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# Bed rest with or without hospitalisation for hypertension during pregnancy (Review)

Meher S, Abalos E, Carroli G

Meher S, Abalos E, Carroli G. Bed rest with or without hospitalisation for hypertension during pregnancy. *Cochrane Database of Systematic Reviews* 2005, Issue 4. Art. No.: CD003514. DOI: 10.1002/14651858.CD003514.pub2.

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#### [Intervention Review]

# Bed rest with or without hospitalisation for hypertension during pregnancy

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**Editorial group:** Cochrane Pregnancy and Childbirth Group. **Publication status and date:** New search for studies and content updated (no change to conclusions), published in Issue 2, 2010.

**Citation:** Meher S, Abalos E, Carroli G. Bed rest with or without hospitalisation for hypertension during pregnancy. *Cochrane Database of Systematic Reviews* 2005, Issue 4. Art. No.: CD003514. DOI: 10.1002/14651858.CD003514.pub2.

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### ABSTRACT

#### Background

Bed rest or restriction of activity, with or without hospitalisation, have been advocated for women with hypertension during pregnancy to improve pregnancy outcome. However, benefits need to be demonstrated before such interventions can be recommended since restricted activity may be disruptive to women's lives, expensive, and increase the risk of thromboembolism.

#### Objectives

To assess the effects on the mother and the baby of different degrees of bed rest, compared with each other, and with routine activity, in hospital or at home, for primary treatment of hypertension during pregnancy.

#### Search methods

We searched the Cochrane Pregnancy and Childbirth Group's Trials Register (January 2010).

#### **Selection criteria**

Randomised trials evaluating bed rest for women with hypertension in pregnancy were selected.

#### Data collection and analysis

Two review authors assessed trials for inclusion independently, and extracted data. Data were entered into RevMan software and double-checked.

#### **Main results**

Four small trials (449 women) were included. Three were of good quality. Two trials (145 women) compared strict bed rest with some rest, in hospital, for women with proteinuric hypertension. There was insufficient evidence to demonstrate any differences between the groups for reported outcomes. Two trials (304 women) compared some bed rest in hospital with routine activity at home for non-proteinuric hypertension. There was reduced risk of severe hypertension (one trial, 218 women; relative risk (RR) 0.58, 95% confidence interval (CI) 0.38 to 0.89) and a borderline reduction in risk of preterm birth (one trial, 218 women; RR 0.53, 95% CI 0.29 to 0.99) with some rest compared to normal activity. More women in the bed rest group opted not to have the same management in future pregnancies, if the choice were given (one trial, 86 women; RR 3.00, 95% CI 1.43 to 6.31). There were no significant differences for any other outcomes.



#### **Authors' conclusions**

Few randomised trials have evaluated rest for women with hypertension during pregnancy, and important information on side-effects and cost implication is missing from available trials. Although one small trial suggests that some bed rest may be associated with reduced risk of severe hypertension and preterm birth, these findings need to be confirmed in larger trials. At present, there is insufficient evidence to provide clear guidance for clinical practice. Therefore, bed rest should not be recommended routinely for hypertension in pregnancy, especially since more women appear to prefer unrestricted activity, if the choice were given.

### PLAIN LANGUAGE SUMMARY

#### Bed rest with or without hospitalisation for hypertension during pregnancy

Not enough evidence to say if bed rest in pregnancy helps women and their babies when women have high blood pressure.

High blood pressure in pregnant women can contribute to babies being small, being born too soon and having considerable health problems. Women with high blood pressure are often advised to rest in bed either at home or in hospital. It is suggested that this might help to reduce the mother's blood pressure and so provide benefits for the baby. However, there may be adverse effects; for example, some women may find it stressful, it may contribute to blood clots in the legs and can put a burden on the woman's family. Although one small trial suggested that there may be some possible benefits, there are insufficient data to be confident. Moreover, trials did not address possible adverse effects of bed rest. More women seemed to prefer normal activity at home rather than resting in hospital, if a choice were given. Further research is needed.



### BACKGROUND

Raised blood pressure during pregnancy is one of the commonest medical complications, occurring in 6% to 8% of all pregnancies (ACOG 1996). There are controversies about the definition of hypertensive disorders during pregnancy, and several classifications have been suggested (ASSHP 1993; Davey 1988; NHBPEP 2000; North 1999; Roberts 1993). These are discussed in more detail in Abalos 2007. However, most include the four main categories (a) chronic hypertension; (b) pre-eclampsia, characterised by hypertension with proteinuria; (c) pre-eclampsia superimposed on chronic hypertension; and (d) pregnancyinduced hypertension or gestational hypertension, transient hypertension without proteinuria. Hypertension is usually defined as blood pressure of at least 140 mmHg systolic or 90 mmHg diastolic, or both, and proteinuria is defined as at least 0.3 g protein in a 24-hour collection, or 30 mg/dL or more (1+ on dipstick) in a random urine sample (NHBPEP 2000). An important issue in classification of hypertensive disorders of pregnancy is the ability to differentiate hypertensive disorders present before pregnancy from those that are pregnancy-specific, of which the more ominous is pre-eclampsia. The impact of the two groups of conditions on the mother and fetus is different, as is their management.

About a quarter of the cases of hypertension during pregnancy are due to chronic hypertension (Lindheimer 1999). Women with uncomplicated mild to moderate hypertension often have uneventful pregnancies, but nearly 20% of women with chronic hypertension develop superimposed pre-eclampsia; development of pre-eclampsia or severe hypertension is responsible for most of the morbidity in these pregnancies (Sibai 1998). Preeclampsia, either presenting 'de novo' or superimposed on chronic hypertension or renal disease is the disorder most likely to endanger both mother and fetus (Lindheimer 1999). The cardinal features of pre-eclampsia are hypertension and proteinuria, but women may also have abnormalities of liver function or coagulation, or both, thrombocytopenia (low platelet counts) or eclampsia (the rare occurrence of seizures superimposed on pre-eclampsia). Potential consequences for the fetus are intrauterine growth restriction, preterm birth, or death. For women with gestational hypertension the outcome is usually good, although hypertension frequently recurs in subsequent pregnancies (Chesley 1978).

The potential worsening from mild/moderate to severe disease and the difficulty in distinguishing between pre-eclampsia, chronic, secondary and gestational hypertension, and combinations of these entities gives strong support for close supervision of all hypertensive pregnant women (NHBPEP 2000; Saudan 1998; Walker 2000).

Systolic blood pressure has been demonstrated to be higher in ambulating women, which is the logic behind the use of bed rest for hypertension during pregnancy. Bed rest, either in hospital or at home, has a long history in the treatment of mild pregnancyinduced hypertension and pre-eclampsia, and has drawn the attention of different authors. Since 1952, when Hamlin reported the virtual disappearance of eclampsia with a policy of vigorous antenatal education and supervision (including early admission to hospital and numerous interventions besides bed rest) (Hamlin 1952), the notion that bed rest might improve the outcome of hypertensive pregnancies became engrained in the mind of practitioners and health service planners. Hospitalisation was then strongly recommended, not only for closer supervision to monitor the mother and fetus for signs of deterioration, but also for bed rest. Thus, hospitalisation to enhance compliance with bed rest became a common practice for decades. It is not known whether bed rest is beneficial, regardless of whether combined with hospitalisation. Over the last ten years, domiciliary or day care facilities have been used in Europe to monitor pregnancies with hypertension or mild pre-eclampsia, thus avoiding hospitalisation. However, use of such facilities appears to occur infrequently in many other settings, especially in developing countries.

The extent to which bed rest is prescribed in pregnancy is difficult to determine, either in hospital or in outpatient clinics, as it is not clear whether the advice to rest in bed is recorded as frequently as it is advised (Goldenberg 1994). Using the 1988 US National Infant Mortality Survey (Sanderson 1991) Goldenberg et al found that 4.8% of pregnant women were advised by their physicians to rest in bed for at least one week because of hypertension (Goldenberg 1994). This represents nearly 200,000 women of the four million annual United States births. A great number of women also reduce their work load or stop working. This could be stressful for the woman and disruptive for her family (Kramer 1986; Maloni 1993), and could also have important consequences in terms of costs for both the families and the health services.

