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Regimens of fetal surveillance for impaired fetal growth

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Abstract

Background—Policies and protocols for fetal surveillance in the pregnancy where impaired fetal growth is suspected vary widely, with numerous combinations of different surveillance methods.

Objectives—To assess the effects of antenatal fetal surveillance regimens on important perinatal and maternal outcomes.

Search methods—We searched the Cochrane Pregnancy and Childbirth Group's Trials Register (April 2008).

Selection criteria—Randomised and quasi-randomised trials comparing the effects of described antenatal fetal surveillance regimens.

Data collection and analysis—R Grivell and L Wong independently assessed trial eligibility and quality and extracted data.

Main results—One trial of 167 women and their babies was included. This trial was a pilot study recruiting alongside another study, therefore a separate sample size was not calculated. The trial compared a twice-weekly surveillance regimen (biophysical profile, nonstress tests, umbilical artery and middle cerebral artery Doppler and uterine artery Doppler) with the same regimen applied fortnightly (both groups had growth assessed fortnightly). There were insufficient data to assess this review's primary infant outcome of composite perinatal mortality and serious morbidity (although there were no perinatal deaths) and no difference was seen in the primary maternal outcome of emergency caesarean section for fetal distress. In keeping with the more frequent monitoring, mean gestational age at birth was four days less for the twice-weekly surveillance group compared with the fortnightly surveillance group. Women in the twice-weekly

DECLARATIONS OF INTEREST None known.

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surveillance group were 25% more likely to have induction of labour than those in the fortnightly surveillance group. The risk ratio was 1.25 (95% confidence interval 1.04 to 1.50).

Authors' conclusions—There is limited evidence from randomised controlled trials to inform best practice for fetal surveillance. regimens when caring for women with pregnancies affected by impaired fetal growth. More studies are needed to evaluate the effects of currently used fetal surveillance regimens in impaired fetal growth

Medical Subject Headings (MeSH)

Fetal Development; Fetal Growth Retardation [*diagnosis]; Fetal Monitoring [*methods]; Middle Cerebral Artery [ultrasonography]; Randomized Controlled Trials as Topic; Ultrasonography, Prenatal; Umbilical Arteries [ultrasonography]

MeSH check words

Female; Humans; Pregnancy

BACKGROUND

Description of the condition

Impaired fetal growth is a disturbance of normal growth, resulting in a baby that does not grow to its potential. Impaired growth may be due to maternal complications (such as high blood pressure or hypertension), fetal complications (including structural or chromosomal problems) or problems with placental development (which is the most common cause of impaired growth) (Baschat 2006). It can be difficult to accurately differentiate between the constitutionally small baby and one with impaired growth when compared with its potential.

There are numerous definitions for growth of the baby in pregnancy. An infant with low birthweight for gestation is classified as very small-for-gestational age (SGA) (less than 3rd percentile) or SGA (less than 10th percentile). SGA includes both those babies that have not reached their growth potential (through growth impairment) and also those babies that are constitutionally small. A baby may be of normal birthweight but still significantly lighter than its genetic growth potential (Baschat 2006).

When impaired fetal growth is suspected in pregnancy, ultrasound may be ordered to assess growth with various standard measurements. These include the abdominal circumference, biparietal diameter, head circumference and femur (upper leg) length, which can then be used to estimate the fetal weight.

Ultrasound allows these measurements to be plotted against known population percentile lines on a growth curve to assess severity and timing of growth restriction. Sequential ultrasounds can also allow identification of the baby who begins pregnancy on a certain percentile and then crosses the percentile lines because of impaired growth.

Impaired growth is associated with increased risk of complications for the baby, including perinatal death (Baschat 2001; Gardosi 1998), adverse neurological outcomes (such as cerebral palsy) and poor long-term neurodevelopmental outcomes (Yanney 2004). There

may also be an association with adverse health events in adult life (such as obesity, type 2 diabetes and high blood pressure). Optimal management of growth restriction has three main aspects: identification of the baby at risk in the general population (pregnancy risk factors may be of assistance); confirmation of the diagnosis of impaired fetal growth and distinction from the healthy small baby; and ongoing care, culminating in decisions about mode and timing of birth (Baschat 2005; Breeze 2007).

Description of the intervention

After a baby has been identified with impaired growth in-utero, antenatal care providers may adopt a variety of methods to monitor the well-being of the baby. The choice of monitoring technique and the frequency of monitoring varies considerably, but may be influenced by the severity of the growth impairment.

Antenatal surveillance aims to provide information about the health of the baby while in the uterus. Decisions based on the results of that surveillance may in turn reduce the risks to both the baby and the mother associated with impaired growth.

