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The impact of HIV infection and ART on the predicted risk of Down's syndrome

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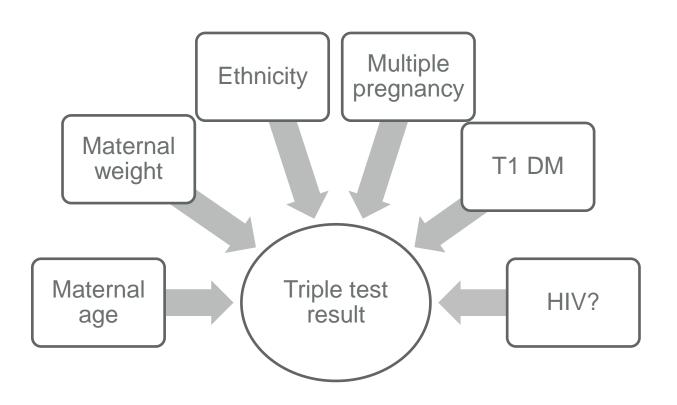
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Background

Down's syndrome screening is routinely offered to all pregnant women in the UK:

- Combined test (10-14 weeks)
- Triple assay (15-20 weeks)
 - √ β- human chorionic gonadotrophin (HCG)
 - \checkmark α– fetoprotein (AFP)
 - ✓ Unconjugated oestriol (UE3)

Background



At the time of this work a "high risk" screening result was >1/250



The story so far

Does HIV infection lead to more high risk DS screening results?

Study	Type (No. Px)	HCG	AFP	UE3	High risk screening vs General population
Neale et al. 2001	Retrospective (76)	-	-	-	↑ (αFP alone, Triple, Quad)
Gross et al. 2003	Retrospective (49)	1	1	\leftrightarrow	_
Yudin <i>et al.</i> 2003	Retrospective (34)	↑	\leftrightarrow	\leftrightarrow	← (Triple) ↑ (Quad)
Spencer 2010	Retrospective (52)	\leftrightarrow	\leftrightarrow	\downarrow	_

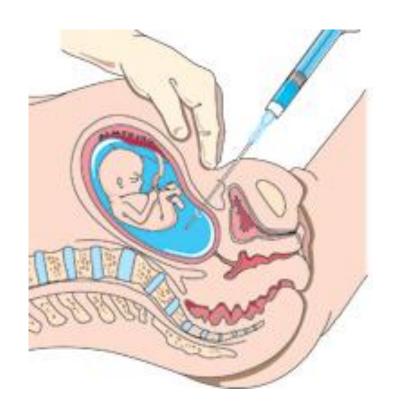


The story so far

Why is this important?

Diagnostic tests (CVS and amniocentesis):

- 0.5-1% foetal loss (Papantoniou et al. 2001)
- HIV transmission (Mandelbrot et al. 1996)
- Anxiety





What we did

Are there more "high risk" screening results in the HIV positive population?

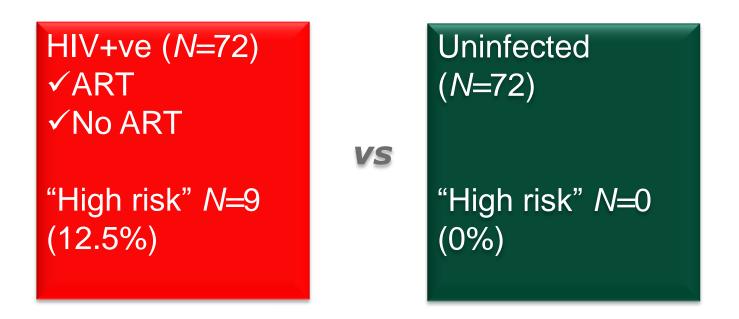
- Retrospective, case-control study
- 72 HIV+ve Vs 72 uninfected controls
- Screening, singleton pregnancy's, 14-18 weeks gestation
- Births took place at St Mary's Hospital, London between January 2002 and July 2009 (Laboratory BWH)

Do these women have an increased risk of a Down's syndrome affected pregnancy?

 National Study of HIV in Pregnancy and Childhood (NSHPC) vs National DS register data

Key findings 1

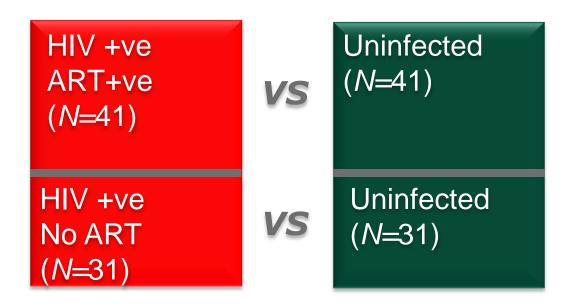
HIV population compared to uninfected controls:



HIV positive women were **twice** as likely to receive "high risk" screening results: OR = 2.14, 95% CI = (1.79 - 2.57) p = 0.002

Key findings 2

HIV population (without ART) compared to uninfected controls:



- Higher mean hCG MoM (1.64 \vee 1.07, p=0.02)
- **Higher risk** (1/909 v 1/33333, *p*=0.03)
- This effect was <u>not seen</u> in patients on ART

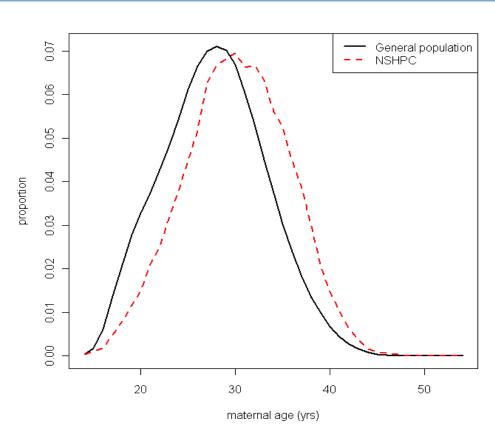
Key findings 3

Population based data:

- Incidence in HIV positive population,
 1/416
- Incidence in general population,1/633

However, the maternal age differed significantly between the two groups.

Corrected risk = 0



Distributions of maternal age at delivery for the general population and among women enrolled in the NSHPC

Summary

- HIV positive women are twice as likely to receive a "high risk" screening result compared to an uninfected population
- A particular problem in patients without ART
- After adjusting for differences in maternal age, population-based evidence does not support a link between HIV serostatus and an increased chance of a DS affected pregnancy

Main take home messages

- 1.If patient presents early enough, offer the combined test ,10-14 weeks (Brossard *et al.* 2008)
- 2. Preconceptual counselling
- 3. Know HIV status before interpreting a screening result and referring for invasive diagnostic tests



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