Even though, for most women with hypertension, the period of bed rest is likely to be relatively short, concerns about its safety have been raised. There may be an increase in the likelihood of adverse events such as thrombosis, muscle atrophy, bone demineralisation and calcium depletion (Maloni 1993).

Bed rest has been prescribed both as a primary therapy, and as an adjunct to other treatments for hypertension during pregnancy, such as antihypertensive drugs. There is a need to evaluate bed rest, in hospital or at home, to determine whether its use improves pregnancy outcome sufficiently to warrant such a recommendation. If bed rest is found to be clearly effective in improving outcome, then the costs, disruptions in normal living and maternal stress should be evaluated further in risk and cost-benefit analyses.

A number of other interventions for women with hypertension during pregnancy are covered by other Cochrane reviews. These include salt restriction (Duley 2005), antiplatelet agents (Duley 2007), abdominal decompression (Hofmeyr 1996), antihypertensive drug treatments for mild to moderate hypertension (Abalos 2007), and drug regimens for severe hypertension (Duley 2006). There is also a separate review assessing the effect of oral beta blockers in mild to moderate hypertension during pregnancy (Magee 2003).

The aim of this review was to assess the relative benefits, risks, and side-effects for the woman and baby of different degrees of bed rest compared with each other or with routine activity, with or without hospitalisation, as a primary therapy for the treatment of hypertension during pregnancy. If bed rest is beneficial overall, a secondary aim was to assess the comparative effects of bed rest in hospital and at home.



# OBJECTIVES

To determine the possible benefits, risks, and side-effects of different degrees of bed rest compared with each other, and with routine activity, in hospital or at home, as primary treatment for raised blood pressure during pregnancy. A secondary aim was to compare the effects of bed rest in hospital with bed rest at home.

## METHODS

#### Criteria for considering studies for this review

#### **Types of studies**

Randomised controlled trials evaluating bed rest as a primary treatment for women with hypertension in pregnancy. Trials with quasi-randomised designs were excluded.

#### **Types of participants**

Pregnant women with raised blood pressure (defined, whenever possible, as systolic blood pressure greater than or equal to 140 mmHg or diastolic blood pressure greater than or equal to 90 mmHg, or both). Women were included regardless of whether they had proteinuria or previous antihypertensive treatment and irrespective of whether they had a singleton or multiple pregnancy.

#### **Types of interventions**

Interventions considered for this review included: (1) any comparisons, in hospital or at home, between:

- strict bed rest: when the woman is confined to bed, and only allowed up to go to the toilet;
- some rest: when the woman is encouraged to restrict activity, whether in bed or not, but is allowed some voluntary activity; and
- routine activity: unrestricted activity.
- (2) comparisons of any rest in hospital versus any rest at home.

We excluded studies evaluating bed rest as adjunctive therapy to other interventions for hypertension in pregnancy.

#### Types of outcome measures

### For the woman

#### **Main outcomes**

(1) Severe hypertension: defined, whenever possible, as either systolic blood pressure greater than or equal to 160 mmHg, or diastolic blood pressure greater than or equal to 110 mmHg. Trials where the definition of severe hypertension is not clear will also be included and clearly documented.

(2) Pre-eclampsia: defined, whenever possible, as new onset proteinuria (greater than or equal to 1+ or greater than or equal to 300 mg/24 hour) after 20 weeks' gestation in pregnant women with hypertension.

#### Other outcomes

(3) Death: during pregnancy, childbirth, or up to 42 days after end of pregnancy.

(4) Severe pre-eclampsia: the following are features of severe disease: severe hypertension, severe proteinuria (usually greater than or equal to 3 g protein in 24 hours, or 3+ on dipstick),

reduced urinary volume (less than or equal to 400 to 500 ml in 24 hours), neurological disturbances such as headache, visual disturbances, and exaggerated tendon reflexes, upper abdominal pain, pulmonary oedema (fluid in the lungs), impaired liver function tests, high serum creatinine, low platelets, intrauterine growth restriction or reduced liquor volume. Trials where the definition of severe pre-eclampsia is not clear will be included.

(5) Severe maternal morbidity such as eclampsia, liver or renal failure, haemolysis, elevated liver enzymes, low platelets syndrome, disseminated intravascular coagulation), and cerebrovascular accident (stroke).

(6) Use of antihypertensive drugs to control blood pressure.

(7) Placental abruption.

(8) Elective delivery: including elective caesarean sections and induction of labour.

(9) Caesarean section.

(10) Use of hospital resources: antenatal hospital admission and length of stay greater than seven days, visit to day care unit.(11) Side-effects such as thromboembolism, muscle atrophy, bone demineralisation, and calcium depletion.

#### For the baby

#### Main outcomes

(1) Death: fetal deaths including miscarriage (fetal losses before viability, usually taken as 20 or 24 weeks), stillbirth (death in utero after 20 or 24 weeks' gestation or however defined), perinatal death (stillbirth and death in the first seven days of life), and neonatal death (death in the first 28 days after birth).

(2) Small-for-gestational age: low birthweight for gestational age, below the third, fifth or 10th percentile, using the most severe reported.

(3) Preterm birth: all births less than or equal to 37 completed weeks, and more severe prematurity, defined as less than 32 or less than 34 weeks.

#### Other outcomes

(4) Apgar score at five minutes: low (less than seven) and very low (less than four).

- (5) Endotracheal intubation.
- (6) Admission to neonatal intensive care nursery.
- (7) Respiratory distress syndrome.

#### Satisfaction outcomes

- 1. Woman's views of the intervention and satisfaction with care.
- 2. Care provider's satisfaction with care.

#### Economic cost outcomes

- 1. Providers' costs.
- 2. Users' (women/families) costs.

#### 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 (January 2010).

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



- 1. 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 Coordinator searches the register for each review using the topic list rather than keywords.

For details of the additional searching we undertook for the initial version of the review, using the search strategy listed in Meher 2005, *see* Appendix 1.

We did not apply any language restrictions.

#### Data collection and analysis

# **Selection of studies**

Assessment of the trials for inclusion was performed independently by two authors. Authors were not blinded to the authors, source of the articles, and results. Differences in opinion were resolved by discussion.

#### Assessment of study quality

Methodological quality was assessed with the following criteria. (1) How selection bias at entry to the trial was avoided: how the random assignment was generated and how allocation concealment was ensured. A quality score for concealment of allocation was assigned to each trial, using the following criteria: (A) adequate concealment of allocation, such as telephone randomisation, consecutively numbered sealed opaque envelopes;

(B) unclear whether adequate concealment of allocation;

(C) inadequate concealment of allocation such as open randomnumber tables, sealed envelopes that are not numbered and opaque.

Trials with quasi-random design were excluded.

(2) How selection bias after entry to the trial was avoided: what was the withdrawal rate and whether the analysis was done on an intention-to-treat basis. For withdrawals, studies were classified as follows:

- (A) less than 5% of participants excluded from analysis;
- (B) 5% to 10% of participants excluded from analysis;

(C) more than 10% and up to 20% of participants excluded from analysis.

Trials were excluded if it was not possible to analyse data on an intention-to-treat basis or 20% or more participants were excluded, or both.

(3) How assessment bias was avoided: how the investigator, the participant or the person assessing the outcome, or both, was blinded to the allocated group. In case of no blinding at all, how objective was the endpoint measured. Blinding was assessed in the following way:

(a) blinding of participant (yes/no/unclear or not specified);

(b) blinding of caregiver (yes/no/unclear or not specified);

(c) blinding of outcome assessment (yes/no/unclear or not specified).

#### Data extraction and entry

Data were extracted by two authors, and discrepancies were resolved through discussion. Data were entered onto the Review Manager software (RevMan 2003), and checked for accuracy.

#### **Statistical analyses**

Statistical analyses were carried out using the Review Manager software (RevMan 2003). All outcomes were dichotomous data, so results are presented as summary relative risk with 95% confidence intervals. The I<sup>2</sup> statistic was used to assess heterogeneity between trials, where relevant. As significant heterogeneity (I<sup>2</sup> greater than 50%) was not detected, results were pooled using a fixed-effect model.

#### Subgroup analyses

A prespecified subgroup was based on the type of hypertensive disease at trial entry: gestational hypertension; hypertension with proteinuria; and chronic hypertension. This analysis was not done because of the small numbers in the review. It will be included in future updates when sufficient data are available.