There are a number of tests that may be used in clinical practice, alone or in combination, to identify the baby at risk of growth restriction. These include clinical estimation of the baby's size by palpating the woman's abdomen, ultrasound assessment of baby's growth, fetal movement counting, fetal heart rate assessment (cardiotocography), ultrasound assessment of the baby's well-being by biophysical profile testing, and assessment of the blood flow through the placenta and various blood vessels in the baby (called Doppler velocimetry). The use of some of these tests alone are the subject of other Cochrane reviews (Alfirevic 1996; Lalor 2008; Pattison 1999). The scope of this review is to examine these surveillance methods when used together in different combinations or at different intervals.

How the intervention might work

By increasing the frequency of monitoring a woman's baby during pregnancy, or using different methods of surveillance in combination, it may be possible to identify the baby who would benefit from earlier birth. This in turn, may be associated with improved health outcomes for the baby.

Why it is important to do this review

Once a baby has been identified as having impaired growth, the appropriate methods and frequency of surveillance is uncertain. Currently, there is considerable variation, based on local hospital policies, and individual practitioner preferences. This systematic review will assess the benefits and harms associated with different combinations of surveillance methods.

OBJECTIVES

To assess the benefits and harms associated with different methods of antenatal surveillance for the baby identified with impaired growth.

METHODS

Criteria for considering studies for this review

Types of studies—Published and unpublished randomised and quasi-randomised trials evaluating the effects of one or more described antenatal fetal surveillance regimens. Studies which compare a particular regimen with 'standard care' were only included if standard care involved what this review has defined as a regimen. We did not include studies that compare to no specific surveillance.

Types of participants—Women with pregnancies deemed by the investigators to be affected by impaired fetal growth or fetal growth restriction. (The minimum requirement for this review is estimated fetal weight of less than 10th percentile or abdominal circumference less than 10th percentile, using local intrauterine growth charts.)

Types of interventions—Described antenatal fetal surveillance regimens. A regimen for the purpose of the review was two or more surveillance methods. Each surveillance method was performed at a specified interval. Possible surveillance methods to be included: fetal movement monitoring, cardiotocography, biophysical profile, Doppler ultrasound of the umbilical artery and fetal venous and arterial circulation (ductus venosus or middle cerebral artery). For the purpose of the review, we expected a study to randomise women to different regimens of surveillance (i.e. different combinations of surveillance methods or applying surveillance methods at different intervals).

Types of outcome measures—Clinically relevant outcomes were prespecified through discussion and agreement among the review authors.

Primary outcomes

Infant:

Perinatal mortality and serious neonatal morbidity, as defined by the trial authors

Maternal:

Emergency caesarean section for fetal distress

Secondary outcomes

Infant:

- Stillbirth
- Neonatal death
- Hypoxic ischaemic encephalopathy
- Admission to neonatal special care and or intensive care unit
- Gestational age at birth
- Low birthweight for gestation at birth (as defined by individual trial)

- Birthweight
- Preterm birth
- Low Apgar scores at five minutes (as defined by trial)
- Cord blood acidosis
- Metabolic acidosis or lactic acidosis
- Respiratory distress syndrome
- Use of ventilatory support
- Use of inotropic support
- Necrotising enterocolitis
- Grade III or IV intraventricular haemorrhage
- Duration of neonatal hospital stay
- Longer-term neurodevelopmental outcomes
- Cerebral palsy

Maternal:

- Antenatal hospital admission
- Induction of labour
- Caesarean section
- Operative vaginal delivery
- Womens views of care

Use of healthcare services:

- For infant: length of stay in neonatal special care and or intensive care unit
- For mother: length of antenatal hospital admission
- Costs of monitoring

Search methods for identification of studies

Electronic searches—We searched the Cochrane Pregnancy and Childbirth Group's Trials Register by contacting the Trials Search Co-ordinator (April 2008).

The Cochrane Pregnancy and Childbirth Group's Trials Register is maintained by the Trials Search Co-ordinator and contains trials identified from:

- quarterly searches of the Cochrane Central Register of Controlled Trials (CENTRAL);
- 2. weekly searches of MEDLINE;

- 3. handsearches of 30 journals and the proceedings of major conferences;
- **4.** weekly current awareness alerts for a further 44 journals plus monthly BioMed Central email alerts.

Details of the search strategies for CENTRAL and MEDLINE, the list of handsearched journals and conference proceedings, and the list of journals reviewed via the current awareness service can be found in the 'Specialized Register' section within the editorial information about the Cochrane Pregnancy and Childbirth Group.

