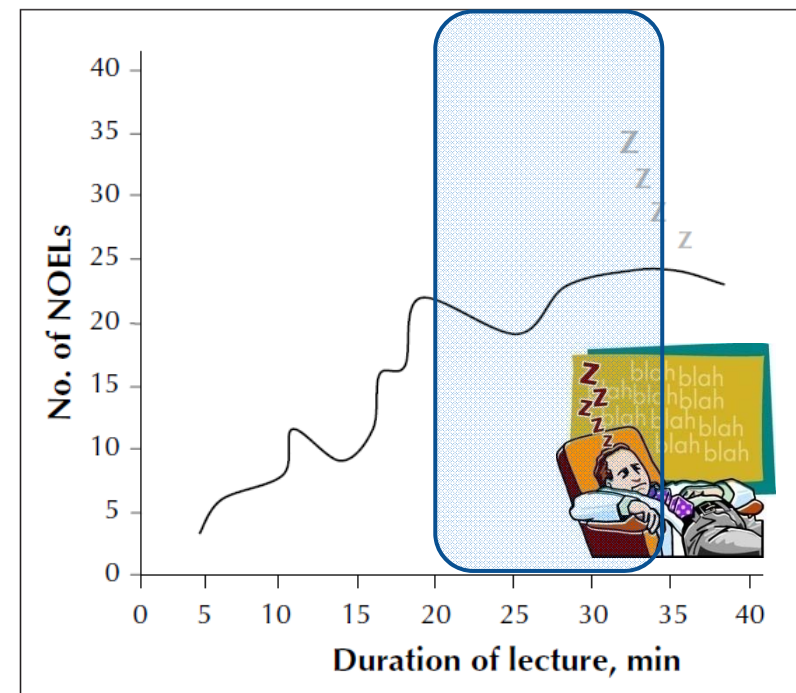


Fig. 1: Special incidence density curve, showing number of nodding-off events per lecture (NOELs) per 100 attendees over length of time of presentation.

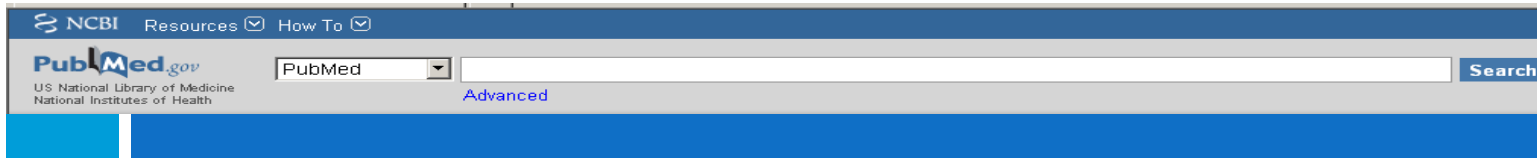
# Définition



- La macrosomie est habituellement définie par un poids de naissance supérieur à 4 000 g
- La macrosomie peut mais doit également se définir par rapport à la répartition des poids de naissance :
  - 90<sup>e</sup> , 95<sup>e</sup> , 97<sup>e</sup> percentiles de courbes de référence selon l'âge gestationnel
    - Leroy Lefort, Fessard ....
    - Intégrée selon le sexe, l'ethnie, parité, mensuration maternelle....
    - Audipog
- Fréquence
  - Courbe de croissance
  - 10% si définition > 90<sup>e</sup> percentile
  - 2,5 % si poids > 4,5 kg

macrosomie : poids $\geq$ 90 <sup>ème</sup> percentile (courbes Audipog)							
Terme (SA)	35	36	37	38	39	40	41
Poids (g)	♀ $\geq$ 3000	♀ $\geq$ 3220	♀ $\geq$ 3400	♀ $\geq$ 3600	♀ $\geq$ 3780	♀ $\geq$ 3850	♀ $\geq$ 3980
	♂ $\geq$ 3000	♂ $\geq$ 3250	♂ $\geq$ 3500	♂ $\geq$ 3720	♂ $\geq$ 3920	♂ $\geq$ 4080	♂ $\geq$ 4180

# Définition



- MEDLINE
  - fetal macrosomia
  - large for gestational age
  - Poids > 4, 4,5, 5 kg
  - Poids > 90<sup>e</sup> percentile
  - Body index en g/cm<sup>3\*</sup>
  - z-score
- *risk, prevention, prediction, management, labour induction, pregnancy outcome, pregnancy complication, delivery and caesarean.*
- Prévalence
  - Quelles courbes de croissance
  - Quelle définition

# Macrosome

## La vision obstétricale?



RESEARCH

2008

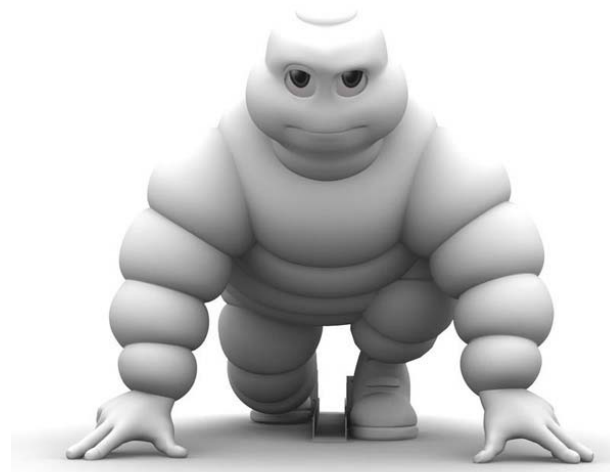
[www.AJOG.org](http://www.AJOG.org)

OBSTETRICS

### How big is too big? The perinatal consequences of fetal macrosomia

Xun Zhang, PhD, Adriana Decker, MD, Robert W. Platt, PhD, Michael S. Kramer, MD

- 4 kg
- 4,5 kg
- 5 kg





# Searching for the Definition of Macrosomia through an Outcome-Based Approach

2014

*Conclusions:* A birthweight greater than 4500 g in Whites, or 4300 g in Blacks and Hispanics regardless of gestational age is the optimal threshold to define macrosomia. A birthweight greater than the 97<sup>th</sup> percentile for a given gestational age, irrespective of race is also reasonable to define macrosomia. The former may be more clinically useful and simpler to apply.

