



*“Will that X-ray harm my unborn child”:  
A meta-analysis of fetal health effects indicates very low risk to fetus  
following occupational exposure of  
pregnant interventional physician.*

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# Disclosures

- ▶ Authors have no disclosures to declare.

# Educational objectives

- ▶ Comprehensive literature review-collected and combined into one source.
  - Review of fetal risk levels
    - radiation dose (occupational radiation exposure)
    - maternal body mass index (BMI)
    - maternal age
    - hormonal therapy during the pregnancy
  
- ▶ Provide literature-based insight which cardiologists and radiologists can use to guide career decisions.

# Take home message

- ▶ For our interventional cardiologists and radiologists with proper radiation safety practices, fetal radiation dose is predicted to be  $< 4$  mGy per term.
- ▶ Compared with known risk of early childhood cancer from other causes, the predicted risk of cancer associated with estimated fetal radiation dose is very low.

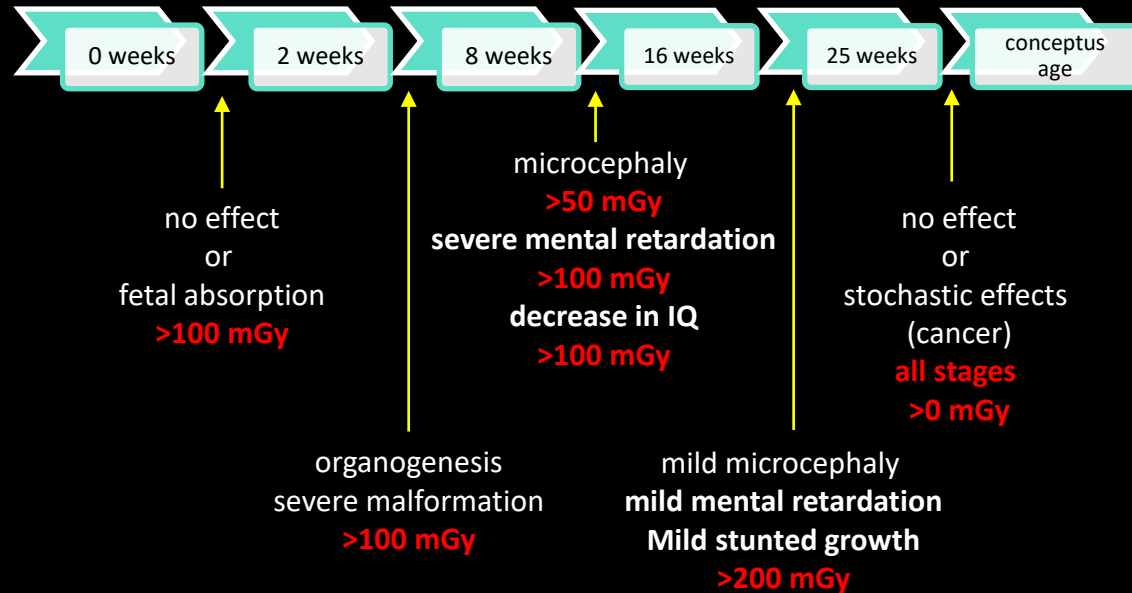
# Radiation dose defined

- ▶ International convention is to express dose to the human fetus in units milli-Gray (mGy), and this convention will be followed herein.
  - Radiation dose to human tissue is assigned units milli-Gray (mGy).
  - Whole body “effective dose” is assigned units milli-Sievert (mSv).
    - *In US regulations, the fetus dose limit is specified in units mrem, where 100 mrem = 1 mSv.*