Trials identified through the searching activities described above are each assigned to a review topic (or topics). The Trials Search Co-ordinator searches the register for each review using the topic list rather than keywords.

We did not apply any language restrictions.

Data collection and analysis

Selection of studies—We assessed for inclusion all potential studies we identified as a result of the search strategy. We resolved any disagreement through discussion or, if required, consulted a third party.

Data extraction and management—We designed a form to extract data. Two review authors (R Grivell and L Wong) extracted the data independently using the agreed form. We resolved discrepancies through discussion. We used the Review Manager software (RevMan 2008) to double enter all the data or a subsample.

When information regarding any of the above was unclear, we attempted to contact authors of the original reports to provide further details.

Assessment of methodological quality of included studies—We assessed the validity of each study using the criteria outlined in the Cochrane Handbook for Systematic Reviews of Interventions (Higgins 2008). Methods used for generation of the randomisation sequence were described for each trial.

(1) Sequence generation (checking for possible selection bias): We described for each included study the method used to generate the allocation sequence in sufficient detail to allow an assessment of whether it should produce comparable groups. We assessed the method as:

- adequate (any truly random process, e.g. random number table; computer random number generator);
- inadequate (any non random process, e.g. odd or even date of birth; hospital or clinic record number);
- unclear.

(2) Allocation concealment (checking for possible selection bias): We described for each included study the method used to conceal the allocation sequence in sufficient detail and

determine whether intervention allocation could have been foreseen in advance of, or during recruitment, or changed after assignment. We assessed the methods as:

- adequate (e.g. telephone or central randomisation; consecutively numbered sealed opaque envelopes);
- inadequate (open random allocation; unsealed or non-opaque envelopes, alternation; date of birth);
- unclear.

(3) Blinding (checking for possible performance bias): We described for each included study the methods used, if any, to blind study participants and personnel from knowledge of which intervention a participant received. Studies were judged at low risk of bias if they were blinded, or if we judged that the lack of blinding could not have affected the results. Blinding was assessed separately for different outcomes or classes of outcomes. We assessed the blinding as: adequate, inadequate or unclear for participants; adequate, inadequate or unclear for outcome assessors.

(4) Incomplete outcome data (checking for possible attrition bias through withdrawals, dropouts, protocol deviations): We described for each included study, and for each outcome or class of outcomes, the completeness of data including attrition and exclusions from the analysis. We state whether attrition and exclusions were reported, the numbers included in the analysis at each stage (compared with the total randomised participants), reasons for attrition or exclusion where reported, and whether missing data were balanced across groups or were related to outcomes. Where sufficient information was reported, or can be supplied by the trial authors, we re-included missing data in the analyses which we undertake. We will assess methods as:

- adequate (i.e. low risk of bias);
- inadequate (i.e. high risk of bias):
- unclear.

(5) Selective reporting bias: We described for each included study how we investigated the possibility of selective outcome reporting bias and what we found. We assessed the methods as:

- adequate (where it is clear that all of the study's prespecified outcomes and all expected outcomes of interest to the review have been reported);
- inadequate (where not all the study's prespecified outcomes have been reported; one or more reported primary outcomes were not prespecified; outcomes of interest are reported incompletely and so cannot be used; study fails to include results of a key outcome that would have been expected to have been reported);
- unclear.

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(6) Other sources of bias: We described for each included study any important concerns we have about other possible sources of bias. We assessed whether each study was free of other problems that could put it at risk of bias:

- yes;
- no;
- unclear.

(7) Overall risk of bias: We made explicit judgements about whether studies are at high risk of bias, according to the criteria given in the Handbook (Higgins 2008). With reference to (1) to (6) above, we assessed the likely magnitude and direction of the bias and whether we considered it likely to impact on the findings. We will explore the impact of the level of bias through undertaking sensitivity analyses - *see* 'Sensitivity analysis'.

Measures of treatment effect—We carried out statistical analysis using RevMan 2008. We used fixed-effect meta-analysis for combining data in the absence of significant heterogeneity if trials were sufficiently similar. If heterogeneity was found, we explored this by sensitivity analysis followed by random-effects if required.

Dichotomous data: For dichotomous data, we presented results as summary risk ratio with 95% confidence intervals.

<u>Continuous data:</u> For continuous data, we used the mean difference if outcomes were measured in the same way between trials. If there was evidence of skewness, this was reported.

Dealing with missing data—We analysed data on all participants with available data in the group to which they were allocated, regardless of whether or not they received the allocated intervention. If, in the original reports, participants were not analysed in the group to which they were randomised, and there was sufficient information in the trial report, we attempted to restore them to the correct group.

