



# Management of Preeclampsia and Eclampsia

Igor Michalec

# Hypertension in pregnancy

- 5 – 10 % all pregnant
- severe impacts on the health of mothers and fetuses
- among the 3 most common causes of death in pregnancy

Report of the National High Blood Pressure Education Program Working Group on High Blood Pressure in Pregnancy

National High Blood Pressure Education Program Working Group on High Blood Pressure in Pregnancy\*

*Bethesda, Maryland*

This report updates the 1990 "National High Blood Pressure Education Program Working Group Report on High Blood Pressure in Pregnancy" and focuses on classification, pathophysiologic features, and

# Classification of hypertension in pregnancy

## Chronic hypertension

- Před těhotenstvím, event. prvně zjištěná v graviditě a přetrvává < 6 týdnů po porodu
- Esenciální nebo sekundární

## Gestational hypertension

- Ve 2. polovině gravidity, vymizí nejpozději do 12 týdnů
- Nemyzli-ii = chronická

## Preeklampsia - eklampsia

- Hypertenze a proteinurie ve 2. polovině gravidity
- Eklampsie – encefalopatie, tonicko-klonické křeče, porucha vědomí, amnézie
- HELLP syndrom

## Preeclampsia superimposed

- Preeklampsie u chronické hypertenze
- Hypertenze před graviditou, proteinurie v 2. polovině gravidity

# Chronic Hypertension vs. preeclampsia

## CHRONIC HYPERTENSION

**Esencial**

**Secondary**

nephrogenic, endocrine, cardiac,  
central

**Assessment of organ disability in the  
introduction**

ventricular hypertrophy,  
retinopathy, renal function

**Risk of pre-eclampsia up to 25%**

# Chronic Hypertension vs. preeclampsia

## CHRONIC HYPERTENSION

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nephrogenic, endocrine, cardiac,  
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## PREEKLAMPSIA