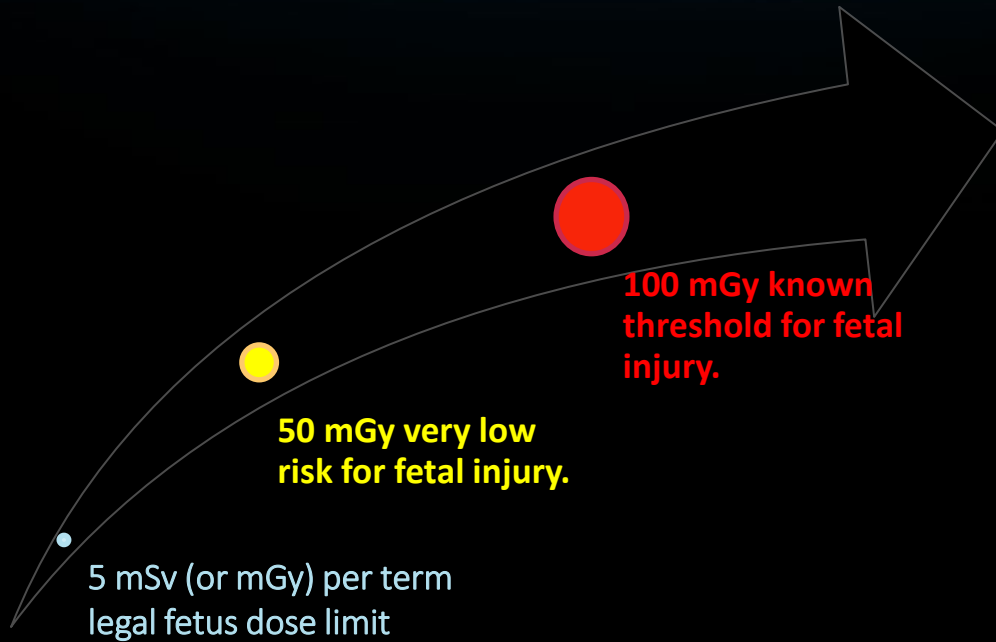
# Deterministic/Stochastic effects

- ▶ Known fetal risks following high fetus radiation dose:
  - prenatal death
  - small head size
  - mental retardation
  - congenital malformation and
  - childhood risk of cancer

Summary of biological effects of prenatal radiation exposure taking in account time point of exposure related to conception as well as the threshold value of risk.



# Putting risks numbers into perspective...



- Exposure to high doses of radiation are known to be detrimental to the health of the fetus.
- Considering the necessary radiation exposure threshold for tissue effect to potentially occur is much higher than what an interventional cardiologists/radiologists would receive.

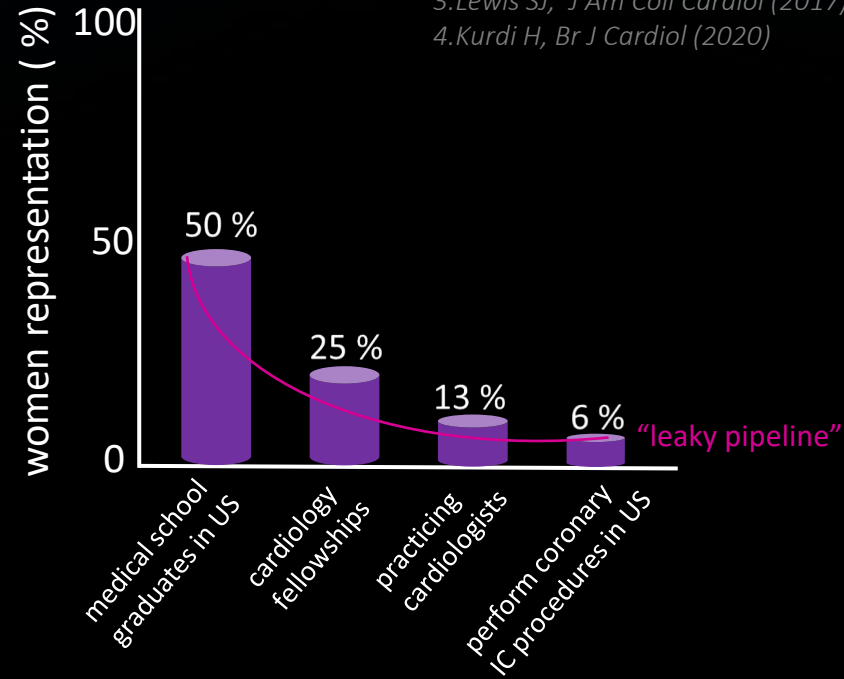
\*ICRP 84 "Pregnancy and Medical Radiation"

\* NCRP 54 "Medical radiation exposure of pregnant and potentially pregnant women"

# Background

- ▶ As a rapidly evolving field interventional cardiology has advanced remarkably since its inception ~50 years ago.
- ▶ Yet, throughout the world, there is a major underrepresentation of women in interventional cardiology (IC).
- ▶ Nonuniformity of the guidelines and often inconclusive data related with fetus health risk continues to be an important factor for women considering careers in IC.