#### RESULTS

#### **Description of studies**

Details for each trial can be found in the 'Characteristics of included studies' table and the 'Characteristics of excluded studies' table.

This review includes four small trials involving a total of 449 women. Two were conducted in Zimbabwe (Crowther 1986; Crowther 1992), one in the UK (Mathews 1982), and one in Hong Kong (Leung 1998). Two were multicentre (Crowther 1992; Mathews 1982).

#### Participants

All participants were women with a singleton pregnancy, between 26 and 38 weeks' gestation at trial entry. Two trials recruited both primigravidae and multigravidae women (Crowther 1992; Leung 1998), and the other two did not report on parity.

The women had diastolic blood pressure between 90 to 110 mmHg. One trial also specified systolic pressure of at least 140 mmHg. Two studies included women with chronic or gestational hypertension without proteinuria (Crowther 1992; Leung 1998). The other two included women with unspecified proteinuric hypertension (Crowther 1986; Mathews 1982). No trials reported on whether women were using antihypertensive therapy at trial entry, and only one reported on use of antihypertensives as an outcome measure (Leung 1998).



#### Interventions

Two trials compared strict bed rest in hospital with some rest in hospital (Crowther 1986; Mathews 1982). Two trials compared some bed rest in hospital with normal activity at home (Crowther 1992; Leung 1998).

#### Outcomes

Definitions of outcomes used by trialists were consistent with those specified in this review.

One trial reported on women's views of the intervention and satisfaction with care (Leung 1998). The other outcomes from this study have not been reported because more than 20% participants were excluded from analysis for all other outcomes.

No trials reported on side-effects of bed rest or costs to health care.

#### **Excluded studies**

Seven studies were excluded from the review. Four were excluded because all comparison groups had bed rest. Three of these studies compared an antihypertensive drug plus bed rest with bed rest alone (Cameron 1985; Catalano 1997; Sibai 1992). The fourth was a three-arm study in which bed rest alone was compared with both compliance enhancement training and a bio-behavioural intervention (Somers 1989). Two studies were excluded because the participants were pregnant women with normal blood pressure (Herrera 1993; Spinapolice 1983). One trial was excluded because women were able to opt out of the trial after randomisation if they were not happy with the allocated treatment, and they were excluded from the analysis. Although the number of women who opted out is not known even after contacting trialists, it seems likely this was greater than 20% (Mathews 1977).

#### **Risk of bias in included studies**

Details for each trial are in the 'Characteristics of included studies' table.

Three trials were of good quality (Crowther 1986; Crowther 1992; Leung 1998) and one was of uncertain quality (Mathews 1982).

#### Allocation concealment

Allocation concealment was adequate in three trials (Crowther 1986; Crowther 1992; Leung 1998) but unclear in the fourth (Mathews 1982).

#### Completeness of follow up

Two trials stated that there were no losses to follow up (Crowther 1986; Crowther 1992). One trial (Mathews 1982) excluded 13% of women from the analysis (equal in both groups). Reasons for exclusion included women not complying with their allocated treatment. The fourth trial (Leung 1998), after recruiting women on a single reading of hypertension, excluded a large number of women (25.6%) mainly because blood pressure was no longer raised in these women after randomisation or because the women went into labour prior to confirmation of the diagnosis of hypertension (13 from bed rest group and 8 from control group). However, only 4.4% women were excluded from analysis for data on women's views of the intervention.

#### Blinding

One trial stated that only the outcome assessor for fetal outcomes was blinded (Crowther 1992). The other three trial reports did not mention blinding. Given the intervention under evaluation, blinding of the participants to this intervention is not possible. Although it would be possible to blind outcome assessment, it is unlikely that this was done because this would be quite a major undertaking, and it is reasonable to assume that it would have been reported had it been done.

#### **Effects of interventions**

Four small trials (449 women) were included.

# Comparison one: strict bed rest in hospital versus some rest in hospital

Two trials (145 women) evaluated strict bed rest in hospital versus some rest in hospital.

#### Severe hypertension

One trial (105 women) reported the risk of severe hypertension for strict bed rest in hospital compared to some rest in hospital (relative risk (RR) 1.18, 95% confidence interval (CI) 0.93 to 1.49).

# Death of baby: miscarriage, stillbirth, perinatal death, or neonatal death

Two trials (145 women) reported on stillbirth, perinatal death, and neonatal death. Even taking all deaths together there were insufficient data for any reliable conclusions about the possible differential effect (RR 1.07, 95% CI 0.52 to 2.19).

#### Preterm birth

One trial (105 women) reported the risk of preterm birth for strict bed rest in hospital compared to some rest in hospital (RR 0.98, 95% CI 0.71 to 1.35).

#### Other outcomes

There were no statistically significant differences for any other outcomes when comparing strict bed rest in hospital with some rest in hospital: two trials (145 women) reported on severe preeclampsia (RR 1.50, 95% CI 0.98 to 2.30), and one trial (105 women) reported on eclampsia (RR 0.33, 95% CI 0.01 to 7.85), elective delivery (RR 1.08, 95% CI 0.87 to 1.34), placental abruption (RR 4.91, 95% CI 0.24 to 99.82), endotracheal intubation (RR 0.65, 95% CI 0.11 to 3.76) and admission of the baby to neonatal intensive care nursery (RR 0.75, 95% CI 0.49 to 1.17).

# Comparison two: some rest in hospital versus routine activity at home

Two trials (304 women) evaluated some rest in hospital versus routine activity at home in women with non-proteinuric hypertension.

#### Severe hypertension

There was a reduction in the risk of severe hypertension (one trial, 218 women; RR 0.58, 95% CI 0.38 to 0.89) with some rest in hospital compared to normal activity at home.



#### Pre-eclampsia

In one trial (218 women), the RR for developing pre-eclampsia was 0.98 (95% CI 0.80 to 1.20).

# Death of baby: miscarriage, stillbirth, perinatal death, or neonatal death

In one trial (218 women) reporting on stillbirth, perinatal death, and neonatal death, there were insufficient data for any reliable conclusions even when all deaths were taken together (RR 1.96, 95% CI 0.18 to 21.34).

#### Preterm birth

In one trial (218 women), the RR for preterm birth was 0.53 (95% CI 0.29 to 0.99) and for very preterm birth (less than 34 weeks' gestation) 0.49 (95% CI 0.09 to 2.62) with some rest in hospital compared to normal activity at home.

#### Small-for-gestational age

The RR of small-for-gestational-age babies was 0.98 (95% CI 0.51 to 1.91; one trial (218 women)).

#### Other outcomes

#### Women's views

One trial (88 women) reported on women's views. Although both groups seemed to be equally satisfied with their care (RR 0.98, 95% CI 0.93 to 1.02), more women in the bed rest group opted not to have the same management in future pregnancies, if the choice were given (RR 3.00, 95% CI 1.43 to 6.31).

There were no statistically significant differences for any other reported outcomes including elective delivery (RR 0.98, 95% CI 0.70 to 1.37), caesarean section (RR 1.41, 95% CI 0.79 to 2.52), and admission to neonatal intensive care unit (RR 0.82, 95% CI 0.37 to 1.81).

#### DISCUSSION

Bed rest or restriction of normal activity either at home or in hospital have long been advocated for women with high blood pressure during pregnancy. Despite this, only four small adequately controlled trials have evaluated the potential effects of these interventions. The evidence available from these trials is insufficient to draw reliable conclusions for clinical practice.

Two trials compared strict bed rest in hospital with some rest in hospital for pregnant women with proteinuric hypertension. We have no evidence that there is any difference in outcome between the two interventions. One trial was of good quality (Crowther 1986), and the other was of uncertain quality (Mathews 1982). The findings between trials were mostly consistent, except for perinatal mortality where the point effect of the good quality trial showed worse outcome with strict bed rest and the point effect of the other trial showed benefit from strict bed rest. However, the confidence intervals for both trials were wide and crossed the line of no effect for this outcome, implying that available evidence is insufficient, and the true answer may lie anywhere.