**Systemic disease**

Decreased organ perfusion

**Damage to kidney, placenta, liver,  
brain and other organs**

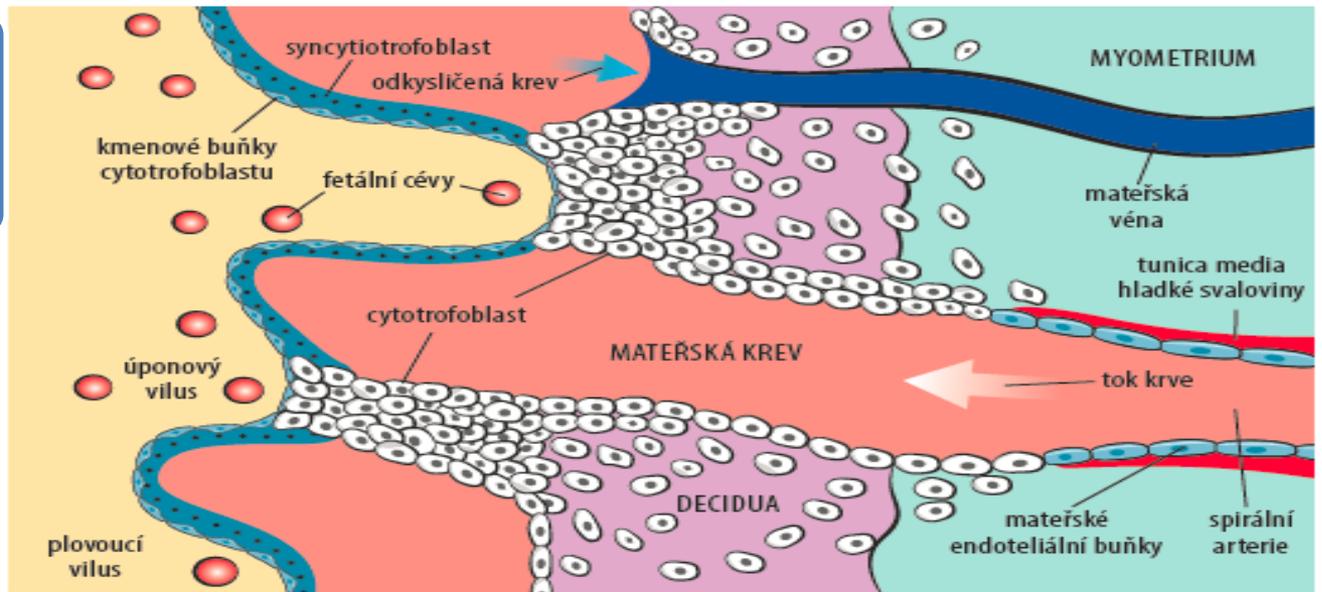
Etiology - multifactorial

**Disorders of trophoblast invasion**

**Endothelium activation**

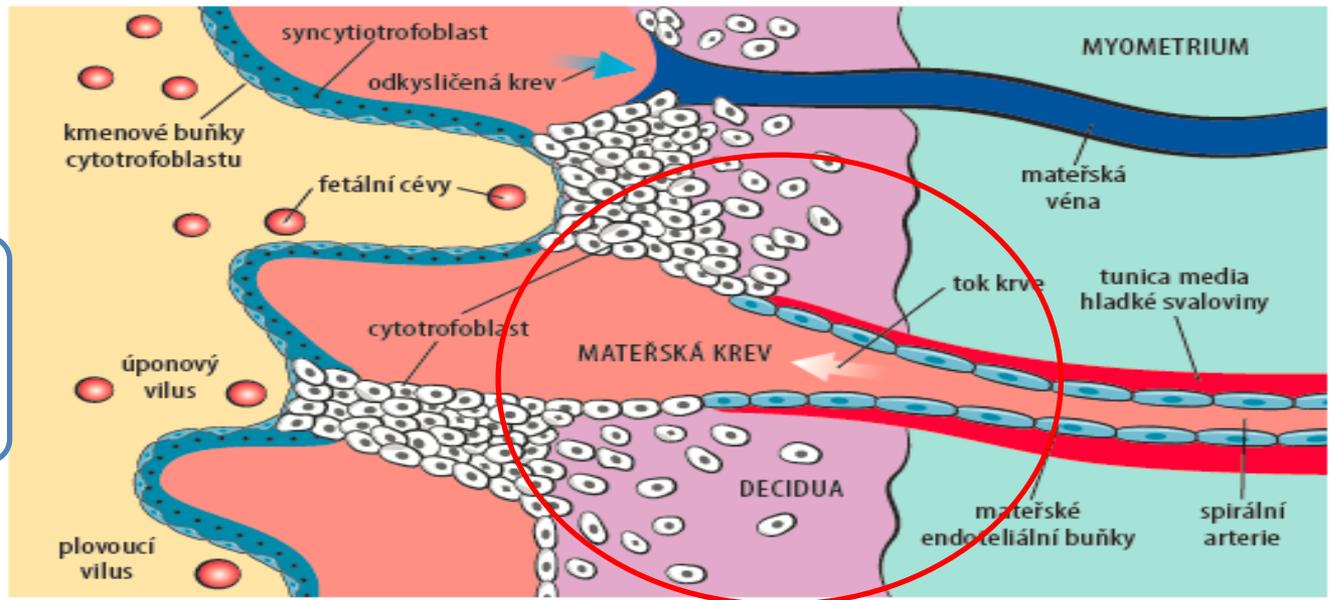
# INVASION OF TROFBLAST

Normal situation

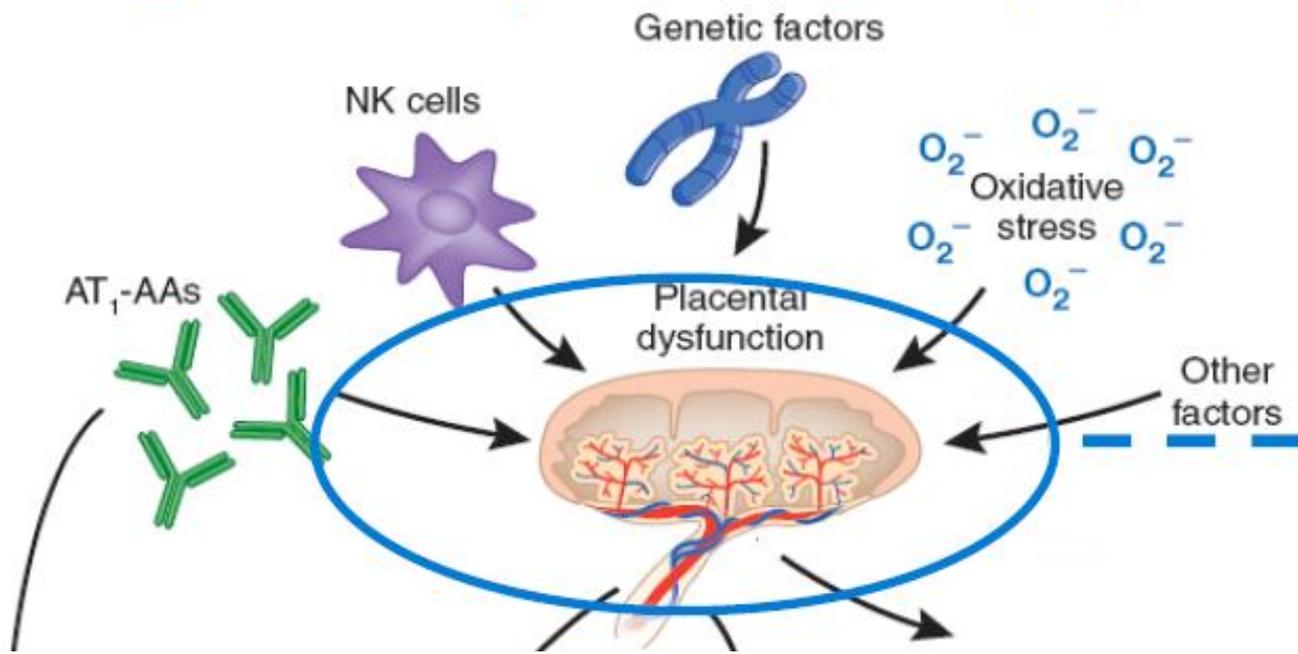


Obr. 4.2a Fyziologická placentace

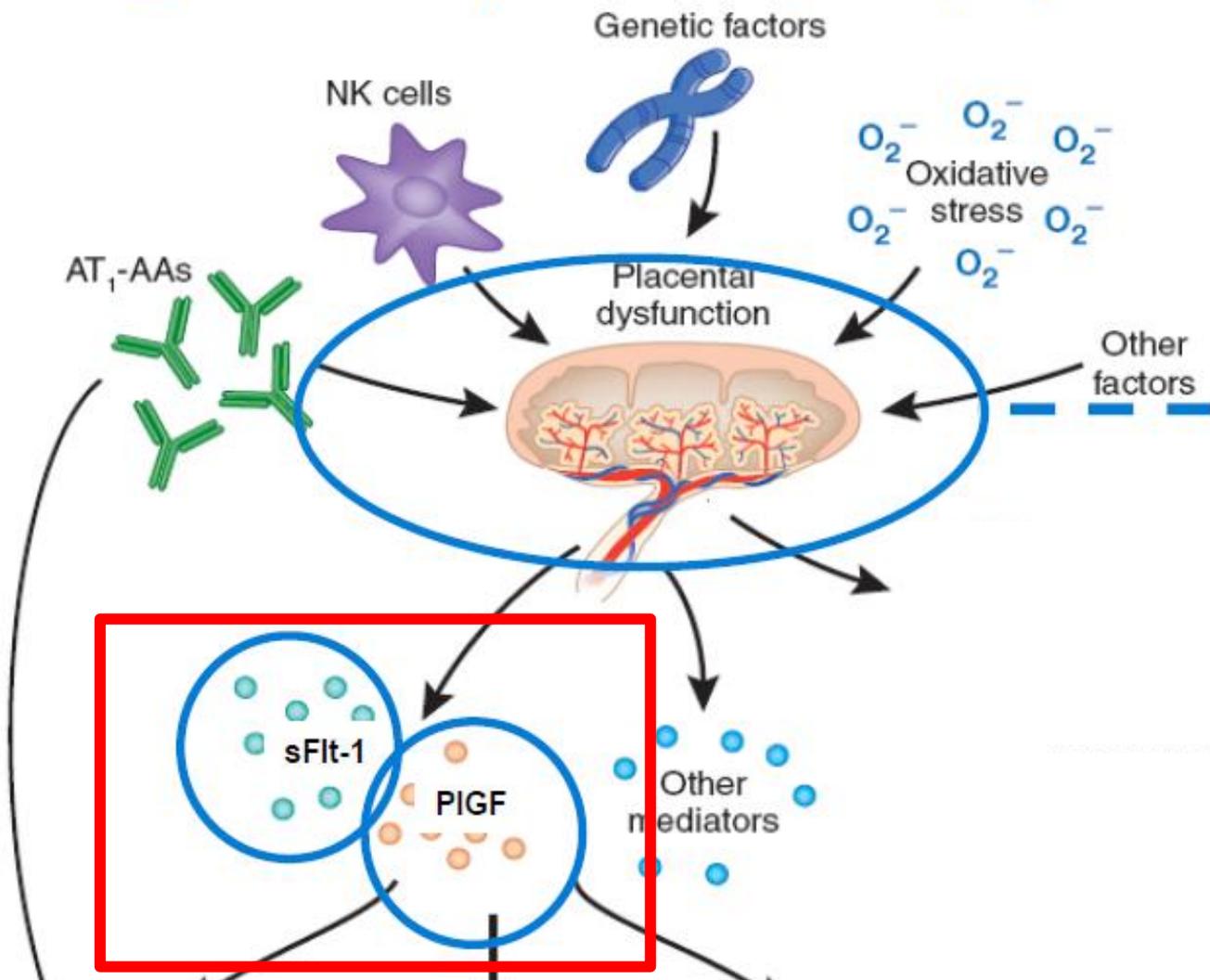
Preeklampsia



Obr. 4.2b Abnormální placentace u preeklampsie



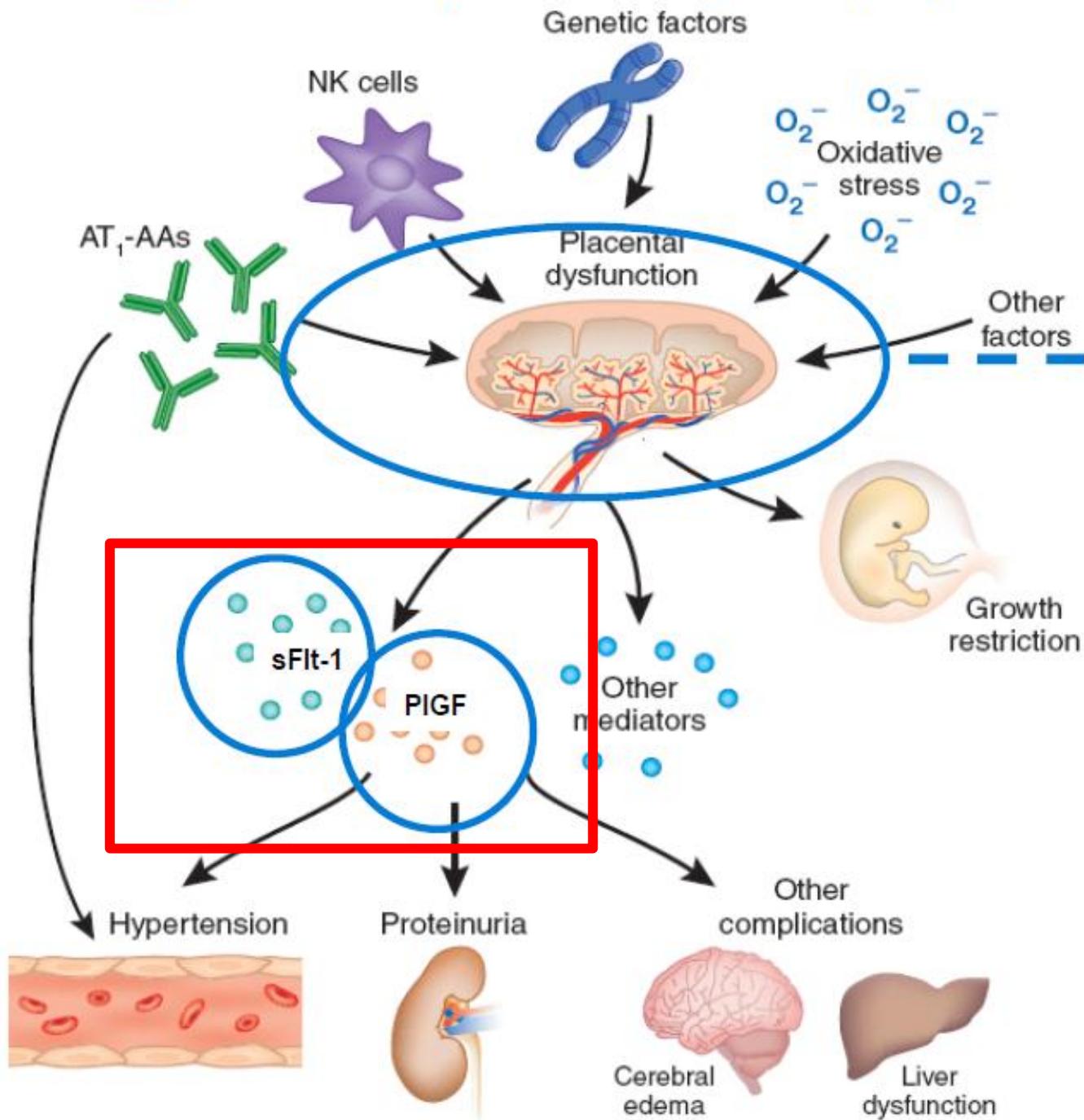
The first half of pregnancy  
placental disorder



The first half of pregnancy  
placental disorder



Endothelial dysfunction  
Antiangiogenic factors



The first half of pregnancy  
placental disorder



Endothelial dysfunction  
Antiangiogenic factors



The second half of pregnancy

- generalized disorder
- preeclampsia / eclampsia / HELLP
- Growth restriction of the fetus

Premature delivery  
Abrupte of the placenta  
Intrauterine death

# Preeklampsia

**4 - 8% of the pregnant population**

**9 - 26% of maternal deaths**

**15% of premature births**

hypertension + proteinuria + edema

**Early / Late**

**The most serious complications**

**Eclampsia**

**HELLP**

**Current best practice in the management of hypertensive disorders in pregnancy.**

[Townsend R<sup>1</sup>](#), [O'Brien P<sup>2</sup>](#), [Khalil A<sup>1</sup>](#).

⊕ **Author information**