## □ cut-offs

■ 4500 g (ACOG)

■ > 97<sup>e</sup> percentile

■ Comparaison : MFIU, Apgar < 4, Mortalité néonatale, césarienne



**Optimal birth weight percentile cut-offs in defining small- or large-for-gestational-age**

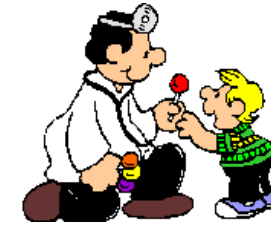
*Acta Pædiatrica/Acta Pædiatrica* 2010 **99**, pp. 550–555

Conclusion: The commonly used 10th and 90th birth weight percentile cut-offs for defining SGA and LGA respectively seem largely arbitrary. The 15th and 97th percentiles may be the optimal cut-offs to define SGA and LGA respectively.

# Macrosomie

Versant pédiatrique :

Période néonatale /suivi



- **Quelle est la cause?**
  - ▣ Peu d'influence sur la prise en charge obstétricale
  - ▣ Diabète maternel +++ : complication de l'hyperinsulinisme ?
- **Quelles complications?**
- **Qui surveiller ?**
  - ▣ Quelle surveillance?
  - ▣ Qui transférer?
- **Évolution à long terme**



# Macrosomie

## Cause

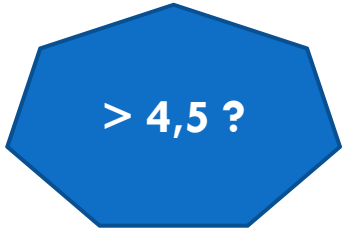


- Diabète++
  - Pré-existant, gestationnel
- Constitutionnelle
  - Courbe intégrée : parité, âge, taille, poids.....
- Syndromique exceptionnel
  - Génétique
    - **Syndrome de Wiedemann-Beckwith (WBS)**
    - **Syndrome de Sotos ou gigantisme cérébral**
    - **Syndrome de Weaver**
    - **Syndrome de Marschall-Smith**
    - **Syndrome de Banayan**





# How big is too big? The perinatal consequences of fetal macrosomia



**TABLE 1**  
**Sociodemographic and clinical characteristics by birthweight (g)**

	2500-3499	3500-4499	4500-4999	5000 or greater	
Gestational age in completed weeks (mean ± SD)	39.2 ± 1.5	39.7 ± 1.4	39.9 ± 1.4	39.9 ± 1.4	
Infant sex (% male)	45.6	56.8	68.4	70.7	
Multiparous (%)	55.3	61.8	68.6	71.2	↑
Mother's age 35 years or older (%)	13.9	16.1	19.8	22.7	↑
Mother's education 13 years or more (%)	53.8	61.6	63.0	61.6	
Marital status (% legally married)	75.3	81.9	85.4	84.9	↑
Smoking (%)	19.5	9.9	5.9	5.7	↓
Maternal diabetes (%)	2.4	2.8	5.4	11.5	→

All P values < .001.

Zhang. How big is too big? The perinatal consequences of fetal macrosomia. Am J Obstet Gynecol 2008.



# Tumour surveillance in Beckwith–Wiedemann syndrome and hemihyperplasia: A critical review of the evidence and suggested guidelines for local practice

Journal of Paediatrics and Child Health 42 (2006) 486–490



Tumour surveillance	Frequency	Until age
Abdominal ultrasound	3 monthly	8 years
Serum alpha-fetoprotein	3 monthly	4 years
Complete physical examination including abdominal palpation by physician	6 monthly†	Indefinite

## EPIGENOTYPING AS A TOOL FOR THE PREDICTION OF TUMOR RISK AND TUMOR TYPE IN PATIENTS WITH BECKWITH-WIEDEMANN SYNDROME (BWS)

The Journal of Pediatrics 2004

	Group I (UPD)	Group II (IC1)	Group III (IC2)	Group IV
Methylation pattern	KCNQIOT1 ↓ HI9 ↑	HI9 ↑	KCNQIOT1 ↓	Normal
Total: tumor frequency in sample 16% (63/287)				
Percentage within group (%)	20/56 (36)	15/29 (52)	8/124 (6)	17/78 (22)
P value*	.010	.0003	<.0001	.848
Wilms' tumor/total number of tumors	10/18	11/11	0/7	9/13
P value†	.624	.0010	.0003	.547

# Macrosome

## La vision obstétricale?



> 4,5 ?