Two trials compared some rest in hospital with routine activity at home for pregnant women with non-proteinuric hypertension, and were of good quality (Crowther 1992; Leung 1998). In one trial (218 women), there was a 42% reduction in the risk of severe hypertension for women who rested in hospital. This finding must be interpreted with caution, as the number of women in the trial was small, so results are more susceptible to the play of chance. The confidence interval is also wide, indicating that the true effect may be anywhere between an 11% reduction in risk to a 62% reduction. Women allocated routine activity at home were monitored with weekly blood pressure measurements in antenatal clinics. It is not reported whether the diagnosis of severe hypertension was based on persistent elevation of blood pressure, or a single episode, and whether antihypertensive medication was required. The clinical significance of this outcome is therefore unclear. It may, at least in part, be related to 'white coat hypertension', where the stress of coming to a hospital leads to transient elevation in blood pressure. This same trial also showed a borderline statistically significant reduction in the risk of preterm birth (RR 0.53, 95% CI 0.29 to 0.99) for women with some bed rest in hospital. However, there was no apparent effect on very preterm birth. As above, the numbers are small, and the confidence interval is wide. It is therefore difficult to draw any firm conclusions about the possible differential effects on preterm birth. While statistically significant results are present in this review, they need to be confirmed, and then clinically significant benefits still need to be demonstrated.

Questions remain about the quantification of rest and activity in the trials. We have made an attempt to quantify 'rest' by dividing it into 'strict' and 'some' based on definitions used by trialists, but the hours of bed rest each day are not reported. Standardising definitions for 'routine' or unrestricted activity is even more difficult. One woman's routine activity may be strenuous work all day, whereas another woman's routine activity may be light work with frequent restful breaks. Although it can be difficult to address these issues in trials, better descriptions of what constitutes rest and baseline level of activity would add to the validity and applicability of results obtained from this review.

The potential effects of bed rest may be related to the type of hypertensive disease a woman has. Although the effects of bed rest on women with proteinuric hypertension (first comparison) are unclear, women with non-proteinuric hypertension (second comparison) seemed to benefit from some rest, with a reduced risk of severe hypertension. Further trials are needed to assess the influence of type of hypertensive disease on effects of bed rest.

All three trials evaluated the effects of bed rest in hospital. It is arguable whether this rest is actually 'restful' for all women, or does the added stress of hospitalisation in certain women undermine any benefits of the prescribed bed rest. Although no trials specifically reported on women's views of bed rest with regards to the disruption to their lives, or stress associated with hospitalisation, in one trial, less women in the bed rest group opted to have the same treatment in future pregnancies, possibly as a result of these factors.

Bed rest in hospital or at home has financial implication for women and their families, and for healthcare services, but included trials did not report on these costs.

Prolonged bed rest may be associated with complications such as thrombosis, muscle atrophy or bone demineralization. Bed rest may be prescribed for a variable duration in pregnancy, but it is likely that the risk of adverse effects would be higher with lengthier periods of immobility. However, included trials did not report on

adverse effects. Greater awareness of the hazards of prolonged periods of immobility may well make interventions such as strict bed rest obsolete for hypertension in pregnancy. What merits further evaluation is whether some rest or restriction of activity, either at home or in hospital, is beneficial.

Bed rest in hospital allows increased surveillance of the woman, and it is proposed that timely access to medical care in hospital may improve pregnancy outcome. At present we have little evidence to support or refute this argument, but if such benefits do exist, they must be weighed against the numbers needed to treat to avert any adverse outcome. On the other hand, being in hospital may predispose to detection bias, and provoke a more interventionalist approach. Moreover, where increased surveillance is required, day care units, where available, are now becoming widely recognised as alternatives to hospital admission for management of complicated pregnancies (Kröner 2001).

Given the paucity of trials, and the absence of relevant and important information from available trials, no reliable conclusions can be made from this review to guide clinical practice regarding bed rest for management of women with hypertension in pregnancy. No conclusive, clinically significant benefits of bed rest have been demonstrated, the possibility of adverse effects remains, and women appear to prefer normal activity and outpatient care in comparison to bed rest with hospitalisation as a care plan.

## AUTHORS' CONCLUSIONS

#### **Implications for practice**

Women with hypertension in pregnancy are often advised to rest in bed, either at home or in the hospital, to prevent

progression of hypertension and improve pregnancy outcome. This intervention has been evaluated in few well-controlled trials, and there is insufficient evidence to provide clear guidance for clinical practice. Although one small trial suggests that some bed rest may be associated with a reduced risk of severe hypertension and preterm birth, these findings need to be confirmed in other larger trials. Moreover, it appears that more women prefer outpatient management with unrestricted activity in comparison to hospitalisation and bed rest if the choice were given. Evidence currently available from randomised trials does not support routine recommendation of bed rest for hypertension in pregnancy.

#### **Implications for research**

Large well-controlled randomised trials are needed to evaluate the benefits and risks of rest in hospital and at home for women with proteinuric and non-proteinuric hypertension in pregnancy. Trials need to report not only on pregnancy outcome, but also on the potential side-effects of bed rest, women's views of the intervention and the costs involved, as these factors are also important determinants in formulating recommendations for clinical practice.

#### ACKNOWLEDGEMENTS

Our thanks to Lelia Duley for her helpful comments on the review.

As part of the pre-publication editorial process, this review has been commented on by four peers (an editor and three referees who are external to the editorial team), one or more members of the Pregnancy and Childbirth Group's international panel of consumers and the Group's Statistical Adviser.

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\* Indicates the major publication for the study

Crowther 1986									
Methods	Randomisation: block randomisation using variable blocks and random number tables. Allocation concealment: consecutively numbered opaque, sealed envelopes. Follow up: no loss of participants (A). Blinding: none.								
Participants	-	105 women with a singleton pregnancy at 28-38 weeks, with proteinuric hypertension (diastolic BP 90-109 mmHg and proteinuria 1+ or more). No other complications of pregnancy.							
Interventions		Exp: admission to hospital for strict bed rest until delivery. Ambulation only to toilet. Controls: allowed to move around the hospital ward as desired.							
Outcomes	eclampsia; eclampsia; Baby: preterm delivery	Women: severe hypertension (diastolic BP > 109 mmHg); increase in proteinuria; fulminating pre- eclampsia; eclampsia; placental abruption; IOL. Baby: preterm delivery; low birthweight; very low birthweight; passage of meconium; Apgar score; perinatal death; early neonatal death; stillbirth; endotracheal intubation; admission to special care nursery.							
Notes	Setting: Zimbabwe. 1 centre.								
Risk of bias									
Bias	Authors' judgement	Support for judgement							
Allocation concealment?	Low risk	A - Adequate							

#### Crowther 1992

crowther 1992									
Methods	Randomisation: block randomisation, stratified into 3 groups: primiparous, multiparous without chronic hypertension, and multiparous with chronic hypertension. Allocation concealment: consecutively numbered, opaque, sealed envelopes. Follow up: no loss of participants (A). Blinding: for fetal outcomes only.								
Participants	teinuric hypertension	218 primigravidae and multigravidae women with a singleton pregnancy at 28-38 weeks, with non-pro- teinuric hypertension (BP at least 140/90 mmHg). Excluded if diastolic BP >/= 110 mmHg, symptomatic, caesarean section scar, or APH during the pregnancy.							
Interventions	daily urinalysis. Controls: normal activ	Exp: admission to hospital for rest. Allowed to move around the ward voluntarily. 4 hourly BP check and daily urinalysis. Controls: normal activity at home with no restrictions. Daily self analysis of urine for protein. Reviewed weekly for BP, weight, bloods.							
Outcomes	Woman: severe hypertension (>/= 160/110 mmHg); proteinuria; caesarean section; IOL. Baby: birthweight (mean); birthweight (< 2500 g); small-for-gestational age (< 10%); admission to in- tensive care nursery; length of stay in hospital; perinatal death; Apgar score at 1 and 5 minutes.								
Notes	Setting: Zimbabwe. 1 maternity hospital and 13 peripheral clinics.								
Risk of bias									
Bias	Authors' judgement	Support for judgement							
Allocation concealment?	Low risk	A - Adequate							