**Abstract**

Preeclampsia is a potentially serious complication of pregnancy with increasing significance worldwide. Preeclampsia is the cause of 9%-26% of global maternal mortality and a significant proportion of preterm delivery, and maternal and neonatal morbidity. Incidence is increasing in keeping with the increase in obesity, maternal age, and women with medical comorbidities entering pregnancy. Recent

# Diagnosics and monitoring

## HYPERTENZE

*140/90 - WHO 1993*

*severe HT > 160/110*

*heavy HT > 180*



## PROTEINURIE

PCR - 30 mg / mmol - negative predictive value

> 300 mg / day - 24 hour collection



# **Diagnosics and monitoring**

## **LABORATORY EXAMINATION**

**BC, coagulation, urinary urea, urea, creatinine, albumin,  
LDH, (bilirubin, haptoglobin), diuresis**

## **MONITORING FETUS STATUS**

# Diagnostics and monitoring

## NEW SCREENING TEST

- The PROGNOSIS study
- sFlt-1/PLGF ratio



## Predictive Value of the sFlt-1:PLGF Ratio in Women with Suspected Preeclampsia

Harald Zeisler, M.D., Elisa Llorba, M.D., Ph.D., Frederic Chantraine, M.D., Ph.D., Manu Vatish, M.B., Ch.B., D.Phil.,  
Anne Cathrine Staff, M.D., Ph.D., Maria Sennström, M.D., Ph.D., Matts Olovsson, M.D., Ph.D.,  
Shaun P. Brennecke, M.B., B.S., D.Phil., Holger Stepan, M.D., Deirdre Allegranza, B.A., Peter Dilba, M.Sc.,  
Maria Schoedel, Ph.D., Martin Hund, Ph.D., and Stefan Verlohren, M.D., Ph.D.

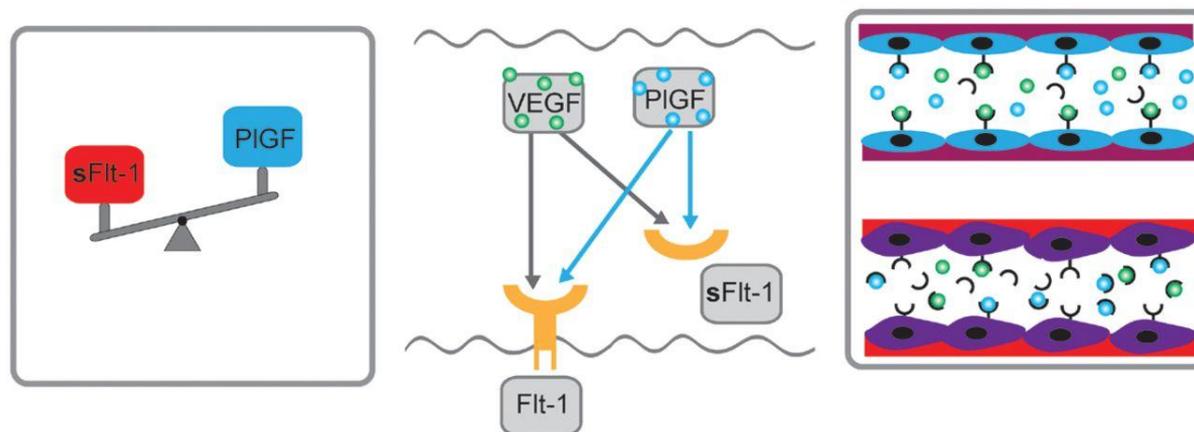
# sFlt-1 / PlGF

sFlt-1 is the dominant anti-angiogenic factor

Its serum levels increase significantly several weeks before clinical manifestation of preeclampsia.