1. Lau ES, *Clin Cardiol* (2018)
2. Douglas PS, *JAMA Cardiol* (2018)
3. Lewis SJ, *J Am Coll Cardiol* (2017)
4. Kurdi H, *Br J Cardiol* (2020)





# Estimating fetus dose in our practice

- ▶ Occupational exposure values (E, mGy) as measured at the left collar, outside the apron for working groups, radiologists and cardiologists, per year.

$$E_{\text{collar, ave.}} = 18.5 \text{ mGy, (N = 191)}$$
$$\text{Range: } 0.5 - 46.1 \text{ mGy}$$

- ▶ Multiply by 2 to estimate exposure at the abdomen, outside the apron.

$$E_{\text{abd., outside}} = 36.9 \text{ mGy}$$

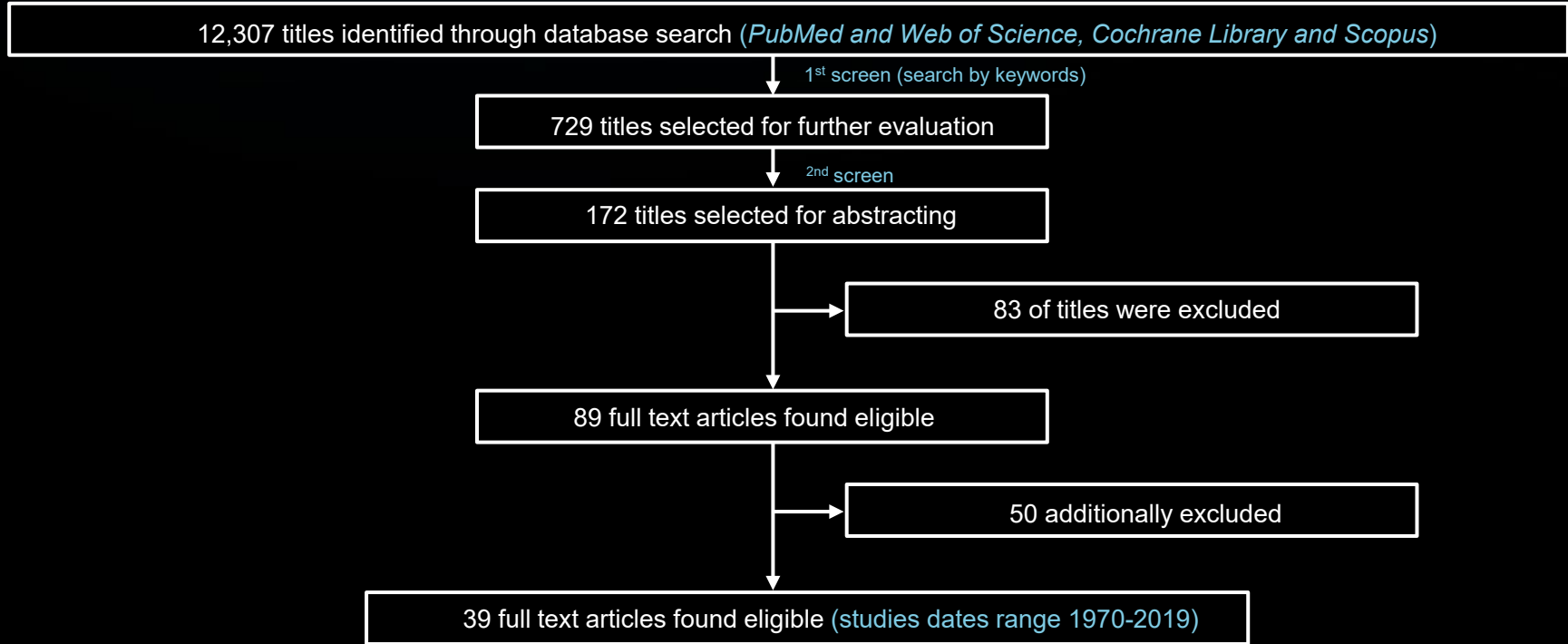
- ▶ Estimate abdomen exposure inside a 0.5 mm Pb apron, assuming 3 % transmission.