RESEARCH

2008

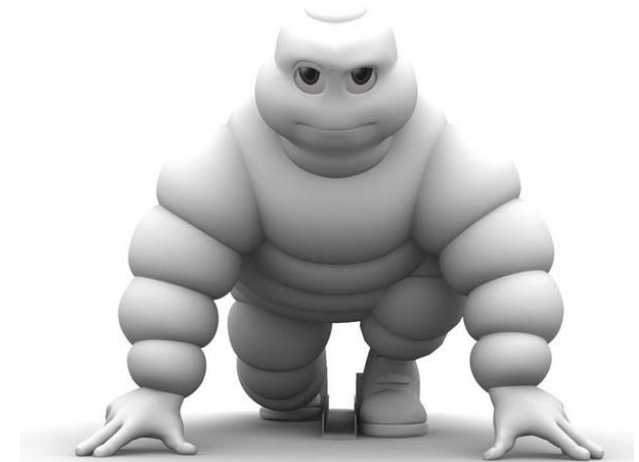
[www.AJOG.org](http://www.AJOG.org)

OBSTETRICS

### How big is too big? The perinatal consequences of fetal macrosomia

Xun Zhang, PhD, Adriana Decker, MD, Robert W. Platt, PhD, Michael S. Kramer, MD

517.e1



**CONCLUSION:** Birthweight greater than 4500 g, and especially greater than 5000 g, is associated with increased risks of perinatal and infant mortality and morbidity.

**Key words:** birth asphyxia, birth injury, macrosomia, stillbirth, sudden infant death syndrome

# Macrosomie

## Les causes de décès



> 4,5 ?

**TABLE 3**  
Major causes of infant deaths by birthweight

Birthweight (g)	Asphyxia	Congenital anomalies	Infection	Postneonatal SIDS
<b>2500-3499</b>				
n (%)	353 (0.01)	1422 (0.05)	142 (0.00)	2230 (0.07)
OR (95% CI)	1.1 (0.9 to 1.3)	2.8 (2.5 to 3.2)	1.5 (1.1 to 2.1)	1.4 (1.3 to 1.6)
<b>3500-4499</b>				
n (%)	295 (0.01)	448 (0.02)	70 (0.00)	1100 (0.04)
OR (95% CI)	1.0	1.0	1.0	1.0
<b>4500-4999</b>				
n (%)	26 (0.02)	12 (0.01)	4 (0.00)	32 (0.03)
OR (95% CI)	2.3 (1.5 to 3.5)	0.6 (0.3 to 1.2)	1.8 (0.7 to 5.0)	0.9 (0.6 to 1.3)
<b>5000 or greater</b>				
n (%)	12 (0.11)	4 (0.04)	2 (0.02)	8 (0.07)
OR (95% CI)	10.5 (5.7 to 19.2)	2.3 (0.9 to 6.3)	8.9 (2.2 to 36.6)	2.3 (1.1 to 4.6)

ORs estimated from multiple logistic regression models adjusted for gestational age, sex, parity, maternal age, maternal education, maternal marital status, maternal diabetes, and smoking.

Zhang. How big is too big? The perinatal consequences of fetal macrosomia. *Am J Obstet Gynecol* 2008.

# Macrosomie et Adaptation

> 4,5 ?

: Apgar, DR, Cn, traumatisme obstétrical...

TABLE 4  
Neonatal morbidity by birthweight

Birthweight (g)	5-minute Apgar score less than 4	Neonatal seizures	Ventilation less than 30 min	Ventilation 30 minutes or more	Birth injury	Meconium aspiration syndrome
<b>2500-3499</b>						
%	0.12	0.06	2.01	0.41	0.24	0.15
OR (95% CI)	1.3 (1.2 to 1.3)	1.1 (1.0 to 1.2)	0.9 (0.9 to 0.9)	1.2 (1.1 to 1.2)	0.6 (0.5 to 0.6)	0.9 (0.8 to 0.9)
<b>3500-4499</b>						
%	0.09	0.05	2.13	0.34	0.40	0.18
OR (95% CI)	1.0	1.0	1.0	1.0	1.0	1.0
<b>4500-4999</b>						
%	0.17	0.08	2.87	0.57	0.91	0.26
OR (95% CI)	1.8 (1.5 to 2.1)	1.6 (1.3 to 2.0)	1.4 (1.3 to 1.4)	1.7 (1.5 to 1.8)	2.4 (2.2 to 2.5)	1.5 (1.3 to 1.7)
<b>5000 or greater</b>						
%	0.60	0.17	4.20	1.31	1.38	0.37
OR (95% CI)	6.4 (4.9 to 8.4)	3.3 (2.1 to 5.3)	2.0 (1.8 to 2.2)	3.7 (3.1 to 4.5)	3.5 (3.0 to 4.2)	2.1 (1.5 to 2.9)

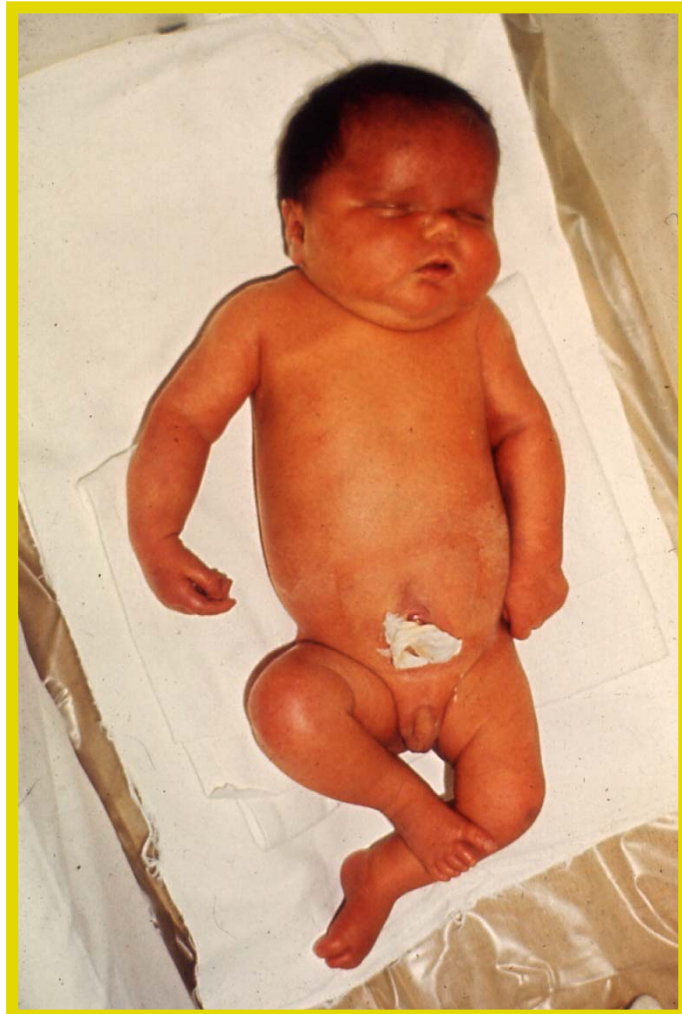
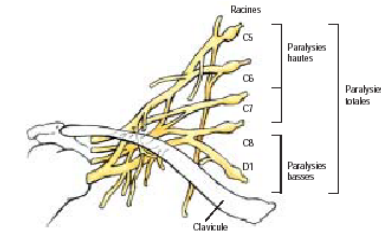
22%