**In contrast, PlGF is an important proangiogenic factor.**

**Its serum levels are significantly reduced already in early stages of pregnancy in patients with subsequent clinical manifestation of preeclampsia.**



# Diagnostics and monitoring

**sFlt-1/PLGF < 38**

Rule out PE within 1 week % (95% CI)	
NPV	99.3 (97.9–99.9)
Sens.	80.0 (51.9–95.7)
Spec.	78.3 (74.6–81.7)

**sFlt-1/PLGF > 38**

Rule in PE within 4 weeks % (95% CI)	
PPV	36.7 (28.4–45.7)
Sens.	66.2 (54.0–77.0)
Spec.	83.1 (79.4–86.3)

# HELLP syndrom

**Low incidence** (**<0.5%**)

**Non-specific symptoms**

**cefalea**

**visa disorders**

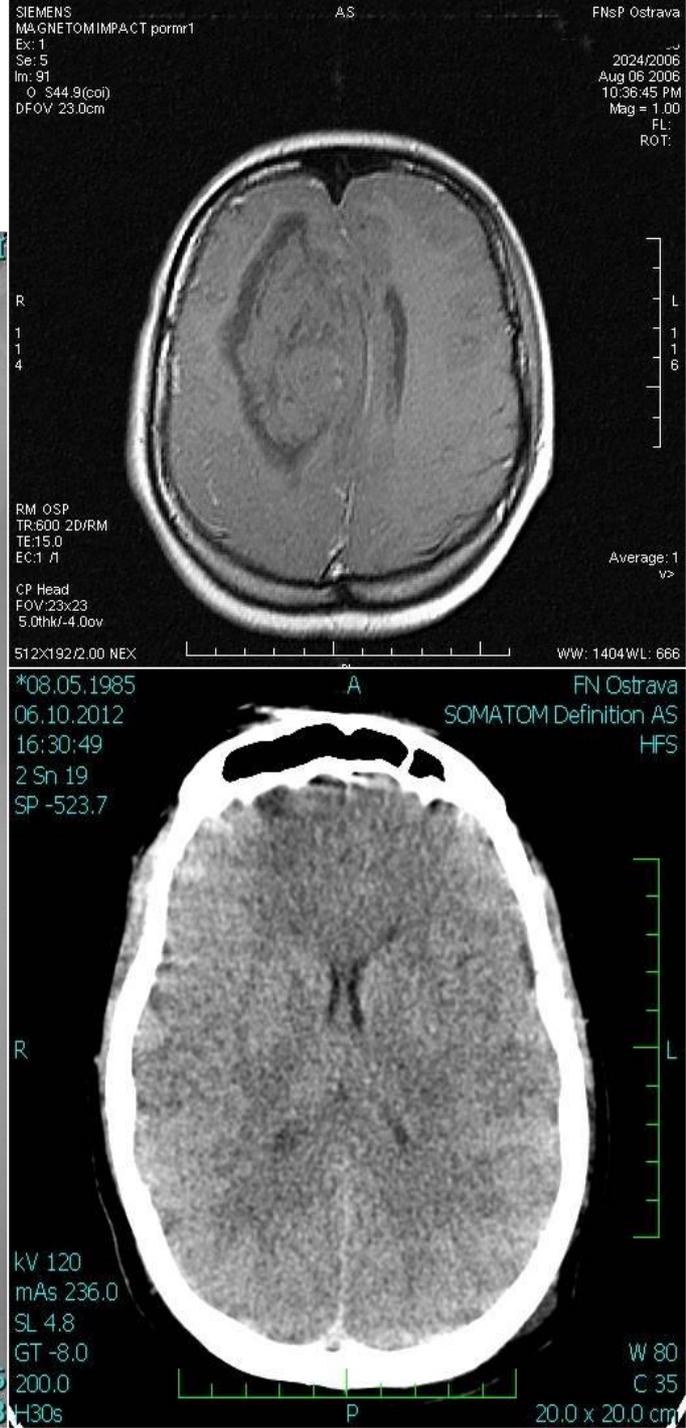
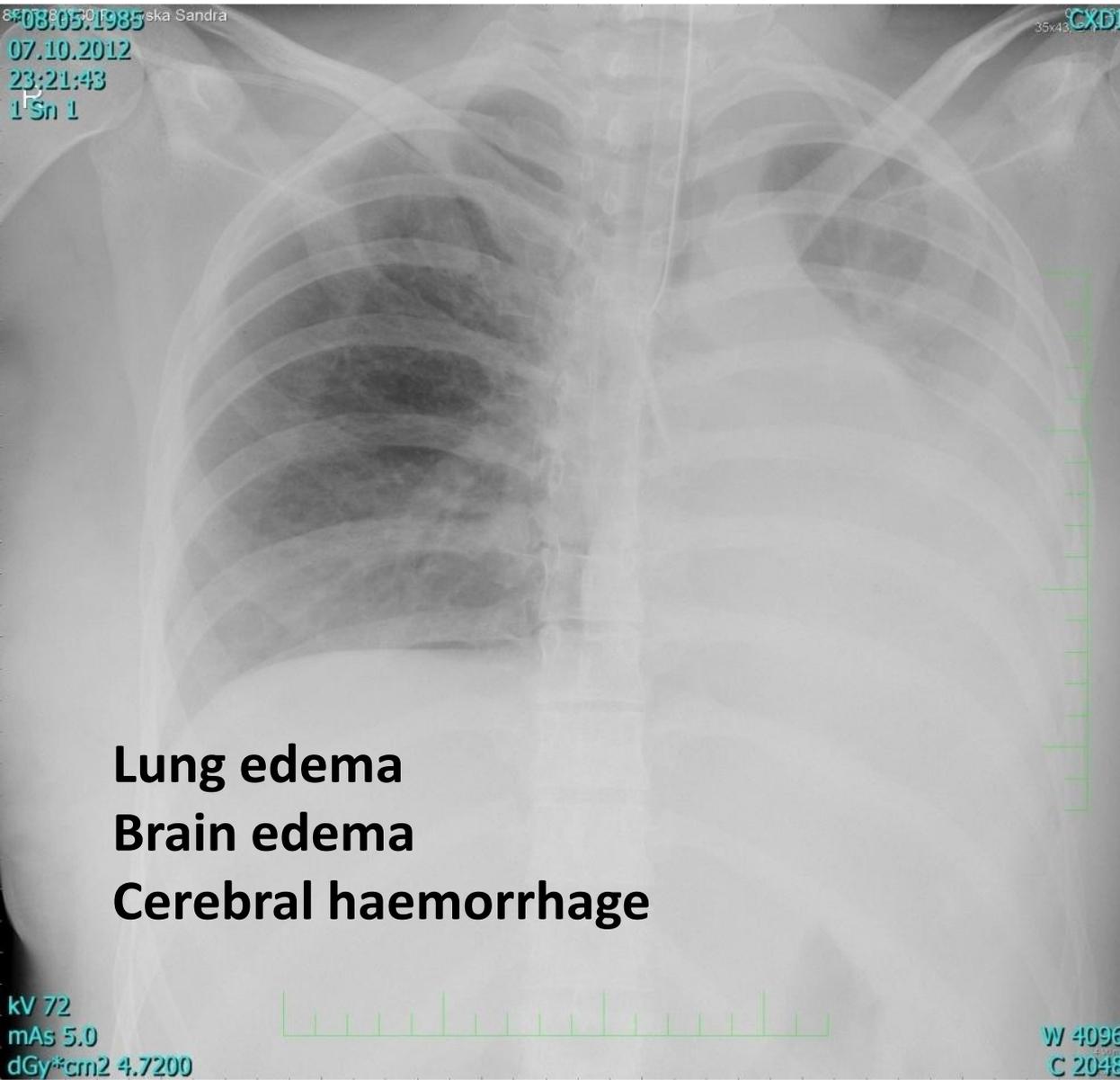
**epigastric pain**