$$E_{\text{abd., inside}} = 1.1 \text{ mGy}$$

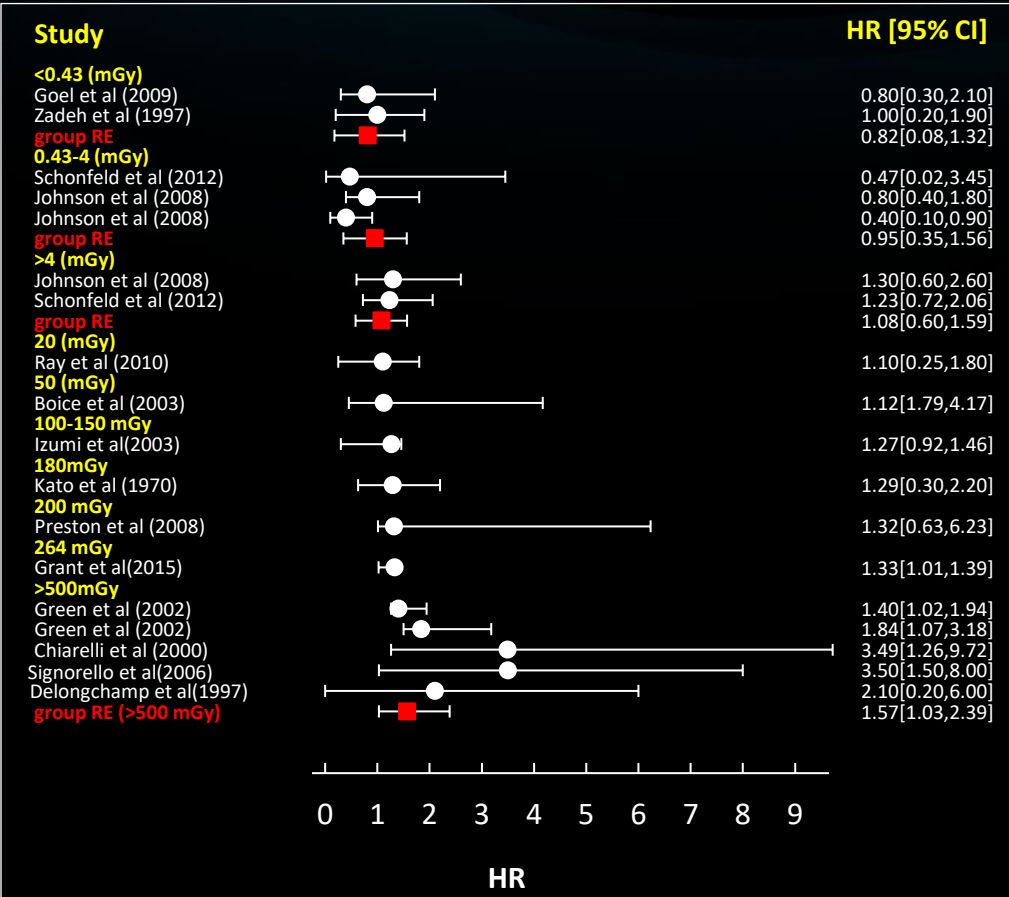
- ▶ Assign exposure under the apron as conservatively high estimate of fetus dose (D).