36%

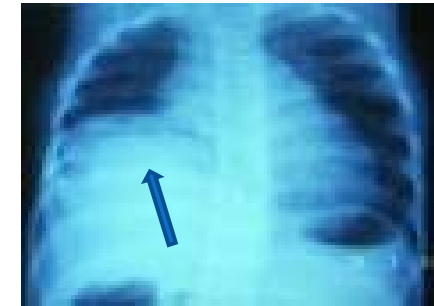
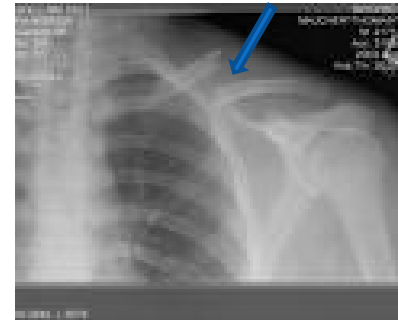
51%

ORs estimated from multiple logistic regression models adjusted for gestational age, sex, parity, maternal age, maternal education, maternal marital status, maternal diabetes, and smoking.  
Zhang. How big is too big? The perinatal consequences of fetal macrosomia. *Am J Obstet Gynecol* 2008.

# Macrosomie et Paralysie du Plexus brachial

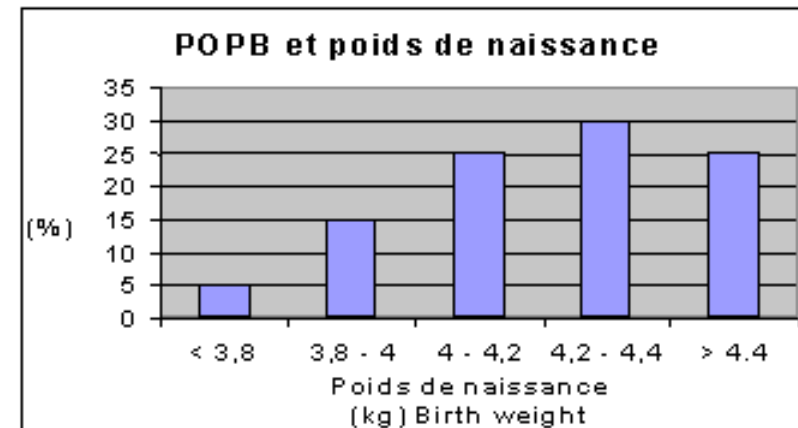


> 4,5 kg



2%

6 % vs 1.5‰



# Macrosomie et Plexus Brachial

- **Information des parents +++**
- **Description initiale +++**
  - ▣ **Bilan initial**
    - **Mobilité des doigts, extension, opposition du pouce**
    - **ROT (Bicipital C5-6)**
  - ▣ **Recherche de lésions associées**
    - **Fracture de clavicule**
      - **Autres fractures**
    - **Paralysie diaphragmatique**
    - **Claude Bernard horner**
- **Surveillance quotidienne**
  - ▣ **Signes de récupération**
  - ▣ **Testing musculaire**
- **Nursing, Kiné, orthopédie**



# Macrosomie et troubles métaboliques



- Surtout macrosomie pour l'âge gestationnel
  - ▣ > 90<sup>e</sup> percentile
  - ▣ En cas de diabète maternel préexistant ou gestationnel
- Moins fréquent
  - ▣ chez le nouveau-né de mère diabétique à terme eutrophe
  - ▣ En cas de diabète gestationnel traité par régime seul
- Hypoglycémie
  - ▣ Apport / poids
  - ▣ Capacité d'ingestion /oralité
- Hypocalcémie



# Neonatal outcome of macrosomic infants: an analysis of a two-year period

Zita Gyurkovits<sup>a,\*</sup>, Karola Kálló<sup>b</sup>, Judit Bakki<sup>a</sup>, Márta Katona<sup>c</sup>, Tamás Bitó<sup>a</sup>, Attila Pál<sup>a</sup>, Hajnalka Orvos<sup>a</sup>

Outcome measures of the macrosomic neonates of diabetic and non-diabetic mothers.