**(Extremely) fast progress**

**Possible death of the mother despite**

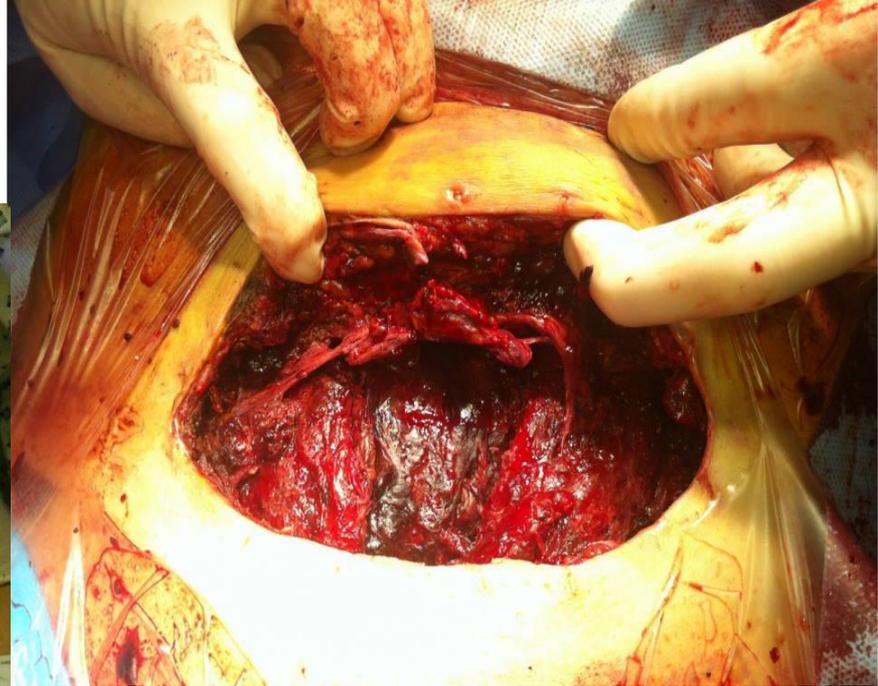
**adequate therapy**

# Clinical manifestations

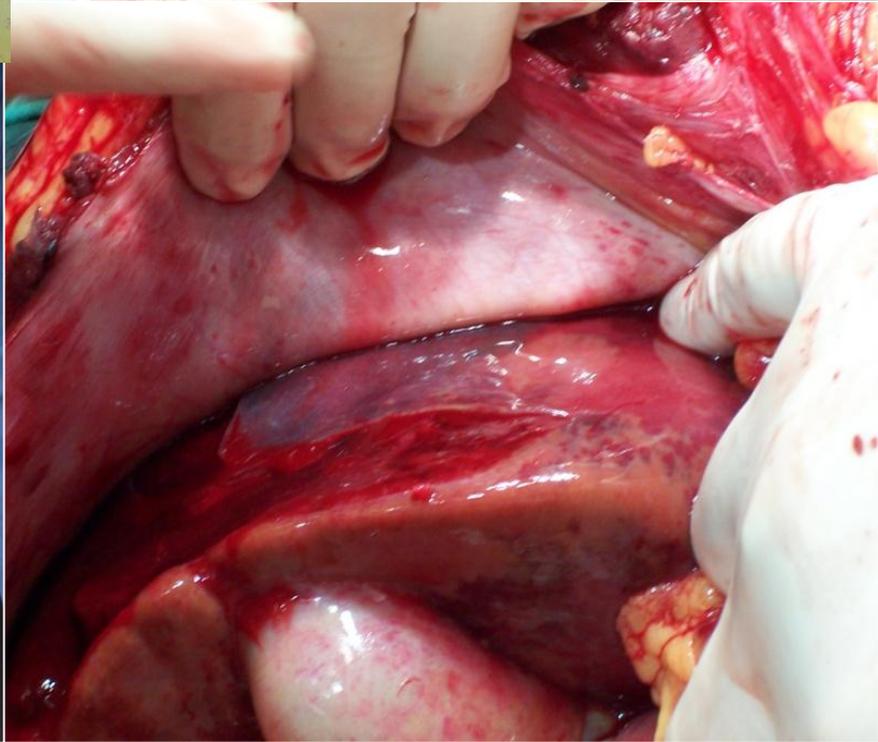


Lung edema  
Brain edema  
Cerebral haemorrhage

# Clinical manifestations



Bleeding manifestations  
DIC  
Liver rupture



# Clinical manifestations



Fluid retention  
Oliguria / anuria  
Maternal death  
Prematurita



# Care for pregnant women with chronic hypertension

## Essential

Light forms often without medication

Target BP 150/100 mmHg

Termination of pregnancy only when impossibility of compensation

## Secondary

As recommended by the specialist and the status of organ functions

## Therapy

YES: methyldopa, beta blockers, calcium channel blockers

NO: diuretics, ACEI

# Care for pregnant women with chronic hypertension

## Light

Monitoring, resting regimen, antihypertensives

Induction in term pregnancy

## Severe (incl. Eclampsia)

End of pregnancy

MgSO<sub>4</sub>, antihypertensive therapy, consistent adjustment of fluid balance

## HELLP

Termination of pregnancy within 24 hours

MgSO<sub>4</sub>, antihypertensives, consistent fluid balance adjustment, DIC treatment

Corticosteroids? Plasmaferesis?