Assessment of heterogeneity—We were unable to apply tests of heterogeneity between trials using the I^2 statistic because only one trial was included in the reveiw.

Subgroup analyses—We planned to conduct subgroup analyses classifying whole trials by interaction tests as described by Deeks 2001. We planned to carry out the following subgroup analyses: diagnosis of impaired fetal growth before or after 32 weeks, normal or abnormal umbilical artery doppler at time of diagnosis. These analyses were not possible on the information from this study.

Sensitivity analyses—We planned sensitivity analysis to explore the effect of trial quality. This involved analysis based on an a rating of selection bias and attrition bias. We planned to exclude studies of poor quality in the analysis in order to assess for any substantive difference to the overall result. Sensitivity analysis was not possible as there was only one trial included.

RESULTS

Description of studies

See: Characteristics of included studies; Characteristics of excluded studies.

Three trials were identified and one trial met our inclusion criteria (McCowan 2000) - see table of Characteristics of included studies. Two trials were excluded as they did not compare antenatal fetal surveillance regimens (i.e. combination of methods of surveillance, each at a specified frequency). Williams 2003 randomised women to either nonstress testing or umbilical artery velocimetry, while Sood 2007 looked at the addition of vibroacoustic stimulation to biophysical profile testing.

Participants—McCowan 2000 was a pilot study and recruitment occurred concurrently with another randomised trial assessing the use of low-dose aspirin treatment for small-for-gestational-age fetuses with abnormal Doppler studies. The pilot trial McCowan 2000 recruited pregnant women who were outpatients with a singleton pregnancy between 24 and 36 weeks' gestation, with ultrasonographic evidence of small-for-gestational-age fetuses (i.e. abdominal circumference < 10th percentile), normal results of umbilical artery Doppler velocimetry studies and no oligohydramnios. A total of 167 women were randomly assigned, 85 to the twice-weekly surveillance group and 82 to the fortnightly surveillance group.

Settings—The included study was conducted in Auckland, New Zealand at a tertiary referral center (acute hospital setting).

Interventions—Twice-weekly or fortnightly fetal surveillance involving biophysical profile, nonstress tests, umbilical artery and middle cerebral artery Doppler, growth scans and uterine artery Doppler.

Risk of bias in included studies

Formal randomisation was reported in McCowan 2000 which used sequence generation via computer-generated numbers and allocation concealment via telephone randomisation. It was not feasible to blind the participant and care giver and it was not stated whether the outcome assessors were blinded or not. No loss to follow up was documented.

Effects of interventions

Primary outcomes

Infant:

• Perinatal mortality and serious neonatal morbidity, as defined by the trial authors

The authors state that there were no perinatal deaths in either study group. McCowan 2000 did not assess composite perinatal morbidity and mortality as a primary outcome but instead used markers of neonatal morbidity. Perinatal mortality is mentioned in the body of the text, this information has been used for this review's outcome of perinatal mortality.

Maternal:

• Emergency caesarean section for fetal distress

No statistically significant difference was seen for this primary maternal outcome (risk ratio (RR) 0.96; 95% confidence interval (CI) 0.35 to 2.63; 167 women) (Analysis 1.22).

Secondary outcomes

Infant: Data were available for several of the secondary outcomes that relate to infant health. In keeping with the more frequent monitoring, mean gestational age at birth was four days less for the twice-weekly surveillance group compared with the fortnightly surveillance group, mean difference - 4 days (95% CI –7.79 to -0.21) (Analysis 1.7) No statistically significant differences were seen for any of the other infant secondary outcomes where data were available, for admission to neonatal special care unit or intensive care unit (Analysis 1.6); low birthweight (Analysis 1.8); mean birthweight (Analysis 1.9); and cord blood acidosis (Analysis 1.12).

No data were available for some of the serious neonatal morbidity and mortality outcomes including preterm birth, low Apgar scores at five minutes (as defined by trial), and other complications of prematurity, such as respiratory morbidity, necrotising enterocolitis and serious intraventricular haemorrhage. Only two infants required mechanical ventilation (at 30 and 32 weeks), both babies were in the twice-weekly group (Analysis 1.15). No follow up was performed. Consequently, no data are available on longer-term neurodevelopmental outcomes and cerebral palsy.

Maternal: Women in the twice-weekly surveillance group were 25% more likely to have labour induced than those in the fortnightly surveillance group (RR 1.25; 95% CI 1.04 to 1.50) (Analysis 1.24).