	Diabetic n = 43	Non-diabetic n = 367	p
★ Caesarean section	23 (53%)	179 (49%)	0.56
Males	24 (56%)	256 (70%)	0.06
Umbilical cord pH < 7.2	2 (4.7%)	➔ 68 (19%)	0.02*
Apgar score < 7 at 5 min	0 (0%)	4 (1.1%)	0.49
Congenital anomalies	4 (9.3%)	14 (3.8%)	0.10
★ Hypoglycaemia ➔	11 (26%)	14 (3.8%)	<0.001*
Polycythaemia	4 (9.3%)	15 (4.1%)	0.06
★ Hyperbilirubinaemia	13 (30%)	63 (17%)	0.04*
Clavicle fracture	0 (0%)	7 (1.9%)	0.36
Cephalhaematoma	3 (7%)	12 (3.3%)	0.22
★ Adrenal haemorrhage	1 (2.3%)	3 (0.8%)	0.34
Respiratory disorder	4 (9.3%)	17 (4.6%)	0.19
NICU admission	4 (9.3%)	17 (4.6%)	0.19
Mechanical ventilation	1 (2.3%)	1 (0.3%)	0.07
★ Cardiomyopathy ➔	2 (4.7%)	1 (0.3%)	0.01*
Neurological disorder	1 (2.3%)	8 (2.2%)	0.96

\* Significance at  $p < 0.05$ .

Contents lists available at SciVerse ScienceDirect

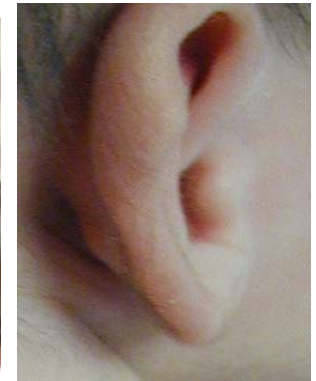
European Journal of Obstetrics & Gynecology and  
Reproductive Biology

# Nouveau-né de mère diabétique : Hyperinsulinisme / Morphologie



## ▣ Macrosomie

- accroissement du panicule adipeux
- Répartition facio-tronculaire
- Pilosité, oreilles
- Césarienne



## ▣ Retard de résorption du liquide pulmonaire

## ▣ Hypertrophie myocardique : 15-30%

- SIV > +2DS

## ▣ Cytostéatonécrose (rare)

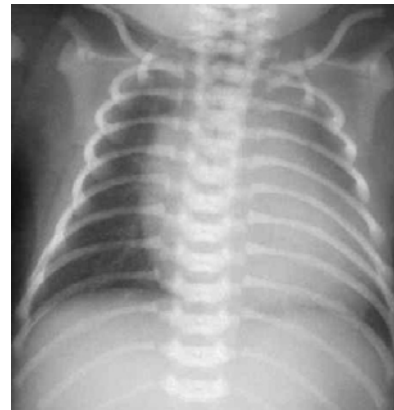
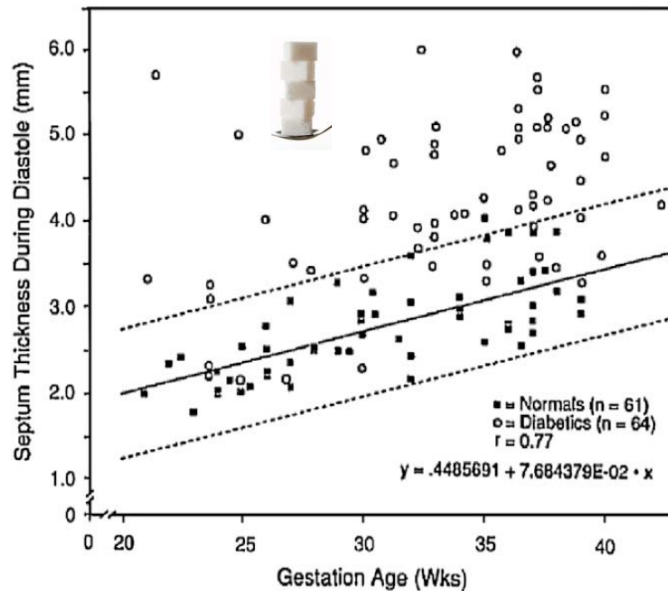
- Lipolyse, inflammation sous cutanée, hypercalcémie tardive



# Complications cardio-pulmonaires



## Corrélation SIV/ AG



- HSV 75%
- Linéaire
- 12 vs 8%
  - P<0.05
- Symptomatique : 12%

- DRT
  - Césarienne
  - Hyperinsulinisme
- Polyglobulie
- HTAP



Tachypnée T →



# Nouveau-né de mère diabétique

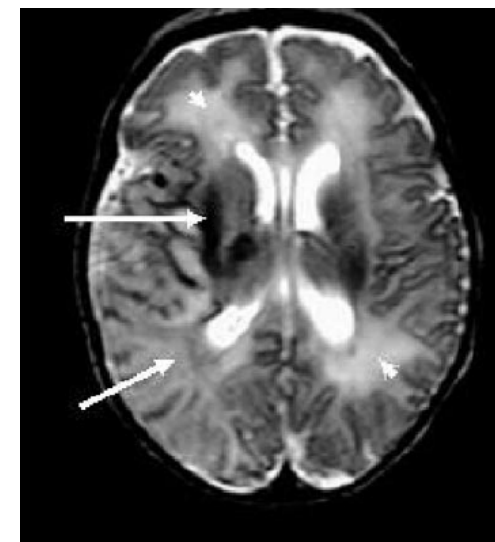
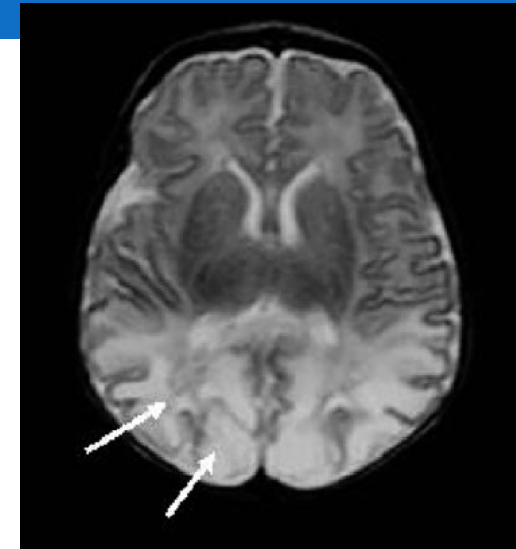
## Métabolisme