# Risk of cardiovascular disease

## Cardiovascular mortality after pre-eclampsia in one child mothers: prospective, population based cohort study

 OPEN ACCESS

Rolv Skjaerven *professor*<sup>1,2</sup>, Allen J Wilcox *senior investigator*<sup>3</sup>, Kari Klungsoyr *associate professor*<sup>1,2</sup>, Lorentz M Irgens *professor*<sup>1,2</sup>, Bjørn Egil Vikse *associate professor*<sup>4,5</sup>, Lars J Vatten *professor*<sup>6</sup>, Rolv Terje Lie *professor*<sup>1,2</sup>

# Risk of cardiovascular disease

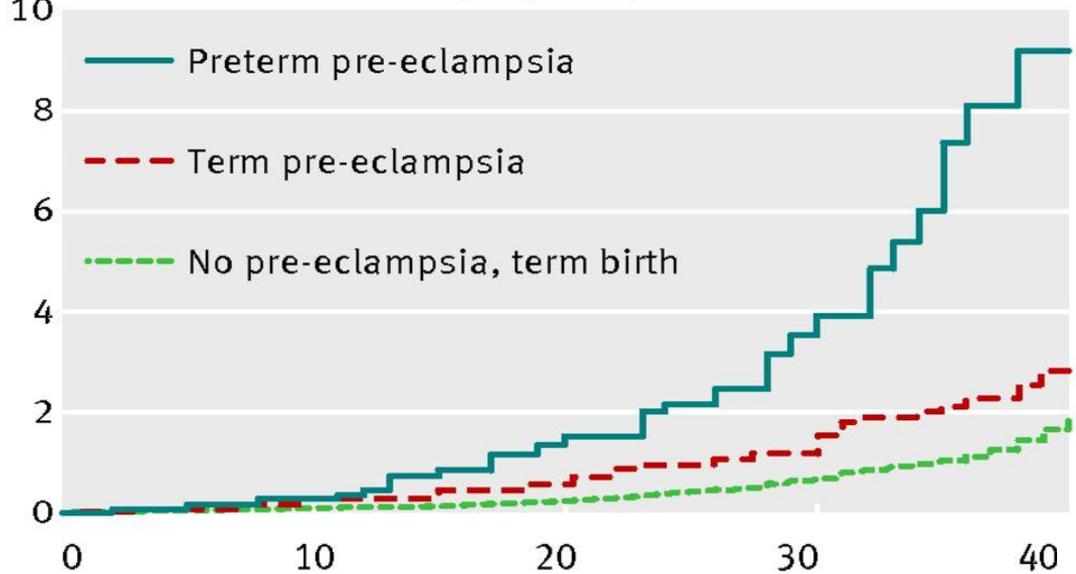
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Cardiovascular death (%)

Women with 1 lifetime pregnancy



# Risk of cardiovascular disease

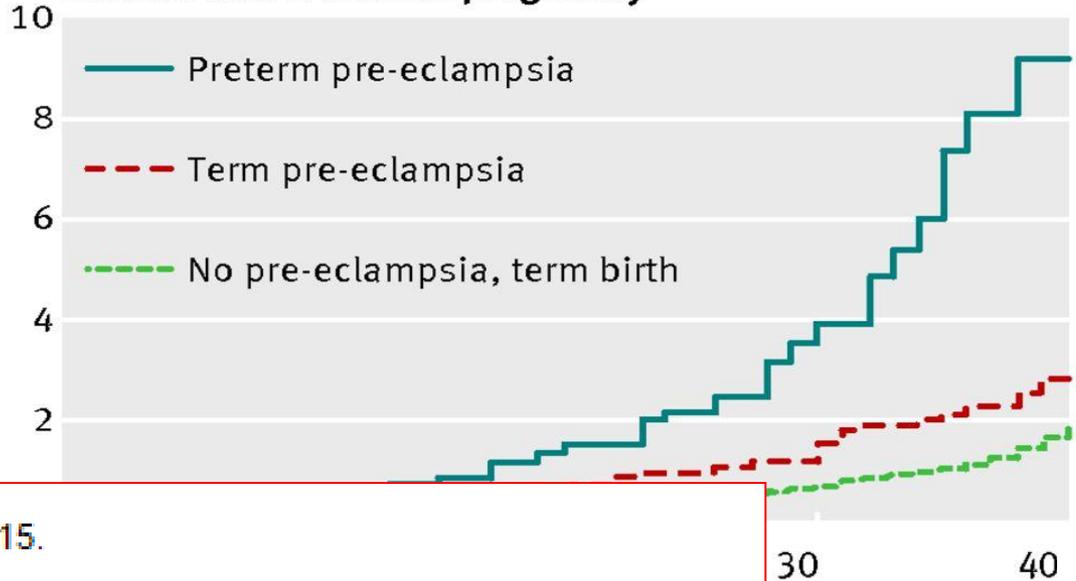
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[Epidemiology](#). 2005 Mar;16(2):206-15.

## Long-term mortality after preeclampsia.

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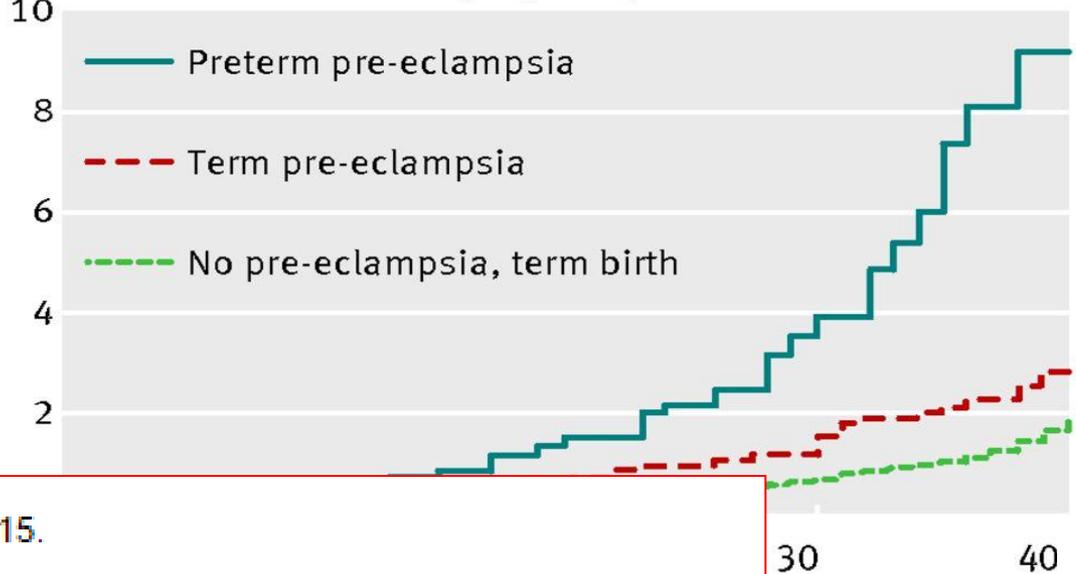
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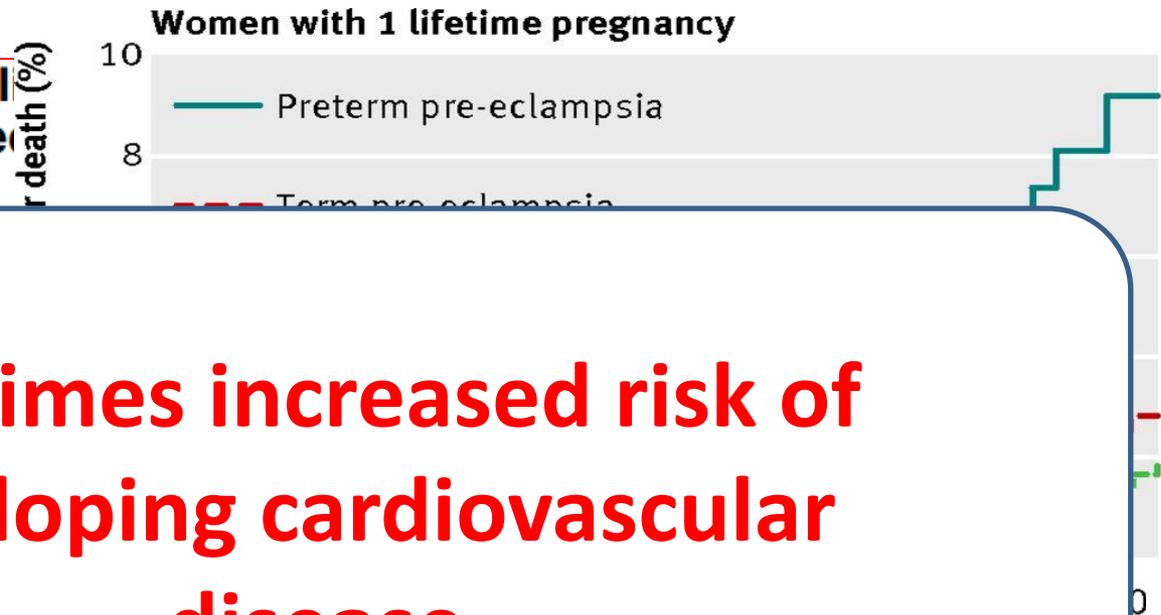
[Lancet](#). 2005 Nov 19;366(9499):1797-803.