Bias	Authors' judgement Support for judgement
Risk of bias	
Notes	Setting: Hong Kong. 1 centre. The only outcome we have reported from this paper is women's views and satisfaction with care as > 20% women have been excluded from analysis for all other outcomes.
Outcomes	Women: hypertension (dBP > 90 x 2), severe hypertension (dBP >/= 110 mmHg x 2); proteinuria; mode of delivery; IOL; use of antihypertensive medication; women's views and preferences (questionnaire). Baby: birthweight (mean); small-for-gestational age (not defined); admission to intensive care nursery; length of stay in hospital; stillbirth/neonatal death; Apgar score at 1 and 5 minutes.
Interventions	Exp: admission to antenatal ward in hospital. Advised to rest in bed as much as possible. Controls: normal activity at home with no restrictions. Daily self analysis of urine for protein. Reviewed weekly in day care centre (day ward with 12 beds) or outpatient clinic for BP, fetal monitoring, urinaly- sis, bloods.
Participants	90 primigravidae and multigravidae women with a singleton pregnancy at 28-38 weeks, with non-pro- teinuric hypertension (diastolic BP 90-100 mmHg) after 5 minutes rest. Excluded if proteinuria >/= 1+ o symptoms of severe PE.
Methods	Randomisation: women were "allocated randomly" to 2 groups. No further information. Allocation concealment: consecutively numbered, opaque, sealed envelopes. Follow up: 25.6% excluded from analysis. 2 defaulted follow up and delivered somewhere else. Others were excluded either because hypertension was absent in these women postrandomisation or because the women went into labour prior to confirmation of the diagnosis of hypertension (13 from bed rest group and 8 from control group). However, only 4.4% women were excluded from analysis for data on women's views of the intervention (2 woman in each group). Blinding: none.
eung 1998 Methods	

Allocation concealment?	Low risk	A - Adequate

Mathews 1982	
Methods	Randomisation: volunteers allocated "at random" to either of the 2 groups. No further information. Allocation concealment: sealed envelopes. Follow up: 13% participants excluded from analysis for various reasons (3 participants in each group) (C). Blinding: none.
Participants	40 women with a singleton pregnancy at 26 to over 36 weeks, with proteinuric hypertension (diastolic BP 90 to 109 mmHg, and proteinuria more than a trace), and asymptomatic.
Interventions	Exp: admission to hospital for strict bed rest. Controls: allowed to move around the ward in hospital. A pedometer was attached to both groups of women as a crude measure of whole body activity.
Outcomes	Women: plasma urea and urate; serum human placental lactogen and oestriol; development of pre- monitory symptoms of eclampsia; hypertension; proteinuria; mode of delivery. Baby: perinatal death (stillbirths and neonatal death), gestation at delivery, birthweight, small-for-ges- tational age, sex of baby.
Notes	Setting: UK. 2 centres.

#### Mathews 1982 (Continued)

Missing data: complete data on all 40 participants available only for 2 outcomes in the published report: perinatal death and development of premonitory symptoms. For other outcomes, data reported for only 10 'high-risk' participants. Trialist contacted for missing data for other outcomes but data were not available.

Risk of bias							
Bias	Authors' judgement	Support for judgement					
Allocation concealment?	Unclear risk	B - Unclear					

APH: antepartum haemorrhage BP: blood pressure Exp: experiment IOL: induction of labour PE: pre-eclampsia dBP: diastolic blood pressure

## Characteristics of excluded studies [ordered by study ID]

Study	Reason for exclusion							
Cameron 1985	Comparison of labetalol with bed rest versus bed rest alone, in hospital. Abstract only.							
	Methods: "randomised". No further information. Participants: 85 pregnant women with hypertension. Intervention: as above. Outcomes: blood pressure, proteinuria, biochemical and haematological tests, tests of placental function, adverse effects of therapy on mother and baby.							
Catalano 1997	Comparison of nifedipine with bed rest versus bed rest alone, in hospital for women with mild pre- eclampsia.							
	Methods: "randomly allocated". No further information. Participants: 100 women at 26-36 weeks' gestation, with mild pre-eclampsia (not defined in trans- lated summary). Intervention: as above. Outcomes: blood pressure; prolongation of pregnancy; days of hospitalisation; preterm birth; mean birthweight; small-for-gestational age.							
Herrera 1993	Participants: normotensive women.							
	Methods: randomisation by computer-generated list. Allocation concealment by closed envelopes. Participants: 74 primigravida women at 28-29 weeks' gestation with normal blood pressure and a positive roll-over test. Intervention: bed rest at home and nutritional supplements (soy protein, linoleic acid, calcium) versus no bed rest at home and placebo (iron tablets). Outcomes: pregnancy-induced hypertension; pre-eclampsia; caesarean section; gestational age at birth; mean birthweight.							
Mathews 1977	Participants excluded after randomisation if they opted out of the study because they were not happy with allocated treatment. Trialist contacted to determine the number of participants excluded after randomisation, but this information was not available.							
	Methods: participants "allocated at random" to 1 of 4 groups using previously prepared cards con- tained in envelopes. No further information. Participants: 135 women with singleton pregnancy between 28 to over 38 weeks, with non-pro- teinuric hypertension (diastolic BP between 90 and 109 mmHg), and asymptomatic.							

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Study	Reason for exclusionIntervention: (4 groups) bed rest in hospital with or without sedation (phenobarbitone 15 mg 3 times daily) versus normal activity at home with or without sedation.Outcomes: severe hypertension (BP > 109 mmHg); proteinuria; eclampsia; induction of labour; op- erative vaginal delivery; caesarean section; biochemical parameters; stillbirths and neonatal death; fetal distress during labour; Apgar score; small for gestation; preterm birth.							
Sibai 1992	Comparison of nifedipine with bed rest versus bed rest alone, in hospital.							
	Methods: randomised controlled trial. Participants: 200 women at 26-36 weeks' gestation with proteinuric and non-proteinuric hyperten- sion. Interventions: as above. Outcomes: blood pressure; haematological and biochemical parameters; days in hospital; prolon- gation of pregnancy; preterm delivery; gestation at birth; mean birthweight; small-for-gestational age; placental weight; cord blood gas.							
Somers 1989	3-arm trial in which all 3 groups of participants had bed rest varying from 15.8 to 18 hours per day along with additional interventions.							
	Methods: "randomly" assigned. No further information. Participants: 45 women at 30-36 weeks with hypertension, all prescribed restricted physical activi- ty for hypertension. Interventions: 3 arms: bed rest at home versus bed rest and compliance enhancement training at home verus bed rest, compliance enhancement training, and bio-behavioral interventions at home. Outcomes: mean arterial pressure; compliance with bed rest.							
Spinapolice 1983	Participants normotensive pregnant women.							
	Methods: "random allocation". No further information available. Participants: 32 nulliparous women at 28-32 weeks' gestation with a normal blood pressure and positive roll-over test. Intervention: bed rest for 4 to 6 hours at home versus normal activity at home. Outcomes: gestational hypertension; pre-eclampsia; gestation at delivery; birthweight; Apgar scores.							

BP: blood pressure

# DATA AND ANALYSES

# Comparison 1. Strict bed rest in hospital versus some rest in hospital

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size	
1 Severe hypertension	1	105	Risk Ratio (M-H, Fixed, 95% CI)	1.18 [0.93, 1.49]	
2 Severe pre-eclampsia	2	145	Risk Ratio (M-H, Fixed, 95% CI)	1.50 [0.98, 2.30]	
3 Eclampsia	1	105	Risk Ratio (M-H, Fixed, 95% CI)	0.33 [0.01, 7.85]	
4 Elective delivery including elec- tive caesarean section or induc- tion of labour, or both	1	105	Risk Ratio (M-H, Fixed, 95% CI)	1.08 [0.87, 1.34]	



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
5 Placental abruption	1	105	Risk Ratio (M-H, Fixed, 95% CI)	4.91 [0.24, 99.82]
6 Death of baby, including mis- carriage, stillbirth, perinatal death, and neonatal death	2	145	Risk Ratio (M-H, Fixed, 95% CI)	1.07 [0.52, 2.19]
7 Death of baby by timing of death	2		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
7.1 Stillbirth	2	145	Risk Ratio (M-H, Fixed, 95% CI)	1.11 [0.45, 2.75]
7.2 Perinatal death	2	145	Risk Ratio (M-H, Fixed, 95% CI)	1.07 [0.52, 2.19]
7.3 Neonatal death	2	145	Risk Ratio (M-H, Fixed, 95% CI)	0.99 [0.28, 3.50]
8 Preterm birth	1	105	Risk Ratio (M-H, Fixed, 95% CI)	0.98 [0.71, 1.35]
9 Endotracheal intubation	1	105	Risk Ratio (M-H, Fixed, 95% CI)	0.65 [0.11, 3.76]
10 Admission to neonatal inten- sive care nursery	1	105	Risk Ratio (M-H, Fixed, 95% CI)	0.75 [0.49, 1.17]