No statistically significant differences were seen between the two groups regarding caesarean section (Analysis 1.25). No data were available on women's views of care, length of antenatal hospital admission and operative vaginal birth.

Use of healthcare services—No statistically significant difference was seen for length of stay in neonatal special care and/or intensive care unit. No data were available for length of antenatal hospital admission or costs of monitoring.

Outcomes reported but not prespecified in the review—McCowan 2000 reported no statistically significant difference in umbilical artery resistance index at delivery, Ponderal index, hypoglycaemia, spontaneous onset of labour, pre-eclampsia and gestational hypertension.

DISCUSSION

This pilot, randomised trial of two regimens did not demonstrate any difference in markers of neonatal morbidity or perinatal mortality between the two study groups. A much larger

study will be required to detect a difference in perinatal mortality and serious neonatal morbidity.

The women in the twice-weekly surveillance group were 25% more likely to have induction of labour compared to those in the fortnightly surveillance group where spontaneous onset of labour was more likely to occur. Despite the earlier intervention, there were no statistically significant benefits in infant or maternal outcomes such as neonatal morbidity, the need for caesarean section or the need for emergency caesarean section for fetal distress.

Women in the twice-weekly surveillance group gave birth on average four days earlier than women in the fortnightly surveillance group. This difference of four days is unlikely to be clinically significant in infants born at term and is unlikely to impact on serious neonatal morbidity or mortality.

There is almost no evidence to date to indicate an optimal antenatal surveillance method for infants identified with impaired growth. Any future studies should include an assessment of women's views of care and should be adequately powered to detect clinically relevant differences in maternal and infant health outcomes.

This study only recruited a subset population of all pregnancies with small-for-gestationalage fetus (that is, pregnancies with normal Doppler velocimetric studies); it excluded pregnancies with abnormal Doppler studies and oligohydramnios. Often babies who are identified as having impared in-utero growth also have abnormal Doppler studies or oligohydramnios (low amniotic fluid volume), or both. There is no evidence to inform the optimal surveillance regimen for this group of women and their babies. Thus, further studies could incorporate subgroup analysis of pregnancies with small-for-gestational-age fetuses associated with abnormal Doppler studies or oligohydramnios, or both.

One international group of authors has embarked on a randomised trial of timing of delivery in early preterm fetal growth restriction based on early and late fetal Doppler venous changes versus cardiotocography. It is noted by the Trial of Umbilical and Fetal Flow in Europe (TRUFFLE) group that current evidence regarding assessment techniques is lacking and that there is a need for a multicenter randomised controlled trial that will help to provide reliable evidence (Lees 2005). It is hoped that the TRUFFLE study will contribute at least in part to our knowledge surrounding the complex question of what fetal surveillance methods to use, how to use them and how the results should be interpreted and acted upon.

AUTHORS' CONCLUSIONS

Implications for practice

There is very little evidence from randomised controlled trials to inform best practice for antenatal surveillance regimens when caring for women whose pregnancy is affected by impaired fetal growth.

Implications for research

More studies are needed to evaluate the effects of current fetal surveillance regimens for impaired fetal growth.

Acknowledgments

The authors would like to acknowledge Philippa Middleton for her advice during the protocol and review phase.

As part of the prepublication editorial process, this review has been commented on by two peers (an editor and referee who is external to the editorial team), a member of the Pregnancy and Childbirth Group's international panel of consumers and the Group's Statistical Adviser.

CHARACTERISTICS OF STUDIES

Characteristics of included studies [ordered by study ID]

McCowan 2000

Methods	Randomised controlled trial, p	bilot study recruiting alongside another larger trial				
Participants	167 women with singleton pregnancies 24 to 36 weeks; outpatients; previous normal anatomy scan; ultrasound showing abdominal circumference < 10th percentile, 2 measures of umbilical artery Doppler resistive index < 95th centil oligohydramnios					
Interventions	Twice-weekly versus fortnightly fetal surveillance. Surveillance methods inclu biophysical profile, nonstress tests, umbilical artery and middle cerebral artery Doppler, growth scans and uterine artery Doppler					
Outcomes	Neonatal; gestational age at delivery, umbilical artery resistive index at delivery, abnormal umbilical artery at delivery, female sex, birthweight, birthweight < 10th percentile, ponderal index, ponderal index < 10th percentile, admission to neonata nursery, neonatal hospital stay, acidosis at birth, hypogylcemia Maternal: spontaneous onset of labour, induction of labour, caesarean delivery, caesarean delivery for fetal distress, pre-eclampsia, gestational hypertension					
Notes	Sample size calculated for main study, no prespecified single primary outcome for this pilot complementary study					
Risk of bias						
Item	Authors' judgement	Description				
Adequate sequence generation?	Yes	Computer-generated numbers.				
Allocation concealment?	Yes	Central telephone randomisation service.				
Blinding? All outcomes	Unclear	Not feasible to blind participant and unclear if clinician or outcome assessors were blinded				
Incomplete outcome data addressed? All outcomes	Yes	No loss to follow up documented.				