- **Hypoglycémie : <math><2.2\text{ mmol/L}</math>**
  - 30-50%
- **Signes cliniques d'hypoglycémie**
  - asymptomatique
  - Trémulation , Hyperexcitabilité
  - Accès de cyanose (apnées, sueurs, convulsions...)
  - hypotonie , Cn
- ***Dextro et confirmer l'hypo par glycémie sg***
- **Hypoparathyroïdisme transitoire**
  - Hypocalcémie : 30% (Mg)
  - Un-alpha
- **Polyglobulie (30%)**
  - HTAP, Ictère, Thrombose

Burns. 2008. p 65

- **Hypoglycémies symptomatiques**
  - < 1,5 mmol/l
  - 2/3 transitoires
  - 80% H0-H48 (57% 0 J1)
- **Convulsion (83%)**
- **IRM avant J10**
- **Lésions de la substance blanche**
  - 94%
    - Severe 43%
    - Postérieure 29%
    - Hémorragie 30%
    - Noyaux gris 51%



# Macrosome en maternité

## Alimentation précoce ou tétée précoce < 30'

- Dextro 1h après
  - Dextro avant le départ de la salle de naissance
  - Dextro > 2, 2 mmoles/l
- Alimentation/3h
- Lait Pré/ liquigen
- Vitamine D précoce : double dose
  - Calcémie à J3
- Surveillance clinique, T° , dextro /48h
  - Arrêt si 4 dextros consécutifs normaux
    - En l'absence de difficultés alimentaires et de troubles du comportement
- Surveillance ictère : polyglobulie : Biliflash

# Outcome

*Seminars in Fetal & Neonatal Medicine* 14 (2009) 119–124  
*Curr Diab Rep* (2014) 14:489



Offspring of diabetic pregnancy: Long-term outcomes

- **Diabète**
- **Obésité**
- **Syndrome métabolique : HTA...**
- **Troubles attentionnels, motricité fine...**

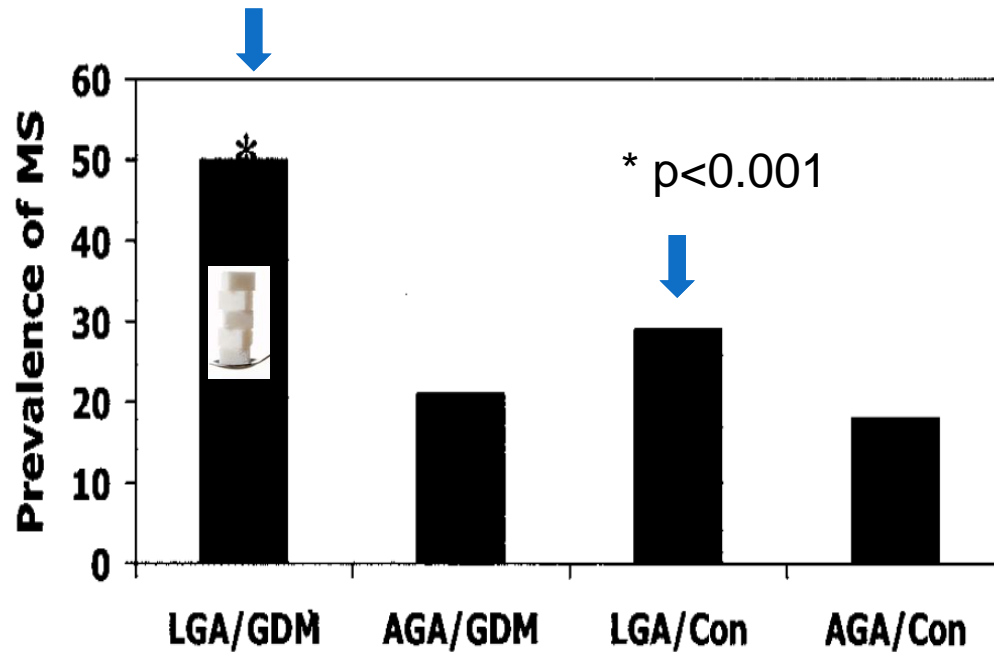


Le risque d'obésité et de diabète non insulino-dépendant est augmenté chez ces enfants. Une surveillance régulière des enfants et une éducation nutritionnelle de la mère et de l'enfant sont donc nécessaires. L'efficacité de ces mesures sur la prévention du diabète non insulino-dépendant doit faire l'objet d'études cliniques tout comme l'évaluation des difficultés neuro-cognitives.