$$D_{\text{fetus}} \approx 3.9 \text{ mGy (mSv)}$$

# Material and methods

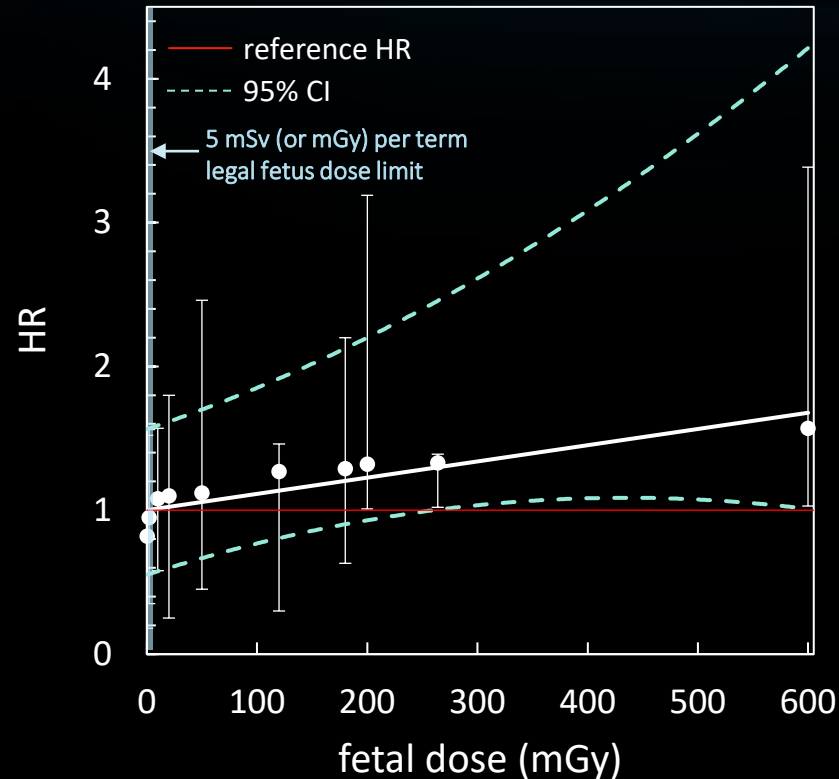


# Results: Meta-analysis of maternal exposure to radiation and early childhood cancer risk



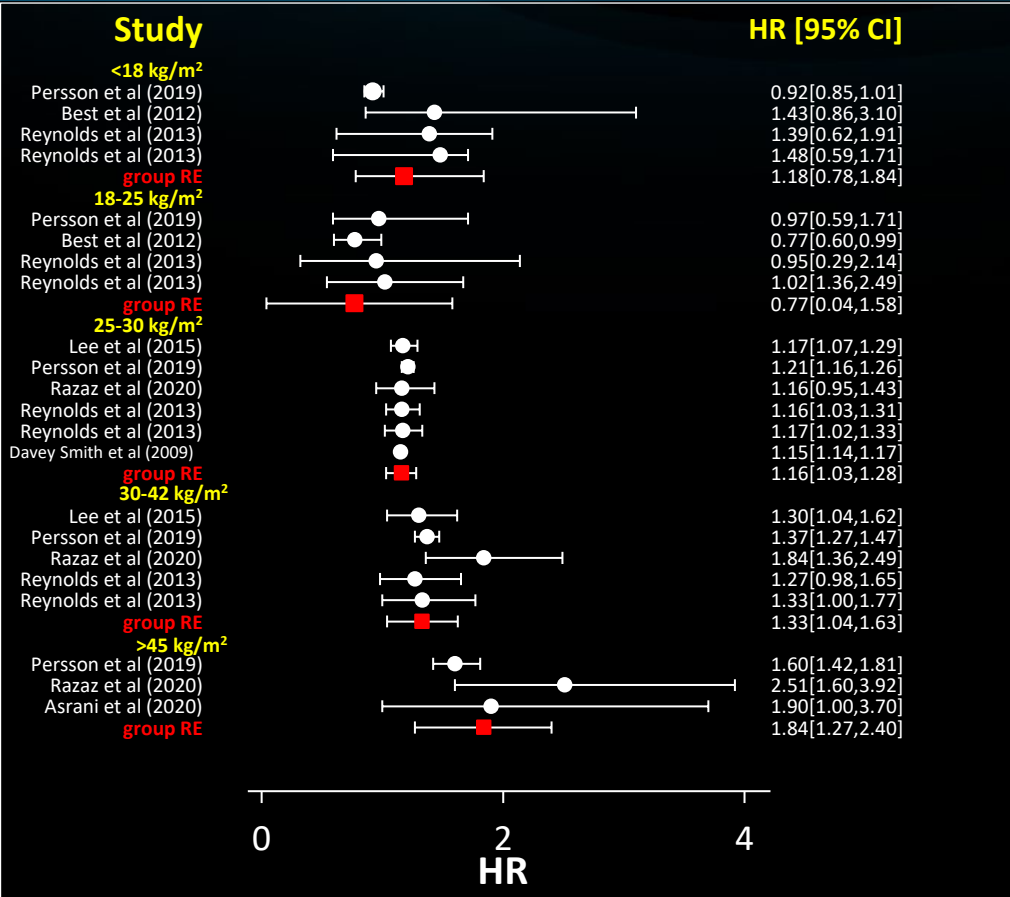
Pooled data using random effect model showed that risk of developing early childhood cancer increase with dose.