## Cardiovascular health after maternal placental syndromes (CHAMPS): population-based retrospective cohort study.

[Ray JG](#)<sup>1</sup>, [Vermeulen MJ](#), [Schull MJ](#), [Redelmeier DA](#).

# Risk of cardiovascular disease

Cardiovascular mortality in child mothers: prospective study



2 - 3 times increased risk of developing cardiovascular disease

Funai EF<sup>1</sup>, Friedlander Y, Paltiel O, Tiram E, Xue X, Deutsch L, Harlap S.

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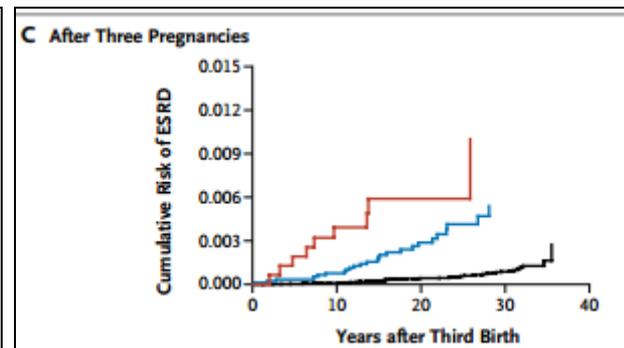
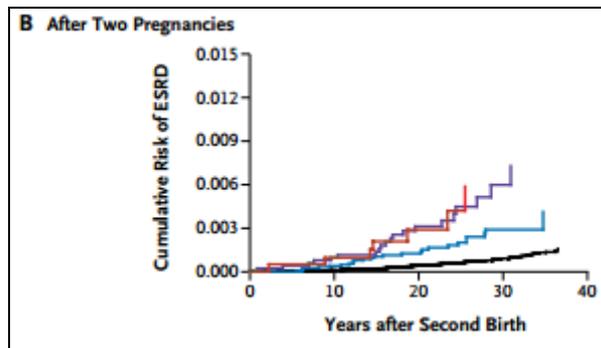
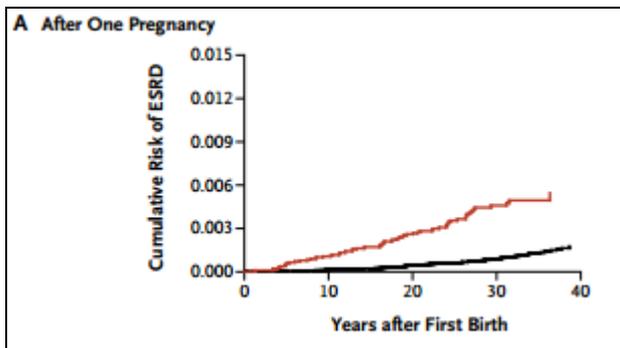
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# Risk of renal disease

N Engl J Med. 2008 Aug 21;359(8):800-9. doi: 10.1056/NEJMoa0706790.

## **Preeclampsia and the risk of end-stage renal disease.**

Vikse BE<sup>1</sup>, Irgens LM, Leivestad T, Skjaerven R, Iversen BM.

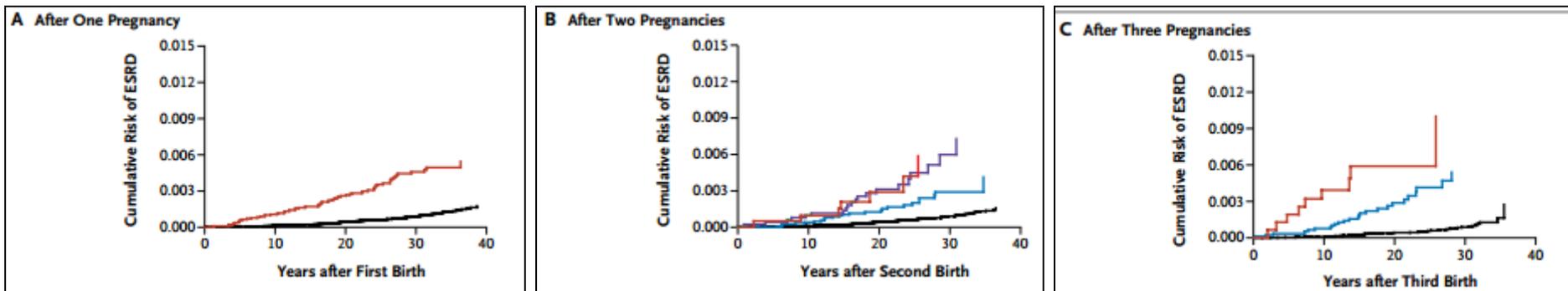


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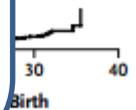
**3 - 5x increased risk of developing chronic renal disease and increased incidence of microalbuminuria**

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A After One P

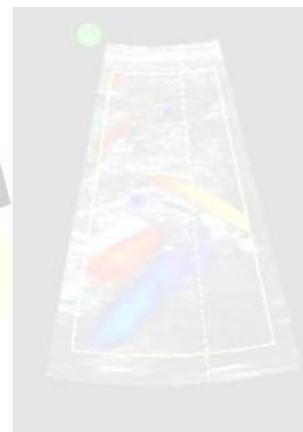
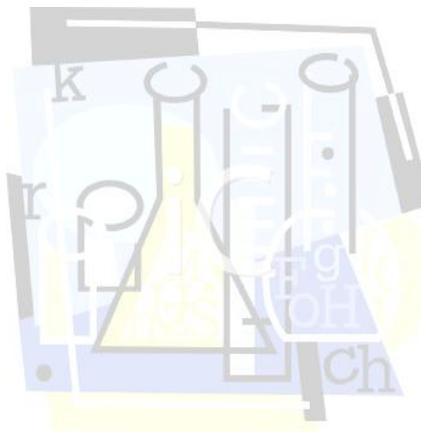


# Epidemiological risk for PE

RF	PE *
Previous PE	8x
BMI over 35, DM	4x
Multipara, twins	3x
Age over 40, HT, renal or autoimmune, interval between pregnancies over 10 years	2x