Characteristics of excluded studies [ordered by study ID]

Study	Reason for exclusion
Sood 2007	Participants were high risk, not specifically growth restricted and did not compare 2 regimens

Study	Reason for exclusion
Williams 2003	Participants were high risk, not specifically growth restricted and compared nonstress test (cardiotocography) to umbilical artery doppler therefore this does not qualify as a regimen per our definition

DATA AND ANALYSES

Comparison 1

Twice-weekly versus fortnightly surveillance

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Perinatal mortality	1	167	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
2 Serious neonatal morbidity	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
3 Stillbirth	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
4 Neonatal death	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
5 Hypoxic ischaemic encephalopathy	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
6 Admission to neonatal special care or intensive care unit	1	167	Risk Ratio (M-H, Fixed, 95% CI)	0.90 [0.58, 1.39]
7 Gestational age at birth	1	167	Mean Difference (IV, Fixed, 95% CI)	-4.0 [-7.79, -0.21]
8 Low birthweight for gestation at birth	1	167	Risk Ratio (M-H, Fixed, 95% CI)	0.80 [0.63, 1.01]
9 Birthweight	1	167	Mean Difference (IV, Fixed, 95% CI)	-53.0 [-184.40, 78.40]
10 Preterm birth	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
11 Low Apgar scores at 5 minutes	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
12 Cord blood acidosis	1	167	Risk Ratio (M-H, Fixed, 95% CI)	1.29 [0.30, 5.57]
13 Metabolic acidosis or lactic acidosis	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
14 Respiratory distress syndrome	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
15 Use of ventilatory support	1	167	Risk Ratio (M-H, Fixed, 95% CI)	4.83 [0.24, 99.02]
16 Use of inotropic support	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
17 Necrotising enterocolitis	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
18 Grade III or IV intraventricular haemorrhage	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
19 Duration of neonatal hospital stay	0	0	Mean Difference (IV, Fixed, 95% CI)	Not estimable
20 Long-term neurodevelopmental outcomes	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
21 Cerebral palsy	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
22 Emergency caesarean section for fetal distress	1	167	Risk Ratio (M-H, Fixed, 95% CI)	0.96 [0.35, 2.63]
23 Antenatal hospital admission	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
24 Induction of labour	1	167	Risk Ratio (M-H, Fixed, 95% CI)	1.25 [1.04, 1.50]
25 Caesarean section	1	167	Risk Ratio (M-H, Fixed, 95% CI)	1.14 [0.54, 2.40]
26 Operative vaginal delivery	0	0	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
27 Length of stay in neonatal special care and/or intensive care unit			Other data	No numeric data
28 Length of antenatal hospital admisison	0	0	Mean Difference (IV, Fixed, 95% CI)	Not estimable
29 Costs of monitoring	0	0	Mean Difference (IV, Fixed, 95% CI)	Not estimable
30 Women's views of care	0	0	Odds Ratio (M-H, Fixed, 95% CI)	Not estimable

Analysis 1.1. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 1 Perinatal mortality

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

Outcome: 1 Perinatal mortality

Study or subgroup	Twice-weekly n/N	Fortnightly n/N			Risk Ratio xed,95% Cl	Weight	Risk Ratio M-H,Fixed,95% C
McCowan 2000	0/85	0/82					Not estimable
Total (95% CI)	85	82					Not estimable
Total events: 0 (Twice-wee	kly), 0 (Fortnightly)						
Heterogeneity: not applica	ble						
Test for overall effect: not	applicable						
Test for subgroup difference	es: Not applicable						
			0.01	0.1	1 10 100		
			Eavours t		Eavours control		

Analysis 1.6. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 6 Admission to neonatal special care or intensive care unit

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

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Outcome: 6 Admission to neonatal special care or intensive care unit

Study or subgroup	Twice-weekly n/N	Fortnightly n/N	M-	Risk Ratio H,Fixed,95% Cl	Weight	Risk Ratio M-H,Fixed,95% CI
McCowan 2000	26/85	28/82		-	100.0 %	0.90 [0.58, 1.39]
Total (95% CI)	85	82		-	100.0 %	0.90 [0.58, 1.39]
Total events: 26 (Twice-we	eekly), 28 (Fortnightly)					
Heterogeneity: not applica	able					
Test for overall effect: Z =	0.49 (P = 0.62)					
Test for subgroup differen	ces: Not applicable					
			0.2 0.5	1 2 5		
			Favours twice week	ly Favours fortni	shtiv	