## Results: Meta-analysis of maternal exposure to radiation and early childhood cancer risk



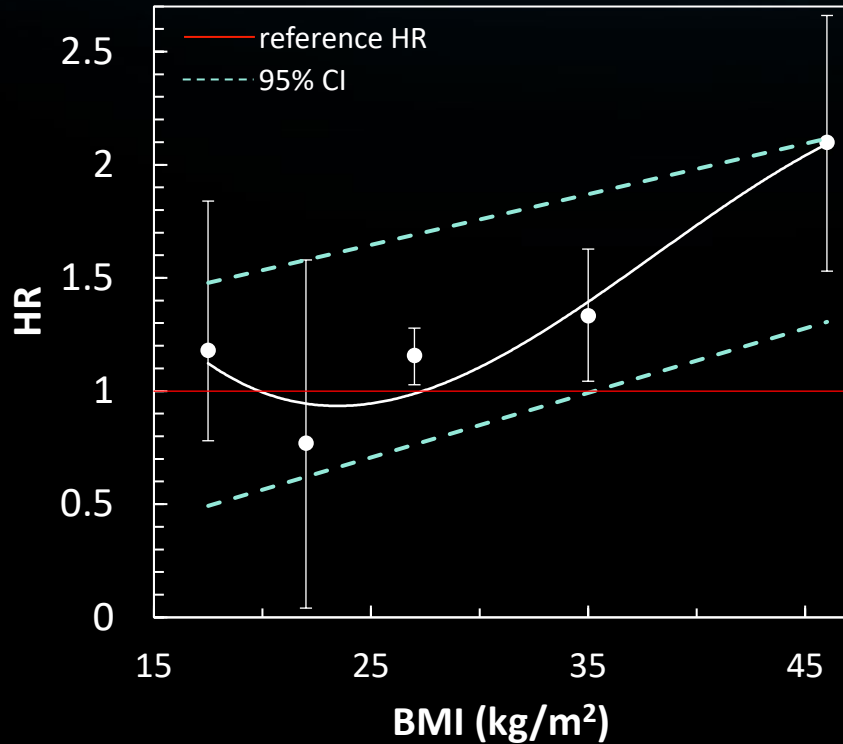
- This meta-analysis suggests that fetus dose greater than ~200 mGy may be associated with increased risk of adverse health effect.
- Other works have suggested that fetus dose >100 mGy may be associated with increased risk.
- Importantly, this meta-analysis demonstrates that HR for fetus dose < 5 mGy is not different than for dose of 0 mGy; HR = 1.001 [95% CI 0.8 to 1.09, p=0.967].

# Results: Meta-analysis of maternal body mass index and risk of congenital heart defects in infants



Meta-analysis among the observational studies showed that maternal BMI is associated with increased risk of CHD in infants.

## Results: Meta-analysis of maternal body mass index and risk of congenital heart defects in infants



Maternal BMI index and hazard risk for CHDs in infants have indicated a positive effect of:

- maternal overweight (BMI >30 kg/m<sup>2</sup>), HR 1.33 [95% CI 1.04, 1.63, p=0.03],
- underweight mothers (BMI <18 kg/m<sup>2</sup>), HR 1.18 [95% CI 0.78, 1.84, p=0.04].

# Results: Meta-analysis of maternal age on preterm birth and low weight newborns

## Study

### under 20 years

Razaz et al (2020)  
 Persson et al (2019)  
 Falster et al (2018)  
 Fuchs et al (2018)  
 Vandekerckhove et al (2021)

### group RE

### 25-35 years

Razaz et al (2020)  
 Persson et al (2019)  
 Falster et al (2018)  
 Fuchs et al (2018)  
 Schummers et al (2018)  
 Vandekerckhove et al (2021)

### group RE

### 35-45 years

Razaz et al (2020)  
 Persson et al (2019)  
 Falster et al (2018)  
 Fuchs et al (2018)  
 Vandekerckhove et al (2021)

### group RE

### over 45 years

Razaz et al (2020)  
 Persson et al (2019)  
 Fuchs et al (2018)  
 Schummers et al (2018)  
 Vandekerckhove et al (2021)

### group RE

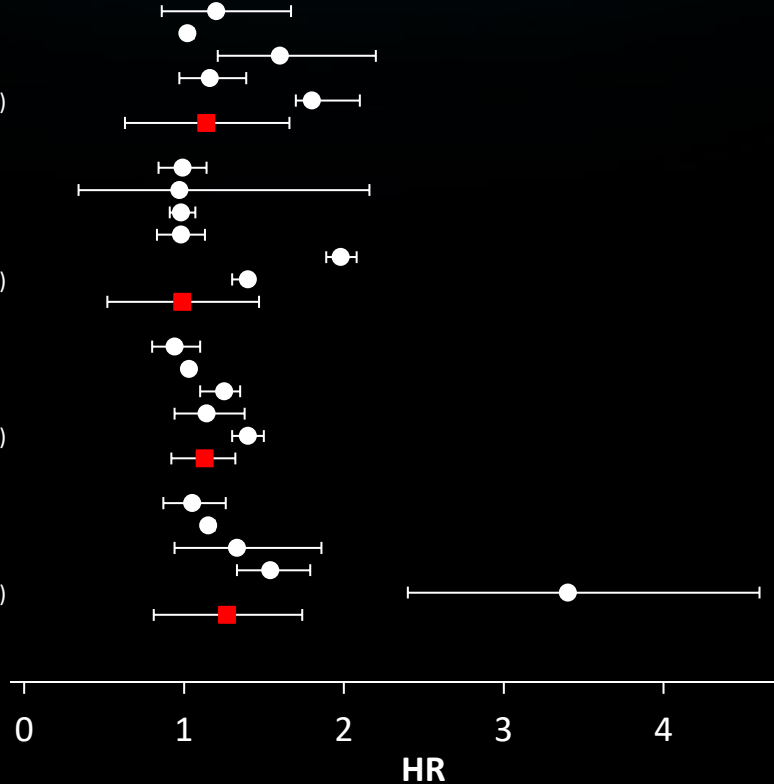
## HR [95% CI]