# Outcome : Sd metabolique

## Follow up : 11 ans

*Pediatrics* 2005;115:e290



- BMI > 85<sup>e</sup> p
- HTA > 90<sup>e</sup> p
- TG/HDL
- Equilibre gly

Variables	Hazard Ratio	P Value	95% CI for Hazard Ratio
LGA versus AGA	2.19	.006	1.25–3.82
Maternal obesity* versus nonobese	1.81	.039	1.03–3.19
GDM versus control	1.44	.191	0.83–2.50
Male versus female	1.52	.133	0.88–2.61

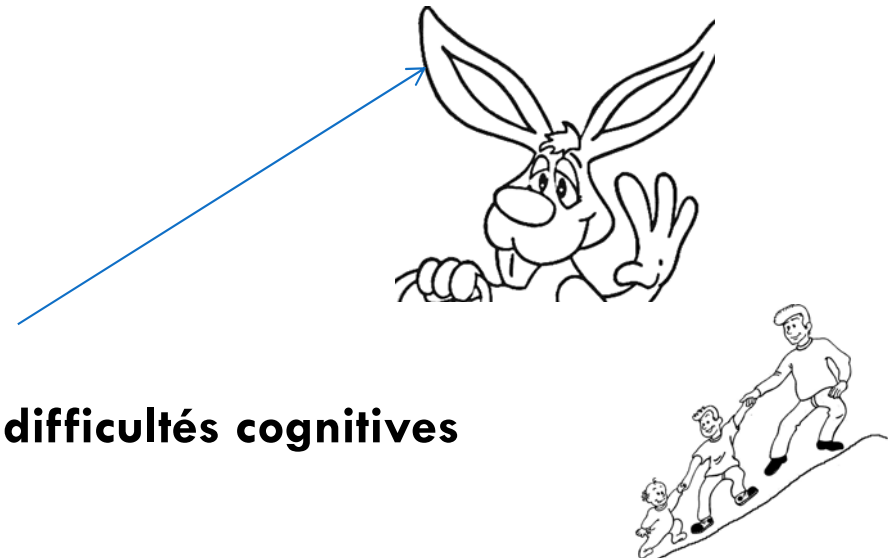
- Facteur aggravant**
  - Obésité maternelle**



# Take home messages

## Surveillance des enfants de plus de 4 kg

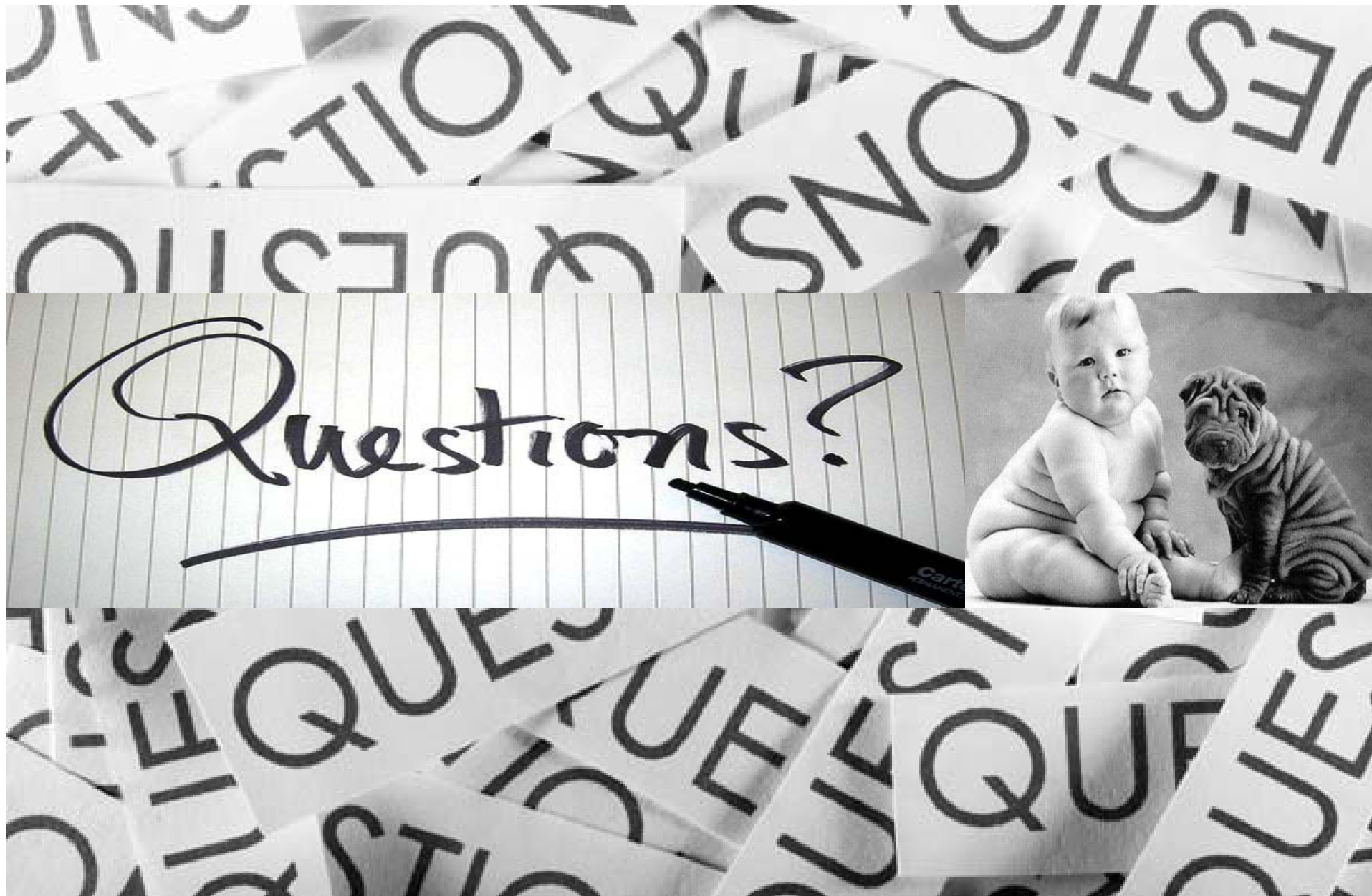
- **Macrosome**
  - ▣ **Colosse aux pieds d'argile**
- **Out come**
  - ▣ **Mortalité périnatale**
  - ▣ **Morbidité obstétricale : traumatisme obstétricale, asphyxie périnatale**
  - ▣ **Infection, détresse respiratoire, transfert, durée d'hospitalisation**
  - ▣ **Trouble métabolique**
- **Nouveau-nés de mère diabétique**
  - ▣ **Homéostasie glucidique**
  - ▣ **Malformation**
  - ▣ **Ictère, Hypertrophie myocardique**
- **Quelques rares syndromes**
- **Suivi à long terme : Sd métabolique, difficultés cognitives**