Analysis 1.7. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 7 Gestational age at birth

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

Outcome: 7 Gestational age at birth

Study or subgroup	Twice-weekly N	Mean(SD)	Fortnightly N	Mean(SD)	Diffe	Mean rence 1.95% Cl	Weight	Mean Difference IV.Fixed,95% CI
	14	Fiean(SD)	IN	riean(SD)	IV,FIXE	1,7 <i>37</i> 6 CI		TV,FIXEO,75% CI
McCowan 2000	85	264 (13)	82	268 (12)	-		100.0 %	-4.00 [-7.79, -0.21]
Total (95% CI)	85		82		-		100.0 %	-4.00 [-7.79, -0.21]
Heterogeneity: not ap	plicable							
Test for overall effect:	Z = 2.07 (P = 0.0)	39)						
Test for subgroup diffe	rences: Not applic	able						
				- 10	-5 0	5	10	
				Eaurouns to	wice weekly	Favours for	trainththy	

Analysis 1.8. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 8 Low birthweight for gestation at birth

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

Outcome: 8 Low birthweight for gestation at birth

Study or subgroup	Twice-weekly n/N	Fortnightly n/N			isk Ratio ed,95% Cl		Weight	Risk Ratio M-H,Fixed,95% Cl
McCowan 2000	47/85	57/82	-	-			100.0 %	0.80 [0.63, 1.01]
Total (95% CI)	85	82	-	-			100.0 %	0.80 [0.63, 1.01]
Total events: 47 (Twice-wee	kly), 57 (Fortnightly)							
Heterogeneity: not applicat	de							
Test for overall effect: Z =	I.88 (P = 0.060)							
Test for subgroup difference	es: Not applicable							
			0.5 I avours twice v	0.7 I	1.5 Favours fo	-		

Analysis 1.9. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 9 Birthweight

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

Outcome: 9 Birthweight

Study or subgroup	Twice-weekly N	Mean(SD)	Fortnightly N	Mean(SD)	Mean Difference IV,Fixed,95% CI	Weight	Mean Difference IV,Fixed,95% CI
McCowan 2000	85	2534 (454)	82	2587 (412)	-	100.0 %	-53.00 [-184.40, 78.40]
Total (95% CI)	85		82		-	100.0 %	-53.00 [-184.40, 78.40]
Heterogeneity: not ap	plicable						
Test for overall effect:	Z = 0.79 (P = 0	.43)					
Test for subgroup diffe	erences: Not app	licable					
				-50	00 -250 0 250	500	
				Eavours t	wice weekly Favours	fortnightly	

Analysis 1.12. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 12 Cord blood acidosis

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

Outcome: 12 Cord blood acidosis

Study or subgroup	Twice-weekly n/N	Fortnightly n/N	Risk Ratio M-H,Fixed,95% CI	Weight	Risk Ratio M-H,Fixed,95% CI
McCowan 2000	4/85	3/82		100.0 %	1.29 [0.30, 5.57]
Total (95% CI)	85	82	-	100.0 %	1.29 [0.30, 5.57]
Total events: 4 (Twice-we	ekly), 3 (Fortnightly)				
Heterogeneity: not applic	able				
Test for overall effect: Z =	0.34 (P = 0.74)				
Test for subgroup differer	ces: Not applicable				
			0.01 0.1 1 10 100		
		Fax	ours twice weekly Favours fortnig	htly	

Analysis 1.15. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 15 Use of ventilatory support

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

Outcome: 15 Use of ventilatory support

Study or subgroup	Twice-weekly n/N	Fortnightly n/N	Risk R M-H,Fixed,9		Weight	Risk Ratio M-H,Fixed,95% CI
McCowan 2000	2/85	0/82			100.0 %	4.83 [0.24, 99.02]
Total (95% CI)	85	82			100.0 %	4.83 [0.24, 99.02]
Total events: 2 (Twice-we	ekly), 0 (Fortnightly)					
Heterogeneity: not applic	able					
Test for overall effect: Z =	= 1.02 (P = 0.31)					
Test for subgroup differer	nces: Not applicable					
			0.01 0.1 1	10 100		
			Favours treatment F	wours control		

Analysis 1.22. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 22 Emergency caesarean section for fetal distress