1.20[0.86,1.67]  
 1.02[0.99,1.06]  
 1.60[1.21,2.20]  
 1.16[0.97,1.39]  
 1.80[1.70,2.10]  
 1.14[0.63,1.66]

0.99[0.83,1.13]  
 0.97[0.29,2.14]  
 0.98[0.85,1.01]  
 0.98[0.83,1.13]  
 1.98[1.89,2.08]  
 1.40[1.30,1.40]  
 0.99[0.52,1.47]

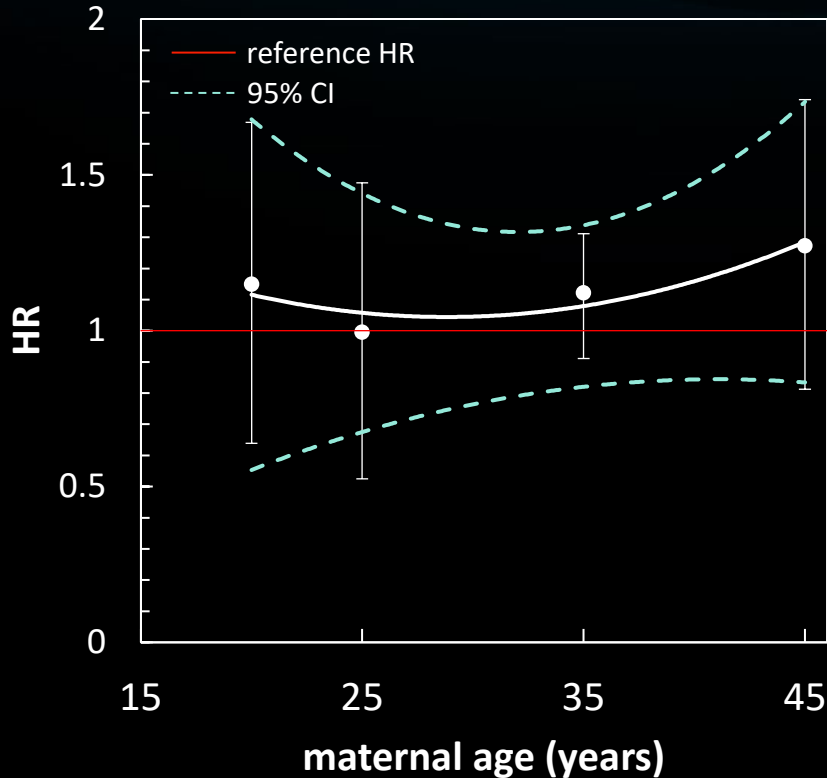
0.94[0.80,1.10]  
 1.03[1.00,1.06]  
 1.25[1.10,1.35]  
 1.14[0.94,1.38]  
 1.40[1.30,1.50]  
 1.13[0.92,1.32]

1.05[0.87,1.26]  
 1.15[1.12,1.19]  
 1.33[0.94,1.86]  
 1.54[1.33,1.79]  
 3.40[2.40,4.60]  
 1.27[0.81,1.74]



Meta-analysis showed age gradient in the probability of giving preterm birth and a low-birth-weight child and was higher at maternal ages older or younger than at the reference category ages (25-35 yrs).