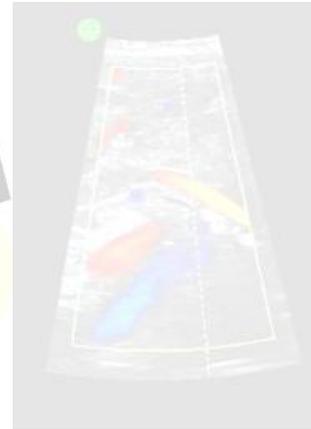
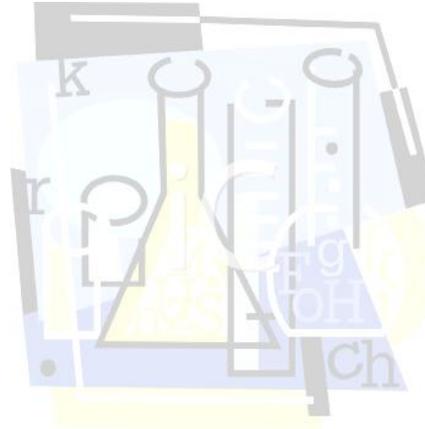
\* Duckitt K. Harington D., Risk factors for preeclampsia at antenatal booking: systematic review of controlled studies. BMJ 2005, 330:565

# Combined screening



METHOD 10 % FPR	11. – 14. week
	Early PE %
?	
?	
?	
?	

# Combined screening

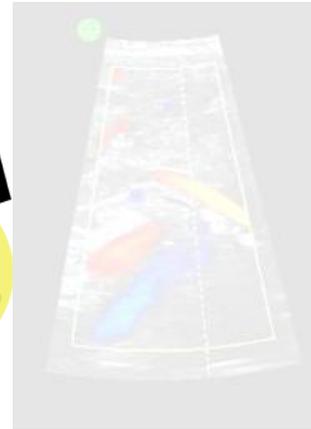
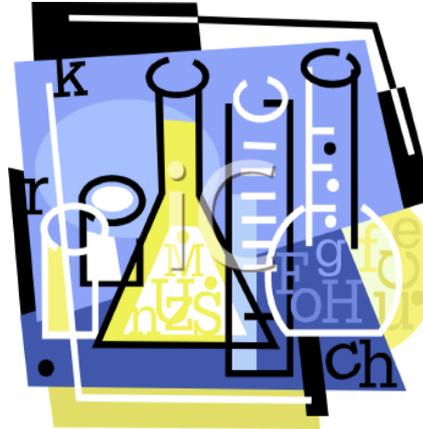


METHOD 10 % FPR	11. – 14. week
	Early PE %
BP*	50
?	
?	
?	

\* Poon LC., Hypertensive disorders in pregnancy: screening by systolic diastolic abd mean arterial pressure at 11-13 week. Hypertens Pregnancy. 2010, 30(1) 93-107

Crovetto F. Performance of first trimester integrated screening for early and late small for gestational age newborns F. IJOG. Accepted 2013

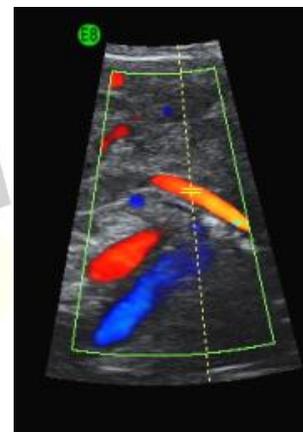
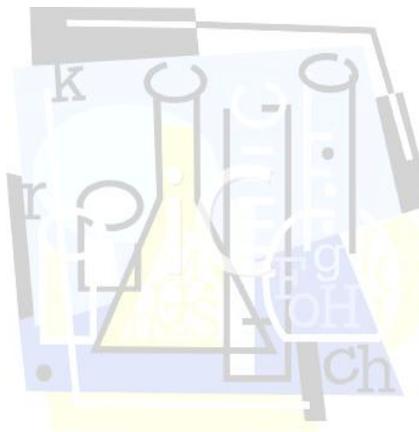
# Combined screening



METHOD 10 % FPR	11. – 14. week
	Early PE %
BP	50
PLGF *	50
?	
?	

\* Chaletz H.2007, Wortelboer EJ. BJOG 2010, Schnauer FI.Placenta 2012, Wortelboer EJ. BJOG 2010, Poon LCY.et al.Prenat.Diag.2008

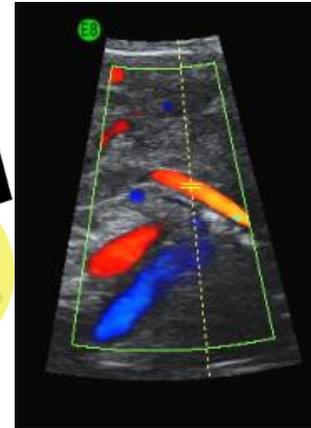
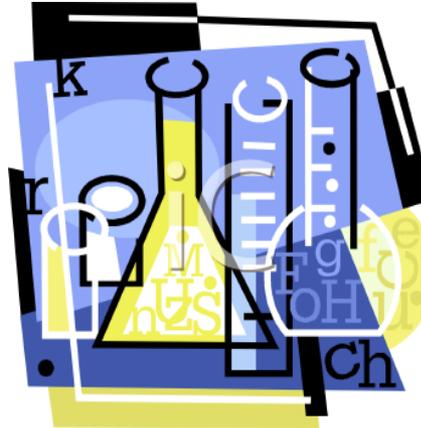
# Combined screening



METHOD 10 % FPR	11. – 14. week
	Early PE %
BP	50
PLGF	50
Doppler UtA *	25
?	

\* Van den Elzen 95, Martin 01, Vainio 05, Ebrashy 05, Gómez 05, Parra 05, Piasencia 07

# Combined screening



METHOD 10 % FPR	11. – 14. week
	Early PE %
BP	50
PLGF	50
Doppler UtA	25
<b>BP, PLGF, UtA *</b>	<b>80</b>

\* Yu CK. Am.J.Obstet.Gynecol.2006, Poon LCY Prenat.Diag.2008, Poon LC.Hypertension 2009, Khall A.IJOG 2010, Poon LC.,J.Horn Hypertensis 2010, Youseff A. Prenat.Diagn.2011, D.Lorenzo. Placenta 2012, Karagiannis G. Fetal Diag.Therapy 2011, Schazzechio E. AJOG 2012

# Prevention

[BMJ Open](#). 2016; 6(6): e011801.

PMCID: PMC4932292

Published online 2016 Jun 28. doi: [10.1136/bmjopen-2016-011801](https://doi.org/10.1136/bmjopen-2016-011801)

## Study protocol for the randomised controlled trial: combined multimarker screening and randomised patient treatment with ASpirin for evidence-based PREeclampsia prevention (ASPRe)

[Neil O'Gorman](#),<sup>1</sup> [David Wright](#),<sup>2</sup> [Daniel L Rolnik](#),<sup>1</sup> [Kypros H Nicolaides](#),<sup>1</sup> and [Liona C Poon](#)<sup>1</sup>

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**acetylsalicylic acid - 150 mg - reduction of early-PE incidence by 38%**



## **Take home message**

***Hypertension in pregnancy is a serious condition and must be properly diagnosed, classified and treated***

**Preeclampsia is a systemic disease with long-term consequences**

**Preeclampsia can be predicted**

**We have relatively effective prevention**

**Serious forms (HELLP) are rare and awareness of their existence is relatively low**

**Thank you for your  
attention**