# Analysis 1.1. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 1 Severe hypertension.

Study or subgroup	Strict bed rest	Some rest	me rest Risk Ratio		Weight		<b>Risk Ratio</b>				
	n/N	n/N			М-Н, F	ixed, 9	95% CI				M-H, Fixed, 95% Cl
Crowther 1986	42/53	35/52				+				100%	1.18[0.93,1.49]
Total (95% CI)	53	52				•	•			100%	1.18[0.93,1.49]
Total events: 42 (Strict bed rest), 35	5 (Some rest)										
Heterogeneity: Not applicable											
Test for overall effect: Z=1.37(P=0.1	17)			1							
		Favours strict rest	0.1	0.2	0.5	1	2	5	10	Favours some rest	

# Analysis 1.2. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 2 Severe pre-eclampsia.

Study or subgroup	Strict bed rest	Some rest		R	isk Ratio			Weight	<b>Risk Ratio</b>
	n/N	n/N		м-н, і	ixed, 95%	CI			M-H, Fixed, 95% CI
Crowther 1986	27/53	20/52			<u> </u>			95.28%	1.32[0.86,2.04]
Mathews 1982	5/20	1/20				+	-	4.72%	5[0.64,39.06]
Total (95% CI)	73	72			•			100%	1.5[0.98,2.3]
Total events: 32 (Strict bed re	est), 21 (Some rest)								
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =	1.63, df=1(P=0.2); I <sup>2</sup> =38.66%								
Test for overall effect: Z=1.85	(P=0.06)			I		i.	1		
	F	avours strict rest	0.01	0.1	1	10	100	Favours some rest	

# Analysis 1.3. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 3 Eclampsia.

Study or subgroup	Strict bed rest	Some rest			Risk Ratio	<b>b</b>		Weight	Risk Ratio
	n/N	n/N		M-H	, Fixed, 95	5% CI			M-H, Fixed, 95% CI
Crowther 1986	0/53	1/52						100%	0.33[0.01,7.85]
Total (95% CI)	53	52						100%	0.33[0.01,7.85]
Total events: 0 (Strict bed rest), 1 (	Some rest)								
Heterogeneity: Not applicable									
Test for overall effect: Z=0.69(P=0.4	19)		1						
	F	Favours strict rest	0.01	0.1	1	10	100	Favours some rest	

# Analysis 1.4. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 4 Elective delivery including elective caesarean section or induction of labour, or both.

Study or subgroup	Strict bed rest	Some rest			Ri	sk Rat	io			Weight	<b>Risk Ratio</b>
	n/N	n/N			М-Н, Р	ixed, 9	5% CI				M-H, Fixed, 95% Cl
Crowther 1986	42/53	38/52				+				100%	1.08[0.87,1.34]
Total (95% CI)	53	52				•				100%	1.08[0.87,1.34]
Total events: 42 (Strict bed rest),	38 (Some rest)										
Heterogeneity: Not applicable											
Test for overall effect: Z=0.74(P=0	.46)			- 1							
		Favours strict rest	0.1	0.2	0.5	1	2	5	10	Favours some rest	

# Analysis 1.5. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 5 Placental abruption.

Study or subgroup	Strict bed rest	Some rest			Risk Ratio			Weight	<b>Risk Ratio</b>
	n/N	n/N		M-	H, Fixed, 959	% CI			M-H, Fixed, 95% Cl
Crowther 1986	2/53	0/52				+		100%	4.91[0.24,99.82]
Total (95% CI)	53	52						100%	4.91[0.24,99.82]
Total events: 2 (Strict bed rest)	, 0 (Some rest)								
Heterogeneity: Not applicable									
Test for overall effect: Z=1.03(F	P=0.3)								
		Favours strict rest	0.01	0.1	1	10	100	Favours some rest	

# Analysis 1.6. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 6 Death of baby, including miscarriage, stillbirth, perinatal death, and neonatal death.

Study or subgroup	Strict bed rest	Some rest		Risk	Ratio			Weight	<b>Risk Ratio</b>
	n/N	n/N		M-H, Fixe	ed, 95% CI				M-H, Fixed, 95% CI
Crowther 1986	11/53	8/52			-			66.88%	1.35[0.59,3.08]
Mathews 1982	2/20	4/20						33.12%	0.5[0.1,2.43]
Total (95% CI)	73	72						100%	1.07[0.52,2.19]
		Favours strict rest	0.1 0.2	0.5	1 2	5	10	Favours some rest	



Study or subgroup	Strict bed rest	Some rest				sk Rat	tio 95% CI			Weight	Risk Ratio M-H, Fixed, 95% Cl
Total events: 13 (Strict bed r	n/N	n/N			м-п, г	ixeu,	95% CI				м-п, гіхец, 95% сі
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =		Ď									
Test for overall effect: Z=0.18	(P=0.86)										
		Favours strict rest	0.1	0.2	0.5	1	2	5	10	Favours some rest	

# Analysis 1.7. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 7 Death of baby by timing of death.

Study or subgroup	Strict bed rest	Some rest	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% CI		M-H, Fixed, 95% Cl
1.7.1 Stillbirth					
Crowther 1986	8/53	4/52	+	50.24%	1.96[0.63,6.12]
Mathews 1982	1/20	4/20		49.76%	0.25[0.03,2.05]
Subtotal (95% CI)	73	72	-	100%	1.11[0.45,2.75]
Total events: 9 (Strict bed rest), 8 (	Some rest)				
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =2.9, d	f=1(P=0.09); I <sup>2</sup> =65.46%				
Test for overall effect: Z=0.23(P=0.8	32)				
1.7.2 Perinatal death					
Crowther 1986	11/53	8/52	<mark></mark>	66.88%	1.35[0.59,3.08]
Mathews 1982	2/20	4/20		33.12%	0.5[0.1,2.43]
Subtotal (95% CI)	73	72	<b>•</b>	100%	1.07[0.52,2.19]
Total events: 13 (Strict bed rest), 12	2 (Some rest)				
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =1.19,	df=1(P=0.27); I <sup>2</sup> =16.19%	)			
Test for overall effect: Z=0.18(P=0.8	36)				
1.7.3 Neonatal death					
Crowther 1986	3/53	4/52		88.98%	0.74[0.17,3.13]
Mathews 1982	1/20	0/20	+	- 11.02%	3[0.13,69.52]
Subtotal (95% CI)	73	72	-	100%	0.99[0.28,3.5]
Total events: 4 (Strict bed rest), 4 (	Some rest)				
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.64,	df=1(P=0.42); I <sup>2</sup> =0%				
Test for overall effect: Z=0.02(P=0.9	98)				
	I	Favours strict rest 0.01	0.1 1 10	<sup>100</sup> Favours some rest	

# Analysis 1.8. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 8 Preterm birth.

Study or subgroup	Strict bed rest	Some rest			Ri	sk Rat	io			Weight	<b>Risk Ratio</b>
	n/N	n/N			M-H, F	ixed, 9	95% CI				M-H, Fixed, 95% Cl
Crowther 1986	31/53	31/52								100%	0.98[0.71,1.35]
Total (95% CI)	53	52				$\blacklozenge$				100%	0.98[0.71,1.35]
Total events: 31 (Strict bed rest), 31	(Some rest)										
Heterogeneity: Not applicable											
Test for overall effect: Z=0.12(P=0.9)	1)										
		Favours strict rest	0.1	0.2	0.5	1	2	5	10	Favours some rest	

# Analysis 1.9. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 9 Endotracheal intubation.

Study or subgroup	Strict bed rest	Some rest		Ris	k Rat	io			Weight	<b>Risk Ratio</b>
	n/N	n/N		M-H, Fiz	ked, 9	95% CI				M-H, Fixed, 95% Cl
Crowther 1986	2/53	3/52		-			_		100%	0.65[0.11,3.76]
Total (95% CI)	53	52					-		100%	0.65[0.11,3.76]
Total events: 2 (Strict bed rest), 3 (	Some rest)									
Heterogeneity: Not applicable										
Test for overall effect: Z=0.48(P=0.6	63)									
		Favours strict rest	0.1 0.2	0.5	1	2	5	10	Favours some rest	

# Analysis 1.10. Comparison 1 Strict bed rest in hospital versus some rest in hospital, Outcome 10 Admission to neonatal intensive care nursery.