# Indications de transfert



- **Pas d'hospitalisation systématique**
  - **Ni en cas de PPB ou de fracture de la clavicule**
- **La macrosomie chez un enfant de mère diabétique signe le plus souvent un mauvais équilibre maternel et justifie une surveillance attentive de l'alimentation et de l'équilibre glycémique**
- **Les difficultés d'adaptation à la vie extra utérine en cas d'asphyxie périnatales majorent le risque d'hypoglycémie**
- **Hypoglycémies rebelles**
  - **→transfert en UK/neonatalogie**
    - **En cas d'hyperinsulinisme: Diazoxide PROGLYCEM 5-20 mg/kg/j (débuter à 10 mg/kg) ou Glucagon (GLUCAGEN en IVC 1mg/j)**



24/10/2014

SFMP Lyon 2014  
AUDIPOG

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Back up

# la dystocie des épaules



## Risk of shoulder dystocia (adapted from Rouse, 1996)

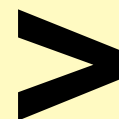
Birth weight (g)

< 4000  
4000–4499  
> 4500



Maternal diabetes: incidence of shoulder dystocia (95% CI)

2.2% (0.6–3.7%)  
13.9% (4.7–23%)  
52.5% (37.5–83.3%)



No maternal diabetes: incidence of shoulder dystocia (95% CI)

0.7% (0.3–1.0%)  
6.7% (3.4–9.0%)  
14.5% (9.8–22.0%)

Outcome	Adjusted odds ratio, 95% CI birthweight 4000–4499 g	Adjusted odds ratio, 95% CI birthweight 4500–4999 g	Adjusted odds ratio, 95% CI birthweight ≥5000 g
Cesarean birth	1.69 (1.62–1.76)	2.99 (2.76–3.24)	5.46 (4.40–6.78)
Shoulder dystocia	6.29 (5.83–6.77)	13.05 (11.70–14.56)	17.52 (13.54–22.68)

	%	Birthweight 2500–3999 g (%)	Birthweight 4000–4499 g (%)	Birthweight 4500–4999 g (%)	Birthweight ≥5000 g (%)
Cesarean birth	13.9	21.1*	32.6*	49.2*	
Shoulder dystocia	1.5	8.8*	17.4*	23.3*	



# Importance du contrôle du diabète maternel pendant la grossesse

Comparison of complications for babies delivered after strict peri-conceptual glycaemic control and normal glycaemic control in women with pre-gestational diabetes mellitus (adapted from Merlob and HOD<sup>52</sup>).

Complication	Prevalence when strict pre-conceptual glycaemic control (%) <sup>52</sup>	Prevalence in other studies of infants of mothers with pre-gestational diabetes
→ Macrosomia	13.7% ↓	21% <sup>1,54</sup>
Hypoglycaemia	26.3%	27–47% <sup>12,55,56</sup>
Hypocalcaemia	7.5% ?	4–50% <sup>5,12,57,58</sup>
Polycythaemia	7.5%	5–29% <sup>12,59,60</sup>
Hyperbilirubinaemia	19.4%	20–25% <sup>12,52</sup>
Respiratory distress syndrome	3.7% ↓	17–92% <sup>25</sup>
→ Ventricular septal hypertrophy	7.5% overall (8.3% in macrosomic infants, 1.8% in normally grown infants) ↓	75% <sup>38</sup>

- Surveillance glycémique +++

# Outcome Macrosome ?



- **Nouveau-nés de mères diabétiques**
  - Trouble de l'attention
  - Trouble de la motricité fine
  - QI similaires si diabète maternel équilibré
    - *Pediatr endocrinol rev* 2005 p 104 Ornoy A 2
  - Nouveau-nés de mère diabétique
    - Co-morbidité fréquente
      - *Curr Diab Rep* 2014. p 489. Fraser A
      - *Diabet. Med.* 2011. p838–844. Clausen TD
- **Des études sont nécessaires**

# Neonatal outcome of macrosomic infants: an analysis of a two-year period

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Outcome measures of control group and macrosomic neonates.

	Control	Macrosomic	<i>p</i>
Total	4757	410	
→ Caesarean section	1898 (39.9%)	202 (49.3%)	↑ <0.001*
Males	2322 (48.8%)	280 (68.3%)	<0.001*
Umbilical cord pH < 7.2	705 (14.8%)	70 (17.0%)	0.22
Apgar score < 7 at 5 min	87 (1.8%)	4 (0.9%)	0.22
Congenital anomalies	185 (3.8%)	18 (4.3%)	0.68
→ Hypoglycaemia	138 (2.9%)	25 (6.1%)	↑ <0.001*
Polycythaemia	161 (3.4%)	19 (4.6%)	0.19
→ Hyperbilirubinaemia	1446 (30.4%)	76 (18.5%)	<0.001*
Clavicle fracture	43 (0.9%)	7 (1.7%)	0.11
Cephalhaematoma	135 (2.8%)	15 (3.6%)	0.34
→ Adrenal haemorrhage	7 (0.15%)	4 (0.98%)	<0.001*
Respiratory disorder	305 (6.4%)	21 (5.1%)	0.26
NICU admission	210 (4.4%)	21 (5.1%)	0.58

\* Significance at  $p < 0.05$ .

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