## Results: Meta-analysis of maternal age on preterm birth and low weight newborns

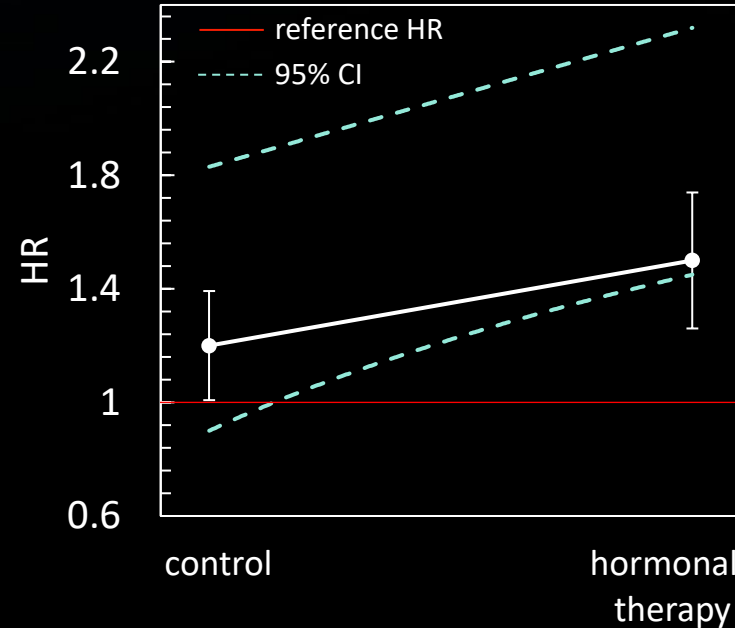
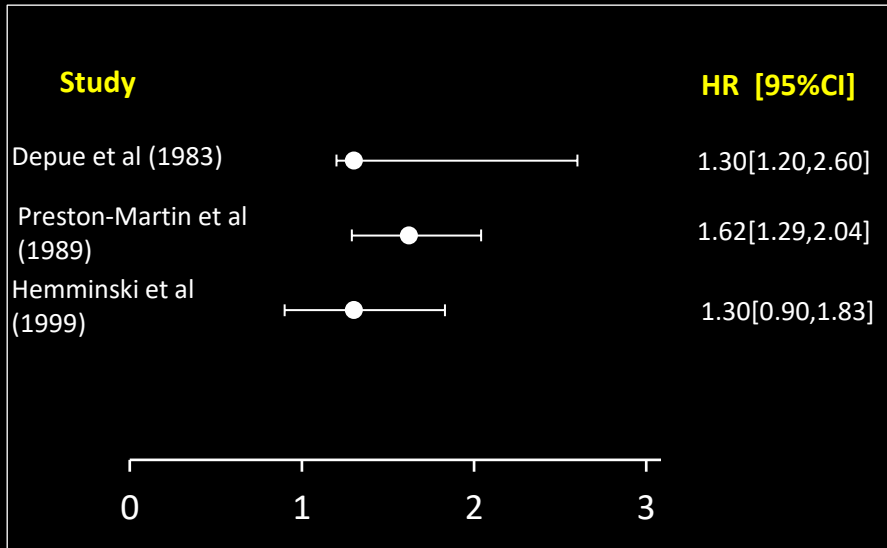


Meta-analysis showed age gradient in the probability of giving preterm birth and a low-birth-weight child and was higher at maternal ages older or younger than at the reference category ages (25-35 yrs).

- (age < 18 yrs HR 1.14 [95%CI 0.63, 1.66, p=0.06])
- (age > 45 yrs HR 1.27 [95% CI 0.81, 1.74, p=0.08])



# Results: Meta-analysis of exposure to female hormone drugs during pregnancy and its effect on malformation in male children



Performed meta-analysis supports hypothesis that oestrogen/progestin drug therapy during pregnancy brings increased risk of malformations in children who were exposed in utero (HR 1.4 [95% CI 0.85, 1.75]. HRs were higher among exposed male children compared to control.

# Summary

1. The meta-analysis demonstrates that HR for fetus dose  $< 5$  mGy is not different than for dose of 0 mGy; HR = 1.001 [95% CI 0.8 to 1.09,  $p=0.967$ ].
2. Maternal BMI  $> 30\text{kg}/\text{m}^2$  was associated with HR 1.33 [95% CI 1.04, 1.63,  $p=0.03$ ], increased risk in CHD in infants.

# Summary

3. Maternal age > 45 years increases risk of preterm birth and underweight newborn with estimated HR 1.27 [95% CI 0.81, 1.74, p=0.08].
4. Hypothesis that oestrogen/progesterone drug therapy during pregnancy brings increased risk of malformation in male children who were exposed *in-utero*, HR 1.4 [95% CI 0.85, 1.75, p=0.08].

# Conclusions

- ▶ This meta-analysis of 14 studies of childhood cancer incidence following in-utero radiation exposure indicates that even dose less than  $\sim 200$  mGy is not associated with adverse health effect. This finding agrees with others that fetus dose less than 100 mGy is unlikely to be associated with adverse health effects.
- ▶ In our practice, dose to the fetus of interventional cardiologists and radiologists is expected to be less than 4 mGy.
- ▶ This work supports the position that radiation risk to the fetus of an interventional physician is exceptionally low.
- ▶ Factors that adversely affect the gestational and early postnatal environment such as maternal BMI, age and some disease treatments can significantly alter fetal development with persistent effects on health.

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