Study or subgroup	Strict bed rest	Some rest			Ri	sk Rat	io			Weight	<b>Risk Ratio</b>
	n/N	n/N			M-H, F	ixed, 9	95% CI				M-H, Fixed, 95% CI
Crowther 1986	20/53	26/52			_					100%	0.75[0.49,1.17]
Total (95% CI)	53	52								100%	0.75[0.49,1.17]
Total events: 20 (Strict bed rest	:), 26 (Some rest)										
Heterogeneity: Not applicable											
Test for overall effect: Z=1.25(P	=0.21)										
		Favours strict rest	0.1	0.2	0.5	1	2	5	10	Favours some rest	

# Comparison 2. Some rest in hospital versus routine activity at home

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Severe hypertension	1	218	Risk Ratio (M-H, Fixed, 95% CI)	0.58 [0.38, 0.89]
2 Pre-eclampsia	1	218	Risk Ratio (M-H, Fixed, 95% CI)	0.98 [0.80, 1.20]
3 Elective delivery, including elective caesarean section or induction of labour, or both	1	218	Risk Ratio (M-H, Fixed, 95% CI)	0.98 [0.70, 1.37]
4 Caesarean section	1	218	Risk Ratio (M-H, Fixed, 95% CI)	1.41 [0.79, 2.52]
5 Death of baby	1	218	Risk Ratio (M-H, Fixed, 95% CI)	1.96 [0.18, 21.34]
6 Death of baby by timing of death	1		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
6.1 Stillbirth	1	218	Risk Ratio (M-H, Fixed, 95% CI)	4.91 [0.24, 101.10]
6.2 Perinatal death	1	218	Risk Ratio (M-H, Fixed, 95% CI)	1.96 [0.18, 21.34]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
6.3 Neonatal death	1	218	Risk Ratio (M-H, Fixed, 95% CI)	0.33 [0.01, 7.95]
7 Small-for-gestational age	1	218	Risk Ratio (M-H, Fixed, 95% CI)	0.98 [0.51, 1.91]
8 Preterm birth	1	218	Risk Ratio (M-H, Fixed, 95% CI)	0.53 [0.29, 0.99]
9 Preterm birth (subgroup by gestational age)	1		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
9.1 Birth < 37 weeks	1	218	Risk Ratio (M-H, Fixed, 95% CI)	0.53 [0.29, 0.99]
9.2 Birth < 34 weeks	1	218	Risk Ratio (M-H, Fixed, 95% CI)	0.49 [0.09, 2.62]
10 Admission to neonatal in- tensive care unit	1	218	Risk Ratio (M-H, Fixed, 95% CI)	0.82 [0.37, 1.81]
11 Women satisfied with man- agement	1	86	Risk Ratio (M-H, Fixed, 95% CI)	0.98 [0.92, 1.04]
12 Women not choosing same management for future preg- nancy	1	86	Risk Ratio (M-H, Fixed, 95% CI)	3.0 [1.43, 6.31]

# Analysis 2.1. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 1 Severe hypertension.

Study or subgroup	Some rest	Routine activity			Ri	sk Rat	tio			Weight	Risk Ratio
	n/N	n/N			M-H, Fi	ixed,	95% CI				M-H, Fixed, 95% Cl
Crowther 1992	25/110	42/108				-				100%	0.58[0.38,0.89]
Total (95% CI)	110	108				•				100%	0.58[0.38,0.89]
Total events: 25 (Some rest), 42 (	Routine activity)										
Heterogeneity: Not applicable											
Test for overall effect: Z=2.52(P=	0.01)										
	F	avours some rest	0.1	0.2	0.5	1	2	5	10	Favours rou activity	

# Analysis 2.2. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 2 Pre-eclampsia.

Study or subgroup	Some rest	Routine activity			Ri	sk Rat	io			Weight	Risk Ratio
	n/N	n/N			М-Н, F	ixed, 9	95% CI				M-H, Fixed, 95% CI
Crowther 1992	69/110	69/108								100%	0.98[0.8,1.2]
Total (95% CI)	110	108				•				100%	0.98[0.8,1.2]
Total events: 69 (Some rest), 69	(Routine activity)										
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0,	df=0(P<0.0001); I <sup>2</sup> =100%										
Test for overall effect: Z=0.18(P=	=0.86)										
	Fa	vours some rest	0.1	0.2	0.5	1	2	5	10	Favours rou activity	



# Analysis 2.3. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 3 Elective delivery, including elective caesarean section or induction of labour, or both.

Study or subgroup	Some rest	Routine activity			Ri	sk Rat	io			Weight	Risk Ratio
	n/N	n/N			М-Н, Р	ixed, 9	95% CI				M-H, Fixed, 95% CI
Crowther 1992	42/110	42/108								100%	0.98[0.7,1.37]
Total (95% CI)	110	108				$\blacklozenge$				100%	0.98[0.7,1.37]
Total events: 42 (Some rest), 42 (Ro	utine activity)										
Heterogeneity: Not applicable											
Test for overall effect: Z=0.11(P=0.93	1)										
	E	avours some rest	0.1	0.2	0.5	1	2	5	10	Favours rou activity	

# Analysis 2.4. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 4 Caesarean section.

Study or subgroup	Some rest	Routine activity			Ri	sk Rat	io			Weight	Risk Ratio
	n/N	n/N			M-H, F	ixed, 9	95% CI				M-H, Fixed, 95% CI
Crowther 1992	23/110	16/108								100%	1.41[0.79,2.52]
Total (95% CI)	110	108								100%	1.41[0.79,2.52]
Total events: 23 (Some rest), 16 (F	Routine activity)										
Heterogeneity: Not applicable											
Test for overall effect: Z=1.16(P=0	.24)										
	E	avours some rest	0.1	0.2	0.5	1	2	5	10	Favours rou activity	

# Analysis 2.5. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 5 Death of baby.

Study or subgroup	Some rest	Routine activity			Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H	, Fixed, 95% C	I			M-H, Fixed, 95% Cl
Crowther 1992	2/110	1/108						100%	1.96[0.18,21.34]
Total (95% CI)	110	108		_				100%	1.96[0.18,21.34]
Total events: 2 (Some rest), 1 (Ro	outine activity)								
Heterogeneity: Not applicable									
Test for overall effect: Z=0.55(P=0	0.58)					1			
	Ĩ	avours some rest	0.01	0.1	1	10	100	Favours rou activity	

# Analysis 2.6. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 6 Death of baby by timing of death.

Study or subgroup	Some rest	Routine activity		Risk R	atio		Weight	Risk Ratio
	n/N	n/N		M-H, Fixed	, 95% CI			M-H, Fixed, 95% Cl
2.6.1 Stillbirth								
Crowther 1992	2/110	0/108		-+	-	-	100%	4.91[0.24,101.1]
Subtotal (95% CI)	110	108				-	100%	4.91[0.24,101.1]
Total events: 2 (Some rest), 0 (Routin	e activity)							
Heterogeneity: Not applicable								
Test for overall effect: Z=1.03(P=0.3)								
2.6.2 Perinatal death								
Crowther 1992	2/110	1/108			+		100%	1.96[0.18,21.34]
Subtotal (95% CI)	110	108					100%	1.96[0.18,21.34]
Total events: 2 (Some rest), 1 (Routin	e activity)							
Heterogeneity: Not applicable								
Test for overall effect: Z=0.55(P=0.58)	)							
2.6.3 Neonatal death								
Crowther 1992	0/110	1/108					100%	0.33[0.01,7.95]
Subtotal (95% CI)	110	108	-				100%	0.33[0.01,7.95]
Total events: 0 (Some rest), 1 (Routin	e activity)							
Heterogeneity: Not applicable								
Test for overall effect: Z=0.69(P=0.49)								
	F	avours some rest	0.001	0.1 1	10	1000	Favours rou activity	