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

Outcome: 22 Emergency caesarean section for fetal distress

Study or subgroup	Twice-weekly n/N	Fortnightly n/N	Risk Ratio M-H,Fixed,95% (Risk Ratio M-H,Fixed,95% C
McCowan 2000	7/85	7/82		100.0 %	0.96 [0.35, 2.63
Total (95% CI)	85	82	-	100.0 %	0.96 [0.35, 2.63]
Total events: 7 (Twice-we	ekly), 7 (Fortnightly)				
Heterogeneity: not applic	able				
Test for overall effect: Z =	= 0.07 (P = 0.94)				
Test for subgroup differer	ices: Not applicable				
			0.01 0.1 1 10	100	
		Fa	wours twice weekly Favour	rs fortnightly	

Analysis 1.24. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 24 Induction of labour

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

Outcome: 24 Induction of labour

Study or subgroup	Twice-weekly n/N	Fortnightly n/N			Risk Ratio ixed,95% Cl	l	Weight	Risk Ratio M-H,Fixed,95% CI
McCowan 2000	70/85	54/82			-		100.0 %	1.25 [1.04, 1.50]
Total (95% CI)	85	82			•		100.0 %	1.25 [1.04, 1.50]
Total events: 70 (Twice-w	eekly), 54 (Fortnightly)							
Heterogeneity: not applica	able							
Test for overall effect: Z =	2.38 (P = 0.017)							
Test for subgroup differen	ces: Not applicable							
			0.2	0.5	1 2	5		
			Favours twic	e weekly	Eavours	fortnightly		

Analysis 1.25. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 25 Caesarean section

Review: Regimens of fetal surveillance for impaired fetal growth

Comparison: 1 Twice-weekly versus fortnightly surveillance

Outcome: 25 Caesarean section

Study or subgroup	Twice-weekly n/N	Fortnightly n/N	Risk Ratio M-H,Fixed,95% CI	Weight	Risk Ratio M-H,Fixed,95% CI
McCowan 2000	13/85	11/82	-	100.0 %	1.14 [0.54, 2.40]
Total (95% CI)	85	82	-	100.0 %	1.14 [0.54, 2.40]
Total events: 13 (Twice-w	eekly), II (Fortnightly)				
Heterogeneity: not applic	able				
Test for overall effect: Z =	= 0.35 (P = 0.73)				
Test for subgroup differer	ices: Not applicable				
			0.1 0.2 0.5 1 2 5 10		
			Favours twice weekly Favours fortnight	ly	

Analysis 1.27. Comparison 1 Twice-weekly versus fortnightly surveillance, Outcome 27 Length of stay in neonatal special care and/or intensive care unit

Length of stay in neonatal special care and/or intensive care unit

Study	Twice weekly	Fortnightly
McCowan 2000	median 5 days (range 0 to 66)	median 4 days (range 1 to 27)

HISTORY

Protocol first published: Issue 2, 2008

Review first published: Issue 1, 2009

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- * Indicates the major publication for the study

PLAIN LANGUAGE SUMMARY

Testing in pregnancy when there is poor growth of the baby

A baby may not grow at the predicted rate during pregnancy and be born with a low birthweight. This can result in problems for both mother and baby. When poor growth of the baby is suspected during pregnancy, antenatal care includes a combination of tests that are carried out at regular intervals to assess the health of the developing baby. Based on knowledge of the progressive changes any deterioration in the baby's condition can be detected, which allows optimal timing of the birth so that the baby is in the best condition possible. The specified frequency and combinations of tests varies with local and personal policies. Tests may include fetal movement monitoring, fetal heart rate, growth scans, Doppler ultrasound to measure blood flow, and changes in fetal heart rate with movement to check for possible stillbirth.

The review authors identified only one controlled trial, from New Zealand. This trial randomised 167 women who were between 24 and 36 weeks' pregnant where ultrasound showed a small-for-gestational-age baby. They received a set combination of tests either twice-weekly or fortnightly. With more frequent testing, women were 25% more likely to have induced labour. Overall their babies were born four days earlier than in the fortnightly surveillance group where spontaneous onset of labour was more likely to occur. The mean gestational age at birth was just under 38 weeks in the twice-weekly group and just over 38 weeks in the fortnightly group, which was unlikely to have an impact on the health of the newborn. The number of caesarean sections, either for fetal distress or because of failure of induction, was no different. No information was available on length of antenatal hospital admission or operative vaginal births and infants were not followed up to determine neurodevelopment and cerebral palsy. This study excluded pregnancies with abnormal Doppler studies and disorders of the amniotic fluid. More studies are needed and the women's views on the testing are also important.