# Analysis 2.7. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 7 Small-for-gestational age.

Study or subgroup	Some rest	Routine activity			Ri	sk Ra	tio			Weight	Risk Ratio
	n/N	n/N			M-H, F	ixed,	95% CI				M-H, Fixed, 95% Cl
Crowther 1992	15/110	15/108								100%	0.98[0.51,1.91]
Total (95% CI)	110	108				$\blacklozenge$				100%	0.98[0.51,1.91]
Total events: 15 (Some rest), 15 (Rou	tine activity)										
Heterogeneity: Not applicable											
Test for overall effect: Z=0.05(P=0.96	)				i						
	F	avours some rest	0.1	0.2	0.5	1	2	5	10	Favours rou activity	

# Analysis 2.8. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 8 Preterm birth.

Study or subgroup	Some rest	Routine activity			Ri	sk Rat	tio			Weight	Risk Ratio
	n/N	n/N		P	M-H, Fi	ixed, 9	95% CI				M-H, Fixed, 95% CI
Crowther 1992	13/110	24/108			-					100%	0.53[0.29,0.99]
Total (95% CI)	110	108				-				100%	0.53[0.29,0.99]
		Favours some rest	0.1	0.2	0.5	1	2	5	10	Favours rou activity	



Study or subgroup	Some rest	Routine activity			Ri	sk Rat	tio			Weight	Risk Ratio
	n/N	n/N			M-H, F	ixed,	95% CI				M-H, Fixed, 95% CI
Total events: 13 (Some rest), 24 (Rou	tine activity)										
Heterogeneity: Not applicable											
Test for overall effect: Z=1.99(P=0.05	)			1	1						
		Equours some rest	0.1	0.2	0.5	1	2	5	10	Eavours rou activity	

Favours some rest 0.1 0.2 0.5 1 2 5 10 Favours rou activity

# Analysis 2.9. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 9 Preterm birth (subgroup by gestational age).

Study or subgroup	Some rest	Routine activity		Risk Ratio		Weight	Risk Ratio
	n/N	n/N		M-H, Fixed, 95	% CI		M-H, Fixed, 95% Cl
2.9.1 Birth < 37 weeks							
Crowther 1992	13/110	24/108				100%	0.53[0.29,0.99]
Subtotal (95% CI)	110	108		•		100%	0.53[0.29,0.99]
Total events: 13 (Some rest), 24 (Rout	ine activity)						
Heterogeneity: Not applicable							
Test for overall effect: Z=1.99(P=0.05)							
2.9.2 Birth < 34 weeks							
Crowther 1992	2/110	4/108				100%	0.49[0.09,2.62]
Subtotal (95% CI)	110	108				100%	0.49[0.09,2.62]
Total events: 2 (Some rest), 4 (Routine	e activity)						
Heterogeneity: Not applicable							
Test for overall effect: Z=0.83(P=0.41)							
	F	avours some rest	0.01	0.1 1	10 1	<sup>00</sup> Favours rou activity	

Analysis 2.10. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 10 Admission to neonatal intensive care unit.

Study or subgroup	Some rest	Routine activity			Ris	sk Rat	tio			Weight	Risk Ratio
	n/N	n/N			M-H, Fi	ixed, 9	95% CI				M-H, Fixed, 95% CI
Crowther 1992	10/110	12/108				-	_			100%	0.82[0.37,1.81]
Total (95% CI)	110	108								100%	0.82[0.37,1.81]
Total events: 10 (Some rest), 12 (Ro	utine activity)										
Heterogeneity: Not applicable											
Test for overall effect: Z=0.49(P=0.6	2)										
		Favours some rest	0.1	0.2	0.5	1	2	5	10	Favours rou activity	

# Analysis 2.11. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 11 Women satisfied with management.

Study or subgroup	Some rest	Routine activity			Ri	sk Rat	io			Weight	Risk Ratio
	n/N	n/N			M-H, F	ixed, 9	95% CI				M-H, Fixed, 95% CI
Leung 1998	42/43	43/43				+				100%	0.98[0.92,1.04]
Total (95% CI)	43	43				•				100%	0.98[0.92,1.04]
Total events: 42 (Some rest), 43 (F	Routine activity)										
Heterogeneity: Not applicable											
Test for overall effect: Z=0.71(P=0	.48)										
	F	avours some rest	0.1	0.2	0.5	1	2	5	10	Favours rou activity	

# Analysis 2.12. Comparison 2 Some rest in hospital versus routine activity at home, Outcome 12 Women not choosing same management for future pregnancy.

Study or subgroup	Some rest	Routine activity			Ris	sk Ra	tio			Weight	Risk Ratio
	n/N	n/N			M-H, Fi	ixed,	95% CI				M-H, Fixed, 95% CI
Leung 1998	21/43	7/43								100%	3[1.43,6.31]
Total (95% CI)	43	43								100%	3[1.43,6.31]
Total events: 21 (Some rest), 7 (Routine activity)											
Heterogeneity: Not applicable											
Test for overall effect: Z=2.9(P=0)											
	F	avours some rest	0.1	0.2	0.5	1	2	5	10	Favours rou activity	

#### Favours some rest

# APPENDICES

# Appendix 1. Additional searching for initial version of review

Sources searched	Search strategy
The Cochrane Central Register of Controlled Trials (The Cochrane Library 2005, Issue 1) and EMBASE (January 2002 to December 2004).	We used the search strategy listed in the generic protocol (see Meher 2005 in 'Additional references' ).

# WHAT'S NEW

Date	Event	Description
18 January 2010	New search has been performed	Search updated. No new trials identified

## HISTORY

Protocol first published: Issue 1, 2002 Review first published: Issue 4, 2005

Date	Event	Description				
10 November 2008	Amended	Contact details amended				
11 February 2008	Amended	Converted to new review format.				
25 October 2007	New search has been performed	Search updated. No new trial reports identified.				
26 May 2005	New citation required and conclusions have changed	Substantive amendment				

# CONTRIBUTIONS OF AUTHORS

Guillermo Carroli wrote the initial version of the protocol. Edgardo Abalos contributed to the final version of the protocol. The protocol was modified by Shireen Meher and Edgardo Abalos following its publication due to the discovery of unexpected but relevant comparisons and definitions of bed rest used by trialists.

Shireen Meher and Edgardo Abalos decided on eligible trials and did the data extraction. Data were entered by Shireen Meher, and double checked by Edgardo Abalos. Shireen Meher drafted the review, and the final report was prepared with contributions from Edgardo Abalos.

### DECLARATIONS OF INTEREST

None known.

# SOURCES OF SUPPORT

#### Internal sources

- Centro Rosarino de Estudios Perinatales-CREP-Rosario, Argentina.
- The University of Liverpool, UK.

#### **External sources**

- Human Reproduction Programme-HRP-World Health Organization. Geneva, Switzerland.
- Health Technology Assessment, UK.

### DIFFERENCES BETWEEN PROTOCOL AND REVIEW

The original published protocol was updated while drafting the 2007 updated review. The differences include:

(1) a new contact author (S Meher) and additional sources of support were added.

(2) the 'Objectives' were reworded and additional comparisons were included. This was done to accommodate for variation of definitions of 'bed rest' and 'normal activity' used by trialists. One trialist's bed rest in hospital (rest in bed but allowed to move around the ward) was another trialist's normal activity in hospital (this trialist defined bed rest as strict bed rest where the participant only got up to go to the toilet). To avoid pooling interventions which crossed arms in different trials, we divided bed rest into different degrees and created new comparisons of different degrees of bed rest. Moreover, the original protocol did not allow for comparisons of bed rest and normal activity across different settings in the same trial, for example, bed rest in hospital versus normal activity at home. We felt that these were important comparisons that needed to be included in the review.

# INDEX TERMS

#### Medical Subject Headings (MeSH)

\*Bed Rest; \*Hospitalization; Hypertension [\*therapy]; Pregnancy Complications, Cardiovascular [\*therapy]; Pregnancy Outcome; Premature Birth [prevention & control]; Randomized Controlled Trials as Topic



# **MeSH check words**

Female; Humans; Pregnancy